

Article

Trends in hypertension management in Type I diabetes across Europe, 1989/1990 – 1997/1999

S. S. Soedamah-Muthu (✉) · H. M. Colhoun · H. Abrahamian · N. N. Chan · R. Mangili · G. P. Reboldi · J. H. Fuller · the EURODIAB Prospective Complications Study Group

S. S. Soedamah-Muthu

EURODIAB, Department of Epidemiology and Public-Health, Royal Free and University College London Medical School, 1–19 Torrington Place, London, WC1E 6BT, UK

S. S. Soedamah-Muthu · H. M. Colhoun · N. N. Chan · J. H. Fuller

Department of Epidemiology and Public-Health, Royal Free and University College London Medical School, London, UK

H. Abrahamian

Medical department, Hospital Vienna Lainz, Vienna, Austria

R. Mangili

Divisions di Medicina, Istituto Scientifico San Raffaele, Milan, Italy

G. P. Reboldi

Dipartimento di Medicina Interna e Scienze, Endocrine e Metaboliche, Perugia, Italy

✉ S. S. Soedamah-Muthu

E-mail: sabita@public-health.ucl.ac.uk

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Abstract *Aims/hypothesis* Our aim was to examine the change in the management of hypertension in patients with Type I (insulin-dependent) diabetes mellitus in Europe, between 1989–1990 and 1997–1999.

Methods Seven-year changes in hypertension treatment and control (defined as blood pressure <130/85 mmHg) were examined in a large sample of Type I diabetic patients recruited from 26 centres involved in the EURODIAB Prospective Complications Study. Hypertension was defined as a systolic and/or diastolic blood pressure greater than 140 and/or 90 mmHg respectively, and/or use of blood pressure lowering drugs.

Results Of 1866 Type I diabetic patients, 412 had hypertension at baseline and 631 at follow-up. A greater proportion of hypertensive patients were treated at follow-up (69% vs 40%, $p<0.0001$), which persisted after adjustment for age or centre. Of those who were treated, a modest increase in the proportion of those controlled for hypertension was found (41% vs 32%, $p=0.048$), which disappeared after adjustment for age. Among hypertensive patients with albuminuria, the proportions treated also increased, from 35% to 76% ($p<0.0001$) in microalbuminuric and 64% to 95% ($p<0.0001$) in macroalbuminuric patients. Control of hypertension in albuminuric patients did not change significantly and was below 50%. The use of more than one anti-hypertensive drug increased over a 7-year period, from 19% to 33% ($p<0.0001$), and a marked increase was shown in the proportion of those taking an ACE inhibitor (from 57% to 82%, $p<0.0001$).

Conclusion/interpretation The management of hypertension in Type I diabetic patients across Europe has improved over a 7-year follow-up period. Optimal levels of blood pressure treatment and optimal levels of control have not yet been achieved.

Keywords Hypertension · Type I diabetes mellitus · albuminuria · control · blood pressure · treatment

Abbreviations

CCB Calcium channel blocker

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In Type I (insulin-dependent) diabetes mellitus, raised blood pressure has been shown to be related to an increased risk of retinopathy [1, 2, 3], and cardiovascular disease [4, 5] in several prospective studies independent of other risk factors. It is estimated that 30 to 75% of diabetic complications can be attributed to hypertension, which is twice as common in diabetic patients than in non-diabetic people [6].

Several BP lowering trials have shown a reduction in microvascular and macrovascular complications in patients with Type II (non-insulin-dependent) diabetes mellitus [7, 8]. Other recent trials have shown that, independent from BP lowering effects, the progression of diabetic nephropathy can be reduced by angiotensin II receptor blockers in Type II diabetic patients with hypertension and albuminuria [9, 10, 11]. In Type I diabetes, the progression of retinopathy [12]

and nephropathy [13, 14] has also been reduced with the use of ACE inhibitors in normotensive subjects.

Since the overall risk of developing cardiovascular complications in diabetic patients is higher than in non-diabetic people and, given the importance of increased BP in the pathogenesis of diabetic complications, several expert groups have made specific recommendations on the management of hypertension in diabetic patients (Table 1). All these guidelines stress that a diagnosis of hypertension should be based on repeated measurements, management decisions should consider the presence of cardiovascular, renal and retinal end-organ damage, and that a trial period of non-pharmacological intervention is appropriate in those with mild hypertension. The guidelines recommend treating diabetic patients to a lower target BP than in the non-diabetic population, especially in the presence of proteinuria.

[Table 1. will appear here. See end of document.]

Previous studies on hypertension treatment and control in non-diabetic cohorts in Europe and in the United States [15, 16, 17, 18, 19, 20, 21] have shown improvements over time, but inadequate management of hypertension. However, few studies have examined the management of hypertension specifically in diabetic patients [22, 23].

The purpose of our study is to evaluate the impact of hypertension guidelines on the management of hypertension in Type I diabetic patients in the EURODIAB Prospective Complications Study. We have previously published data on the management of hypertension at baseline in the EURODIAB IDDM Complications Study cohort [24]. This study examines how hypertension management has changed after a 7-year period and to what extent this is independent of the cohort getting older.

Subjects and methods

Study design and subjects. Full details of the design, methods and recruitment in the EURODIAB PCS have been published elsewhere [25, 26, 27]. The baseline cross-sectional clinic-based study was carried out in 1989–1991. It was set up to explore risk factors for diabetic complications in 3250 randomly selected people with Type I diabetes of 15 to 60 years of age and attending 31 diabetic clinics in 16 European countries. Sampling was stratified by age, sex and diabetes duration. Type I diabetes was defined as diabetes diagnosed before the age of 36 years with a continuous need for insulin within 1 year after diagnosis. Of those invited 85% participated. Those with

diabetes for less than 1 year and pregnant women were not recruited into the study. At each clinic informed consent from all subjects and ethics committee approval for the study was obtained.

