www.nature.com/iio

Barriers to good glycaemic control: the patient's perspective

FJ Snoek¹*

¹Department of Medical Psychology, Vrije Universiteit Medical Centre, Amsterdam, The Netherlands

Diabetes currently affects at least 120 million people worldwide, and this figure is rising steadily. Intensive treatment improves outcome in terms of morbidity from late diabetic complications and quality of life, but in order for patients to reap such benefits, they must commit to major, long-term changes in lifestyle. The physician's concept of diabetes is often very different from the patient's; and the implementation of a treatment plan acceptable to both is only possible when open communication fosters discussion and patient autonomy, and treatment is seen as logical, acceptable and feasible within the daily life of each patient.

Barriers that impair patients' ability to achieve good glycaemic control include those relating to lifestyle, education, psychology and their environment. An appreciation of barriers to good glycaemic control from the patient's perspective underlies the ability to minimise obstacles and improve outcome in terms of quality of life and metabolic control.

International Journal of Obesity (2000) 24, Suppl 3, S12-S20

Keywords: self-management; metabolic control; quality of life; education; hypoglycaemia; barriers to treatment

Is diabetes the same disease for doctor and patient?

Diabetes currently affects at least 120 million people worldwide, and by 2010 this number is estimated to have reached 220 million.¹ With per capita diabetesrelated expenditure running at four-times that for nondiabetics in the USA,² and total costs accounting for 28% of the annual healthcare budget allocated to the over 65-y-olds,³ the cost in human and financial terms is immense. Large epidemiological studies have shown that intensive treatment significantly reduces morbidity from late diabetic complications.4-7 The challenge is to make it possible for the physician and patient to achieve the best outcome in terms of metabolic control, while ensuring a good quality of life for the patient.

A diagnosis of diabetes poses very different challenges to the patient and physician. The physician begins the process of advising on dietary change, exercise, weight control and the most appropriate choice of medication where necessary. He/she will explain the serious nature of the disease, the potential for dangerous long-term complications and the necessity of maintaining tight blood glucose control at all times. The patient hearing a diagnosis of 'diabetes' receives a litany of advice and guidelines, all of which necessitate dramatic changes in every realm of their life. They may have heard of diabetes because a relative became blind or had a limb amputated as a

result of the disease. To some, diabetes will have connotations of eating healthy food or buying diabetic sweets. To others it may signify early death. Whatever the initial image, when presented with the broadranging and life-long changes required, the likelihood is that 'diabetes' conjures up a very different picture for physician and patient.

High-quality diabetes care requires physician and patient to share an understanding of the distinct but complementary roles each will fulfil. Each party approaches the medical consultation with a different agenda, but like any negotiation, success rests on both parties participating in the decision-making process. Basic disease pathology, natural history and the need for active self-management are concepts that the physician must make accessible to the patient before satisfactory glycaemic control is possible.

Diabetes: a life-long self-management issue

The diagnosis of diabetes confers on the individual a life-long requirement to modify their lifestyle, monitor their disease and control their blood glucose via various treatment modalities. The disappointing standards of blood glucose control stand witness to the difficulty of this task: recent data from three countries indicated that $\geq 60\%$ of patients with Type 2 diabetes have $HbA_{1c} > 7.5\%$ (as discussed by Nattrass in this publication⁸).

In order for the daily routines of diabetes care to be effectively implemented, 'diabetes-related' tasks must make sense to the patient and be acceptable and realistic at a personal and cultural level. The targets that we set for diabetic patients include some that

^{*}Correspondence: FJ Snoek, Department of Medical Psychology, Diabetes Research Centre, Vrije Universiteit Medical Centre, Amsterdam, The Netherlands.

This supplement of the International Journal of Obesity was sponsored by Novo Nordisk A/S.

burden even for the young, whose memories are reliable and who are spurred on by an awareness of the investment they are making for their future health; obese patients attempting to lose weight struggle with their undertaking despite specialist advice, medication, and even sometimes surgery. The tasks set for diabetic patients include both daily medication-taking and losing weight, but to this are added dietary changes, the aspect of self-management, which is generally found most difficult by patients,⁹ smoking and alcohol restrictions, and home blood glucose monitoring, among others. Taking medication has been found less burdensome than some other tasks,⁹ perhaps reflecting deeply ingrained beliefs in society about the patient's responsibilities when taking on the 'sick role'.

As is the case with other chronic diseases, the prospect of treatment can seem more punishing than the disease itself. The potential for long-term diabetic complications may appear less real than the restrictions placed on eating and the daily self-injecting with insulin for those who require it. Blood glucose monitoring does not in itself improve diabetes, and yet adds to the array of 'diabetes-related' practices required on a daily basis.

Factors that influence successful patient self-management were investigated using a 41-question questionnaire.¹⁰ Five factors were found to relate significantly to glycaemic control as measured by HbA_{1c}: (1) practical self-management skills; (2) emotional adjustment; (3) perceived goals; (4) perceived self-efficacy; and (5) cost-benefit analysis. These results should engender optimism since, unlike fixed demographic variables accounting for risk, all these factors can be altered. From the patients' point of view, targets in this study appeared to fall into two broad categories: those relating to blood glucose control (medical goals