Follow-up. Approximately 6 to 8 years after the baseline examinations the study participants were traced and invited back for re-examination. Of the 3250 subjects at baseline, 460 were from centres who did not participate at follow-up. Five centres did not participate in assessments on the management of hypertension at follow-up. Of the remaining 2790 subjects, 2334 subjects were traceable, 99 had died and 357 were lost at follow-up, 1880 had vital status data, of which 1866 subjects were examined for hypertension related assessments, from 26 diabetic clinics in 14 European countries (France, Poland, Croatia, Italy, Finland, Portugal, Austria, Germany, Luxembourg, Belgium, the Netherlands, England, the Republic of Ireland and Greece).

Measurements. The same methods were used at baseline and at follow-up. Patients were asked whether they had had high BP, angina, a heart attack or stroke at any time in the past, diagnosed by a doctor or a nurse. Blood pressure was measured by a random zero sphygmomanometer (Hawksley, Lancing, UK). Using an appropriately sized cuff, two BP readings were taken from the right arm with the patient in a seated position after resting for 5 min. Readings were taken from the top of the meniscus and measurement was recorded to the nearest 2 mmHg. Diastolic BP was recorded at the disappearance of sound (Korotkoff phase V). Data presented here are based on the mean of two measurements. Hypertension was defined as a systolic BP of 140 mmHg or more, or a diastolic BP of 90 mmHg or more, and/or the current use of BP lowering drugs. Patients were considered to be treated for hypertension if they were currently taking BP lowering drugs as assessed from clinical records by local investigators. Control was defined as having a systolic BP of less than 130 mmHg and a diastolic BP of less than 85 mmHg.

A single-timed 24-h urine collection was done to calculate albumin excretion rate after excluding proteinuria due to urinary tract infection. Urinary albumin was measured in a single laboratory by an immunoturbidimetric method (Sanofi Diagnostics Pasteur, Minneapolis, Minn., USA) [28]. Albumin excretion rate was categorised as normoalbuminuria at 20 µg/min or less, microalbuminuria between 20 and 200 µg/min and macroalbuminuria at 200 µg/min or more. At baseline, albumin excretion rates were measured once, whereas at follow-up these were measured twice, with the mean taken as data. If only one measurement was available then the result of just that one collection was used, but this was only the case in 11% (170 out of 1594) of the total number of patients.

Statistical analysis. The statistical packages SAS (SAS, Cary, N.C., USA) and STATA (STATA 6.0, Tex., USA) were used to carry out all statistical analyses. Chi-square and Student's *t* tests were used to test for differences in baseline risk factors between those included and those lost at

follow-up. Mc-Nemar and paired t tests were used to analyse differences in risk factors between the baseline and follow-up sample. The crude proportions of patients with hypertension, those who were on treatment and who were controlled were calculated both at baseline and at follow-up. These proportions were presented by 10-year age bands (using the baseline age) and by albuminuric status. Trends with age (age bands) in hypertension parameters were tested for using chi-square tests for trend. The 95% confidence intervals were calculated for the proportions (of the total) and the chi-square test or Mc-Nemar tests (if denominators were exactly similar) were used to test for differences in the proportions between baseline and follow-up. To test if changes over time in the proportions of hypertensives, treated and controlled for hypertension were independent of age, further conditional or logistic regression models were conducted, fitting time (as a dummy variable baseline or follow-up) and age (age at baseline or follow-up) for the subjects who were included at both timepoints. Likelihood ratio tests were used to test differences between models on hypertension, treatment or control, fitting time alone and models with both time and age. Similarly, the effects of centre on changes in hypertension, treatment and control over time were tested with likelihood ratio tests, to compare logistic regression models fitting time and centre (fitted as a dummy variable) with models fitting time alone. In addition, possible risk factors related to control of BP were analysed by using chi-square tests for categorical variables or t tests for continuous variables. A p value of less than 0.05 was considered to be statistically significant.

Results

The mean age at baseline was 33 years (age ranged from 14.9–60.7 years) and the mean duration of diabetes was 15 years (SD=9) (Table 2). The baseline characteristics of those included in this analysis ($n=1866$) and those lost to follow-up ($n=1384$) were mostly similar, except for a slightly higher proportion of current smokers (chi-square $p=0.002$) and a higher frequency of retinopathy (chi-square $p=0.02$), albuminuria (chi-square $p=0.01$), hypertension (chi-square $p=0.001$) and cardiovascular disease (chi-square $p=0.001$) in those who were lost at follow-up (Table 2). Patients who were lost at follow-up reported slightly more use of BP lowering drugs and were less well-controlled than those included in the study, but these findings were not statistically significant.

[Table 2. will appear here. See end of document.]

Sample characteristics changed between baseline and follow-up. By the time of follow-up, patients were on average 7 years older (mean age: 40 years, range from 22.6–67.6 years) and subsequently had a longer duration of diabetes (Table 2). In addition, there was a higher proportion

of patients with cardiovascular disease (McNemar $p=0.001$) and retinopathy at follow-up (McNemar $p=0.001$). All other risk factors were similar between baseline and follow-up.

Proportion of hypertensive patients. The (crude) proportion of hypertensive patients (systolic BP ≥ 140 mmHg or diastolic BP ≥ 90 mmHg or current use of anti-hypertensive drugs) increased from 22% (412/1866) at baseline to 34% (631/1866) at follow-up (McNemar $p=0.001$, Table 2). Of those who were hypertensive at baseline, 73% were still hypertensive at follow-up. As expected the proportion of hypertensive patients increased with age, for both men and women (chi-square test for trend $p=0.001$, Table 3). To show whether this increase over time was greater than expected from the ageing of the cohort, the data is presented by baseline age bands (Table 3). An increase in hypertension prevalence was found across all age groups. However, further modelling with conditional logistic regression models showed that there was no increase in hypertension over time when adjusted for age (Likelihood ratio-test $p=0.51$). Further adjustment for centre did not alter these results. Hence, after adjusting for age, the proportion of hypertensive patients was similar between baseline and follow-up.

[Table 3. will appear here. See end of document.]

Treatment of hypertension. Of those with hypertension, the crude proportion who were treated with BP lowering drugs increased from 40% (163/409) at baseline to 69% (434/628) at follow-up (chi-square $p<0.0001$) (Table 2). Of those on BP lowering drugs at baseline, 93% were reported to be still on drugs at follow-up. The proportions treated at baseline and at follow-up varied with age (chi-square test for trend $p=0.01$, Table 3). Treatment significantly improved comparing baseline and follow-up assessments across all age-bands (chi-square $p<0.001$). Further modelling with logistic regression analysis showed that the increase in treatment with BP lowering drugs over time remained significant, after adjustment for the ageing effect (likelihood ratio-test $p<0.0001$). Adjustment for centre did not change these findings.

Control of hypertension. Control of hypertension (defined as a systolic BP <130 and diastolic BP <85 mmHg) increased from baseline 32% (51/161) to follow-up 41% (176/434), in hypertensive patients who were on treatment, but was borderline significant (chi-square $p=0.048$) (Table 2). Of those controlled at baseline, only 59% were controlled at follow-up. A trend with age in control of hypertension was found (chi-square test for trend $p=0.02$). There were improvements in control of hypertension between baseline and follow-up by age-bands, but these were not statistically significant (Table 3). Further modelling with logistic regression analysis showed that there was no significant increase in the control of hypertension over time (likelihood ratio-test $p=0.09$) after allowing for the ageing effect. Adjustment for centre did not alter these results.

Factors associated with poor control of hypertension at follow-up. Baseline risk factors related to control of hypertension at follow-up ($n=176$ controlled, 258 not controlled) in treated hypertensive patients were analysed. There were no differences in sex, smoking status, albuminuria status, concentrations of glycated haemoglobin, fasting triglycerides and HDL-cholesterol between those controlled and those not-controlled. Those not controlled were more likely to have had complications at baseline, including retinopathy (78% vs 67%, chi-square $p=0.07$) and cardiovascular complications (19% vs 11%, chi-square $p=0.03$) compared to well controlled patients. They were also older (baseline age 39 vs 35 years, $p<0.0001$), had longer duration of diabetes (21 vs 18 years, t test $p=0.001$), and had higher systolic and diastolic BP (137/84 vs 124/78 mmHg, t test $p<0.0001$), BMI (24.8 vs 23.9 kg/m², t test $p=0.005$), WHR (0.87 vs 0.85, t test $p=0.05$), total cholesterol (5.8 vs 5.5 mmol/l, t test $p=0.003$) and LDL-cholesterol (3.8 vs 3.4 mmol/l, t test $p=0.002$).

Management of hypertension in albuminuric patients. Unadjusted proportions of the frequency, treatment and control of hypertension are shown by albuminuria status (Table 4). The proportion of hypertensive patients, those treated and controlled increased with albuminuric status. From baseline to follow-up, there was an increase in the proportion of hypertensive patients across all albuminuric groups, which was not statistically significant after adjusting for age. The proportion of those with microalbuminuria and macroalbuminuria treated for their hypertension increased from baseline to follow-up, from 35% to 76% (chi-square $p<0.0001$) and from 64% to 95% (chi-square $p<0.0001$), respectively. For control of hypertension in albuminuric patients who were hypertensive and who were treated with BP lowering drugs, non-significant increases were shown from 35% to 47% (chi-square $p=0.18$) in those with microalbuminuria and from 22% to 36% (chi-squared $p=0.06$) in those with macroalbuminuria.

[Table 4. will appear here. See end of document.]

Number and type of BP lowering drugs used among those on treatment. A larger proportion of all diabetic patients were treated with BP lowering drugs at follow-up (23%, 435/1866) compared to baseline (17%, 310/1866) (Mc-Nemar $p<0.0001$). The use of ACE-inhibitors increased from baseline (57%) to follow-up (82%) (chi-square $p<0.0001$) (Table 5). At baseline, 81% were on one drug, whereas at follow-up, this proportion had decreased to 67%. More patients were treated with multiple drugs in 1997 to 1999 (33%) than at baseline (19%) (chi-squared $p<0.0001$). An ACE-inhibitor and a calcium antagonist was the predominantly used combination among those receiving two BP lowering drugs at both time-points and increased from 1989–1990 to 1997–1999. Similar patterns in the use of BP lowering drugs were shown for patients with albuminuria. About 77% of those treated for hypertension, but not controlled, were taking one drug at follow-up.

[Table 5. will appear here. See end of document.]

Discussion

We have shown in this EURODIAB PCS cohort of Type I diabetic patients that hypertension treatment has improved over a 7-year follow-up period. However, the number of patients treated and well controlled is still not optimal, especially in patients with both hypertension and albuminuria. Poor control at follow-up is related to a more atherogenic risk profile and a higher frequency of complications at baseline. There is still a long way to go before target BP levels, according to new and more stringent guidelines, will be reached. We have also shown an increase in the use of ACE-inhibitors, a decrease in the use of monotherapy and an increase in the use of multiple drugs from baseline to follow-up.

Clinical trials with BP lowering drugs. Convincing evidence exists of the beneficial effects of reducing BP in Type II diabetic patients, summarised in several reviews [7, 8]. Lower target levels (<130/80 mmHg) were indicated [29, 30, 31] and greater risk reductions were reported in diabetic patients than in non-diabetic subjects [32, 33]. Different anti-hypertensive treatments have been examined, including ACE inhibitors [31, 34, 35, 36], diuretics or beta-blockers [32, 36, 37, 38] and CCB's [33, 35, 36, 37, 38] which all have been shown to be beneficial compared to placebo or conventional treatments. To achieve lower BP goals in diabetic patients with hypertension, the use of ACE inhibitors (or angiotensin II receptor blockers if ACE inhibitors cannot be tolerated) as first-line therapy and the use of multiple agents has been shown to be advantageous [29, 30]. The benefits of angiotensin II receptor blockers [9, 10, 11] on nephropathy has been shown in Type II diabetes. To date, there are no BP lowering trials reporting on microvascular and macrovascular events in Type I diabetes, even though reductions in the progression of retinopathy [12] and nephropathy [13, 14] were reported with the use of ACE inhibitors in normotensive subjects.

Hypertension management guidelines. Several national and international guidelines have been published on the management of hypertension in diabetes. These guidelines, which include recommendations for specific populations such as diabetic patients, could have contributed to the improvement of the hypertension treatment over time. On the other hand, the complexity of the guidelines and the absence of proper guidelines for Type I diabetes could have discouraged physicians from applying them, leading to suboptimal management of hypertension. Most guidelines, but not all, recommend a systolic BP target of less than 130 mmHg and a diastolic BP

target of less than 85 mmHg and recommend earlier intervention in Type I diabetic patients, especially in those with albuminuria.

Observational studies on hypertension treatment and control. Substantial evidence from European and American studies has confirmed improvements in treatment and control of hypertension but still with suboptimal BP levels in non-diabetic subjects, from cross-sectional [39, 40, 41, 42, 43, 44, 45, 46] and prospective studies [15, 16, 17, 18, 19, 20, 47, 48, 49, 50].

In the EUROASPIRE I and II studies (1995–1996 to 1999–2000) in patients with coronary heart disease, the proportion of hypertensive patients was high at both timepoints (55% vs 54%), but the proportion of patients on BP lowering drugs achieving a target of <140/90 mmHg was low and did not change over time (44% vs 45%). In the same study, there was a slight increase in the use of BP lowering drugs (84% to 89%), and an increase in the use of beta-blockers (54% to 66%) and ACE inhibitors (from 30% to 43%) [17].

In the United States, using data from the National Health and Nutrition Examination Survey (NHANES) more than 7000 non-diabetic subjects, increases in the treatment and control of hypertensive patients but with suboptimal levels of management, were shown from 1976–1980 (NHANES II) to 1991–1994 (NHANES III phase 2) [16]. With the same definition of hypertension, the proportion of hypertensive patients treated with blood pressure lowering drugs increased from 31% to 53%, which is similar to our data (increases from 40% to 69%). Control of hypertension (<140/90 mmHg) increased, as a proportion of the hypertensive patients (from 10% to 27%) which is similar to our data (from 12% to 28%) [15, 16]. For control of hypertension using treated hypertensive patients as the denominator, an increase was seen from 32% to 55% (NHANES phase 1 1988–1991), corresponding with our data (32% to 41%) [49].

In Type I diabetes, the Pittsburgh (USA) Epidemiology of Diabetes Complications Study, showed an increase in control of hypertension from 38% in 1986–1988 to 49.5% in 1996–1998 in those treated and aware of hypertension [23].

Possible explanations for poor hypertension control. Multiple factors could explain suboptimal levels of treatment and control of hypertension but both the patient and physician are important. From the patients' perspective, male sex [39, 46], obesity [39, 40], alcohol [18] and patient education [51] have been suggested to explain poor control. In our study, we also found that increased age, duration of diabetes, BP levels, lipid concentrations and the presence of more complications, as well as obesity at baseline partly explained the reasons for poor control at follow-up. In addition, costs and side-effects of drugs, the complexity of the regimen including multiple drugs and dosages,

lack of patients' adherence to medication and lifestyle modifications could contribute to poor control.

From the physicians' point of view, inappropriate or ineffective treatments, lack of acceptance or knowledge of hypertension treatment guidelines, costs and side-effects, and dosing schedules of drugs have been suggested to play a role in poor management [52, 53, 54]. As a result physicians have been shown to initiate treatment at higher BP thresholds than existing guidelines, are less attentive to controlling BP because they are satisfied with the existing BP, seldom intensify drug therapy for BP above target levels and are more likely to treat or control increased diastolic BP than systolic BP [53, 54, 55].

Limitations of the EURODIAB PCS. The EURODIAB PCS provides a useful European-wide summary as the same standardised methods for BP measurements and albumin excretion rate assessments were used in each centre. However, there are some limitations. This is a clinic-based, and not a population-based study, and is therefore not fully representative. The definition of hypertension used in most studies is a problem because a person treated with BP lowering drugs is considered to be hypertensive, which could lead to an overestimation of the proportion of hypertensives. The apparent increase in the proportion of hypertensives over time is explained by the age, as well as a higher proportion of patients being treated. By examining this sample of Type I diabetic patients in 1989 to 1990 the physicians would have been alerted to put patients on treatment for their hypertension therefore influencing the measurements in 1997 to 1999. As BP was measured on a single occasion, it could also have led to overestimations in the proportion of hypertensives, although two recordings were taken. Underestimations in our results were possibly caused by those patients lost at follow-up who had a higher frequency of complications (retinopathy, cardiovascular disease and hypertension) compared with the baseline sample.

The compliance with BP lowering treatment, the actual indication for taking the BP lowering drugs and non-pharmacological lifestyle interventions could not be assessed in this study. Despite the standardization of methods, there are possible differences between countries in the management of hypertension, however, the results did not change after adjustment for centre.

In conclusion, despite the publication of various guidelines, optimal levels of treatment and BP control have not been achieved in hypertensive Type I diabetic patients over a 7-year period. There is more scope for improvement. Factors that are limiting the widespread use of multiple drug therapy to reduce raised BP levels need to be examined.

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centre. J.H. Fuller, N. Chaturvedi, J. Holloway, D. Webb, L. Asbury, L.K. Stevens, M. Shipley, S.J. Livingstone University College London. *Central laboratories*. G.-C. Viberti, R. Swaminathan, P. Lumb, A. Collins, S. Sankaralingham, Guy's and St Thomas Hospital, London, UK. *Retinopathy Grading Centre*. S. Aldington, T. Mortemore, H. Lipinski, Royal Postgraduate Medical School of Imperial College London, London, UK.

References

1. Klein R, Klein BE, Moss SE, Davis MD, DeMets DL (1989) Is blood pressure a predictor of the incidence or progression of diabetic retinopathy? *Arch Intern Med* 149: 2427–2432
2. Janka HU, Warram JH, Rand LI, Krolewski AS (1989) Risk factors for progression of background retinopathy in long-standing IDDM. *Diabetes* 38: 460–464
3. Sjolie AK, Stephenson J, Aldington S et al. (1997) Retinopathy and vision loss in insulin-dependent diabetes in Europe. The EURODIAB IDDM Complications Study. *Ophthalmology* 104: 252–260
4. Forrest KY, Becker DJ, Kuller LH, Wolfson SK, Orchard TJ (2000) Are predictors of coronary heart disease and lower-extremity arterial disease in type 1 diabetes the same? A prospective study. *Atherosclerosis* 148: 159–169
5. Rossing P, Hougaard P, Borch-Johnsen K, Parving HH (1996) Predictors of mortality in insulin dependent diabetes: 10 year observational follow up study. *BMJ* 313: 779–784
6. Epstein M, Sowers JR (1992) Diabetes mellitus and hypertension. *Hypertension* 19: 403–418
7. Agarwal R (2001) Treatment of hypertension in patients with diabetes: lessons from recent trials. *Cardiol Rev* 9: 36–44
8. Kaplan NM (2001) Management of hypertension in patients with Type 2 diabetes mellitus: guidelines based on current evidence. *Ann Intern Med* 135: 1079–1083
9. Brenner BM, Cooper ME, Zeeuw D de et al. (2001) Effects of losartan on renal and cardiovascular outcomes in patients with type 2 diabetes and nephropathy. *N Engl J Med* 345: 861–869
10. Parving HH, Lehnert H, Brochner-Mortensen J et al. (2001) The effect of irbesartan on the development of diabetic nephropathy in patients with type 2 diabetes. *N Engl J Med* 345: 870–878
11. Lewis EJ, Hunsicker LG, Clarke WR et al. (2001) Renoprotective effect of the angiotensin-receptor antagonist irbesartan in patients with nephropathy due to type 2 diabetes. *N Engl J Med* 345: 851–860
12. Chaturvedi N, Sjolie AK, Stephenson JM et al. (1998) Effect of lisinopril on progression of retinopathy in normotensive people with type 1 diabetes. The EUCLID Study Group. EURODIAB Controlled Trial of Lisinopril in Insulin-Dependent Diabetes Mellitus. *Lancet* 351: 28–31
13. The EUCLID Study Group (1997) Randomised placebo-controlled trial of lisinopril in normotensive patients with insulin-dependent diabetes and normoalbuminuria or microalbuminuria. The EUCLID Study Group. *Lancet* 349: 1787–1792
14. ACE Inhibitors in Diabetic Nephropathy Trialist Group (2001) Should all patients with type 1 diabetes mellitus and microalbuminuria receive angiotensin-converting enzyme inhibitors? A meta-analysis of individual patient data. *Ann Intern Med* 134: 370–379
15. Burt VL, Culter JA, Higgins M et al. (1995) Trends in the prevalence, awareness, treatment, and control of hypertension in the adult US population. Data from the health examination surveys, 1960 to 1991. *Hypertension* 26: 60–69
16. Joint National Committee (1997) The sixth report of the Joint National Committee on prevention, detection, evaluation, and treatment of high blood pressure. *Arch Intern Med* 157: 2413–2446
17. EUROASPIRE (2001) Clinical reality of coronary prevention guidelines: a comparison of EUROASPIRE I and II in nine countries. EUROASPIRE I and II Group. European Action on Secondary Prevention by Intervention to Reduce Events. *Lancet* 357: 995–1001

18. Menotti A, Lanti M, Zanchetti A et al. (2001) Impact of the Gubbio population study on community control of blood pressure and hypertension. *J Hypertens* 19: 843–850
19. Gasse C, Hense HW, Stieber J et al. (2001) Assessing hypertension management in the community: trends of prevalence, detection, treatment, and control of hypertension in the MONICA Project, Augsburg 1984–1995. *J Hum Hypertens* 2001: 27–36
20. Kastarinen MJ, Salomaa VV, Vartiainen EA et al. (1998) Trends in blood pressure levels and control of hypertension in Finland from 1982 to 1997. *J Hypertens* 16: 1379–1387
21. Marques-Vidal P, Tuomilehto J (1997) Hypertension awareness, treatment and control in the community: is the ‘rule of halves’ still valid? *J Hum Hypertens*: 213–220
22. Geiss LS, Rolka DB, Engelgau MM (2002) Elevated blood pressure among U.S. adults with diabetes, 1988–1994(1). *Am J Prev Med* 22: 42–48
23. Zgibor JC, Orchard TJ (2001) Has control of hyperlipidemia and hypertension in patients with type 1 diabetes improved over time? *Diabetes* 50: A255: 1049-P (Abstract)
24. Collado-Mesa F, Colhoun HM, Stevens LK et al. (1999) Prevalence and management of hypertension in type 1 diabetes mellitus in Europe: the EURODIAB IDDM Complications Study. *Diabet Med* 16: 41–48
25. EURODIAB IDDM Complications Study group (1994) Microvascular and acute complications in IDDM patients: the EURODIAB IDDM Complications Study. *Diabetologia* 37: 278–285
26. Chaturvedi N, Sjoelie AK, Porta M et al. (2001) Markers of insulin resistance are strong risk factors for retinopathy incidence in type 1 diabetes: The EURODIAB PCS. *Diabetes Care* 24: 284–289
27. Chaturvedi N, Bandinelli S, Mangili R et al. (2001) Microalbuminuria in type 1 diabetes: rates, risk factors and glycemic threshold. *Kidney Int* 60: 219–227
28. Kearney EM, Mount JN, Watts GF, Slavin BM, Kind PR (1987) Simple immunoturbidimetric method for determining urinary albumin at low concentrations using Cobas-Bio centrifugal analyser. *J Clin Pathol* 40: 465–468
29. Turner R, Cull C, Holman R, UKPDS study group (1998) Tight blood pressure control and risk of macrovascular and microvascular complications in type 2 diabetes: UKPDS 38. UK Prospective Diabetes Study Group. *BMJ* 317: 703–713
30. Hansson L, Zanchetti A, Carruthers SG et al. (1998) Effects of intensive blood-pressure lowering and low-dose aspirin in patients with hypertension: principal results of the Hypertension Optimal Treatment (HOT) randomised trial. HOT Study Group. *Lancet* 351: 1755–1762
31. Estacio RO, Jeffers BW, Hiatt WR et al. (1998) The effect of nisoldipine as compared with enalapril on cardiovascular outcomes in patients with non-insulin-dependent diabetes and hypertension. *N Engl J Med* 338: 645–652
32. Curb JD, Pressel SL, Cutler JA et al. (1996) Effect of diuretic-based antihypertensive treatment on cardiovascular disease risk in older diabetic patients with isolated systolic hypertension. Systolic Hypertension in the Elderly Program Cooperative Research Group. *JAMA* 276: 1886–1892
33. Tuomilehto J, Rastenyte D, Birkenhager WH et al. (1999) Effects of calcium-channel blockade in older patients with diabetes and systolic hypertension. Systolic Hypertension in Europe Trial Investigators. *N Engl J Med* 340: 677–684
34. Hansson L, Lindholm LH, Niskanen L et al. (1999) Effect of angiotensin-converting-enzyme inhibition compared with conventional therapy on cardiovascular morbidity and mortality in hypertension: the Captopril Prevention Project (CAPPP) randomised trial. *Lancet* 353: 611–616
35. Tatti P, Pahor M, Byington RP et al. (1998) Outcome results of the Fosinopril Versus Amlodipine Cardiovascular Events Randomized Trial (FACET) in patients with hypertension and NIDDM. *Diabetes Care* 21: 597–603
36. Hansson L, Lindholm LH, Ekblom T et al. (1999) Randomised trial of old and new antihypertensive drugs in elderly patients: cardiovascular mortality and morbidity the Swedish Trial in Old Patients with Hypertension-2 study. *Lancet* 354: 1751–1756
37. Hansson L, Hedner T, Lund-Johansen P et al. (2000) Randomised trial of effects of calcium antagonists compared with diuretics and beta-blockers on cardiovascular morbidity and mortality in hypertension: the Nordic Diltiazem (NORDIL) study. *Lancet* 356: 359–365

38. Brown MJ, Palmer CR, Castaigne A et al. (2000) Morbidity and mortality in patients randomised to double-blind treatment with a long-acting calcium-channel blocker or diuretic in the International Nifedipine GITS study: Intervention as a Goal in Hypertension Treatment (INSIGHT). *Lancet* 356: 366–372
39. De Henauw S, De Backer D, Fonteyne W et al. (1998) Trends in the prevalence, detection, treatment and control of arterial hypertension in the Belgian adult population. *J Hypertens* 16: 277–284
40. Gnasso A, Calindro MC, Carallo C et al. (1997) Awareness, treatment and control of hyperlipidaemia, hypertension and diabetes mellitus in a selected population of southern Italy. *Eur J Epidemiol* 13: 421–428
41. Rywik SL, Davis CE, Pajak A et al. (1998) Poland and U.S. collaborative study on cardiovascular epidemiology hypertension in the community: prevalence, awareness, treatment, and control of hypertension in the Pol-MONICA Project and the U.S. Atherosclerosis Risk in Communities Study. *Ann Epidemiol* 8: 3–13
42. Mancia G, Sega R, Milesi C, Cesana G, Zanchetti A (1997) Blood-pressure control in the hypertensive population. *Lancet* 349: 454–457
43. Lang T, Gaudemaris R de, Chatellier G, Hamici L, Diene E (2001) Prevalence and therapeutic control of hypertension in 30,000 subjects in the workplace. *Hypertension* 38: 449–454
44. Stergiou GS, Thomopoulou GC, Skeva II, Mountokalakis TD (1999) Prevalence, awareness, treatment, and control of hypertension in Greece: the Didima study. *Am J Hypertens* 12: 959–965
45. Wolf K, Tuomilehto J, Kuulasmaa K et al. (1997) Blood pressure levels in the 41 populations of the WHO MONICA Project. *J Hum Hypertens* 11: 733–742
46. De Backer G, Myny K, De Henauw S et al. (1998) Prevalence, awareness, treatment and control of arterial hypertension in an elderly population in Belgium. *J Hum Hypertens* 12: 701–706
47. Leer EM van, Seidell JC, Kromhout D (1994) Levels and trends in blood pressure and prevalence and treatment of hypertension in the Netherlands, 1987–1991. *Am J Prev Med* 10: 194–199
48. Marques-Vidal P, Ruidavets JB, Cambou JP, Ferrieres J (2000) Trends in hypertension prevalence and management in Southwestern France, 1985–1996. *J Clin Epidemiol* 53: 1230–1235
49. Mulrow PJ (1998) Detection and control of hypertension in the population: the United States experience. *Am J Hypertens* 11: 744–746
50. Primatesta P, Brookes M, Poulter NR (2001) Improved hypertension management and control: results from the health survey for England 1998. *Hypertension* 38: 827–832
51. Saounatsou M, Patsi O, Fasoï G et al. (2001) The influence of the hypertensive patient's education in compliance with their medication. *Public Health Nurs* 18: 436–442
52. Huse DM, Roht LH, Alpert JS, Hartz SC (2001) Physicians' knowledge, attitudes, and practice of pharmacologic treatment of hypertension. *Ann Pharmacother* 35: 1173–1179
53. Oliveria SA, Lapuerta P, McCarthy BD et al. (2002) Physician-related barriers to the effective management of uncontrolled hypertension. *Arch Intern Med* 162: 413–420
54. Berlowitz DR, Ash AS, Hickey EC et al. (1998) Inadequate management of blood pressure in a hypertensive population. *N Engl J Med* 339: 1957–1963
55. Cohen JD (2002) Superior physicians and the treatment of hypertension. *Arch Intern Med* 162: 387–388
56. European IDDM Policy Group (1993) Consensus guidelines for the management of insulin-dependent (type 1) diabetes. *Diabet Med* 10: 990–1005
57. The National High Blood Pressure Education Program Working Group (1994) National High Blood Pressure Education Program Working Group report on hypertension in diabetes. *Hypertension* 23: 145–158
58. Krans HMJ, Porta M, Keen H, Staehr Johansen K (1995) Diabetes care and research in Europe: the St. Vincent Declaration action programme. Implementation document, WHO Regional Office for Europe, Copenhagen, pp 1–99
59. American Diabetes Association (1996) Treatment of hypertension in diabetes. *Diabetes Care* 19: S107–S113

60. Comitato per le Linee Guida (1999) Linee guida 1999 per il trattamento dell'Ipertensione arteriosa dell'Organizzazione Mondiale della Sanità (OMS) e della Società Internazionale dell'Ipertensione (ISH). *Ipertensione e Prevenzione Cardiovascolare* 6: 41–88
61. Agabiti Rosei E, Costa F, Morganti A et al. (1999) Appendice alle Linee Guida per il trattamento dell'Ipertensione Arteriosa a cura della Società Italiana dell'Ipertensione Arteriosa. *Ipertensione e Prevenzione Cardiovascolare* 6: 89–98
62. Ramsay LE, Williams B, Johnston GD et al. (1999) Guidelines for management of hypertension: report of the third working party of the British Hypertension society. *J Hum Hypertens* 13: 569–592
63. European Diabetes Policy Group (1999) A desktop guide to Type 1 (insulin-dependent) diabetes mellitus. European Diabetes Policy Group 1998. *Diabet Med* 16: 253–266
64. European Diabetes Policy Group (1999) A desktop guide to Type 2 diabetes mellitus. European Diabetes Policy Group 1999. *Diabet Med* 16: 716–730
65. Guidelines subcommittee (1999) 1999 World Health Organization-International Society of Hypertension Guidelines for the Management of Hypertension. Guidelines Subcommittee. *J Hypertens* 17: 151–183
66. ANAES (2000) Strategie de prise en charge du patient diabetique de type 2 a l'exclusion de la prise en charge des complications. Recommendations de l'ANAES Mars 2000. *Diabete Metab* 26: 1–96
67. American Diabetes Association (2002) Treatment of hypertension in adults with diabetes. *Diabetes Care* 25: 199–201

Table 1. Summary of guidelines on the management of hypertension in diabetes

	Intervention threshold		Target blood pressure	
	Systolic BP (mmHg)	Diastolic BP (mmHg)	Systolic BP (mmHg)	Diastolic BP (mmHg)
Consensus guidelines: European IDDM Policy Group [56]	1993		≤140	≤85
National High Blood Pressure Education Working Group [57]	1994	140	<130	<85
St. Vincent Declaration [58]				
Age <40 years or end-organ damage	1995	140	≤140	≤85
Age ≥ 40 years and no end-organ damage		160	≤140	≤85
American Diabetes Association [59]	1996	140	<130	<85
Joint National Committee VI [16]	1997	130	≤130	≤85
With proteinuria > 1 g/24 hours			≤125	≤75
Italian Hypertension guidelines (SIIA) (www.siiat.it/linee) [60, 61]	1999	140	<130	<85
British Hypertension Society [62]	1999	140	<140	<80
European Diabetes Policy Group: Type 1 diabetes with proteinuria [63]	1999	135	<130	<80
Desktop guide: European NIDDM Policy Group [64]	1999		<140	<85
World Health Organisation/International Society for Hypertension [65]	1999	140	<130	<85
Dutch hypertension guidelines (CBO) (http://www.cbo.nl/richtlijnen)	2000	140	<140	<90
French Hypertension recommendations: ANAES [66]	2000	140	<140	<80

	Intervention threshold	Target blood pressure	
		Systolic BP (mmHg)	Diastolic BP (mmHg)
Finnish Hypertension guidelines 2001 (www.duodecim.fi/kh)	140	<140	<80
German Hypertension guidelines 2002 (AWMF) (http://www.uni-duesseldorf.de, arterielle hypertonie leitlinien)	140	<135	<85
With proteinuria > 1 g/day		<125	<75
American Diabetes Association [67]	140	<130	<80

Table 2. Baseline and follow-up characteristics of subjects included and lost at follow-up

	1989–1990 cohort (n= 1866)	Lost at follow-up (n=1384)	1997–1999 cohort (n=1866)
	Baseline risk factors	Baseline risk factors	Follow-up risk factors
	Means \pm SD	Means \pm SD	Means \pm SD
Age (years)	33 \pm 10	33 \pm 11	40 \pm 10
Duration of diabetes (years)	15 \pm 9	15 \pm 10	22 \pm 9
Systolic BP (mm Hg)	120 \pm 17	123 \pm 19	121 \pm 19
Diastolic BP (mm Hg)	75 \pm 11	76 \pm 12	74 \pm 12
BMI (kg/m ²)	23.6 \pm 2.8	23.4 \pm 3.0	24.6 \pm 3.3
WHR	0.84 \pm 0.10	0.85 \pm 0.12	0.89 \pm 0.15
HbA _{1c} (%) ^a	8.3 \pm 1.9	8.7 \pm 2.0	8.3 \pm 1.4
Cholesterol (mmol/l)	5.3 \pm 1.1	5.4 \pm 1.2	5.3 \pm 1.2
LDL-cholesterol (mmol/l)	3.3 \pm 0.9	3.5 \pm 1.1	3.2 \pm 1.0
Fasting triglyceride (mmol/l) ^e	1.0 \pm 1.6	1.1 \pm 1.7	1.1 \pm 1.7
	% (95%CI)	% (95%CI)	% (95%CI)
Men	52	51	52
Current smokers	30	36	29
Albuminuria	30	32	28
Retinopathy	45	48	73
Cardiovascular disease	8	12	11
Hypertensives ^b	22 (20–24)	27 (25–29)	34 (32–36)
Treated with BP lowering drugs ^d	40 (35–45)	45 (40–50)	69 (65–73)
Controlled ^{c,d}	12 (9–16)	10 (7–14)	28 (24–32)
Controlled ^{c,e}	32 (25–39)	23 (17–30)	41 (36–45)

^aCorrected HbA_{1c} values according to DCCT method

^bHypertension: defined as a SBP \geq 140 mmHg or DBP \geq 90 mmHg or the use of blood pressure lowering drugs

^cControl is defined as BP<130/85 mmHg

^dProportion of hypertensives

^eGeometric mean (log-transformation)

Table 3. Frequency of hypertensives and management of hypertension by age groups

	1989–1990				1997–1999				
	Age at baseline	<i>n</i>	Men	Women	Total	<i>n</i>	Men	Women	Total
Hypertensives ^a	≤25	471	12.6%	10.8%	11.7% (9–15)	471	20.9%	15.5%	18.3% (15–22)
	25–35	727	18.3%	13.7%	16.2% (14–19)	727	31.4%	21.9%	27.1% (24–30)
	35–45	431	33.0%	25.0%	29.0% (25–34)	431	48.4%	41.2%	44.8% (40–50)
	45–55	202	49.0%	46.2%	47.5% (40–55)	202	64.6%	63.2%	63.9% (57–70)
	>55	35	33.3%	78.6%	51.4% (34–69)	35	66.7%	85.7%	74.3% (57–88)
Treated hypertensives ^b	≤25	55	27.6%	37.5%	32.1% (20–46)	86	52.0%	82.9%	64.7% (54–75)
	25–35	118	31.5%	28.9%	30.5% (22–40)	197	67.2%	68.1%	67.5% (60–74)
	35–45	125	50.0%	35.2%	43.6% (35–53)	193	77.9%	66.3%	72.5% (66–79)
	45–55	96	51.1%	55.1%	53.1% (43–63)	129	68.9%	78.8%	74.0% (65–81)
	>55	18	14.3%	36.4%	27.8% (10–53)	26	35.7%	58.3%	46.2% (27–67)
Controlled (treated) hypertensives ^{c, d}	≤25	17	25.0%	77.8%	52.9% (28–77)	55	57.7%	58.6%	58.2% (44–71)
	25–35	36	30.4%	53.9%	38.9% (23–57)	133	44.1%	51.0%	46.6% (38–55)
	35–45	53	23.5%	26.3%	24.5% (14–38)	140	30.9%	42.4%	35.7% (28–44)
	45–55	50	25.0%	34.6%	30.0% (18–45)	94	33.3%	30.8%	31.9% (23–42)
	>55	5	0%	0%	0%	12	0%	28.6%	16.7% (2–48)

% (95%CI)

^aHypertension: defined as a SBP≥140 mmHg or DBP≥90 mmHg or the use of blood pressure lowering drugs

^bAs a percentage of those with hypertension

^cControl is defined as BP<130/85 mmHg

^dAs a percentage of treated hypertensives

Table 4. Change in prevalence, treatment and control of hypertension by albuminuric status, between 1989–1990 and 1997–1999

Albuminuria	1989–90			1997–99				
	<i>n</i>	Men	Women	Total	Men	Women	Total	
Hypertensives ^a		925	869	1794	925	869	1794	
	Normo	1260	14.4%	16.0%	15.2% (13–17)	1260	24.3%	23.8% (21–26)
	Micro	398	29.5%	28.2%	28.9% (24–34)	398	48.7%	46.0% (41–51)
Treated hypertensives ^b	Macro	136	72.3%	52.8%	64.7% (56–73)	136	94.0%	89.7% (83–94)
	Normo	192	27.3%	35.3%	31.6% (25–39)	300	47.0%	55.0% (49–61)
	Micro	115	35.4%	34.7%	35.1% (26–45)	183	73.4%	75.8% (69–82)
Controlled (treated hypertensives) ^{c,d}	Macro	88	65.0%	60.7%	63.6% (53–74)	122	94.9%	95.1% (90–98)
	Normo	60	33.3%	40.0%	37.3% (25–51)	164	41.4%	38.4% (31–46)
	Micro	40	26.1%	47.1%	35.0% (21–52)	138	43.8%	47.1% (39–56)
Macro	56	21.1%	23.5%	21.8% (12–35)	116	31.1%	36.2% (27–46)	

% (95%CI)

^aHypertension: defined as a SBP \geq 140 mmHg or DBP \geq 90 mmHg or the use of blood pressure lowering drugs

^bAs a percentage of those with hypertension

^cControl is defined as BP<130/85 mmHg

^dAs a percentage of treated hypertensives

Table 5. Distribution of blood pressure lowering drugs in 1989/90 and 1997/99

	1989–1990 (n=310)	1997–1999 (n=435)
ACE inhibitor	57% (n=178)	82% (n=355)
CCB	22% (n=67)	25% (n=109)
Beta Blocker	17% (n=54)	11% (n=48)
Diuretic	9% (n=27)	15% (n=69)
Alpha Blocker	4% (n=11)	6% (n=24)
Other	4% (n=11)	6% (n=27)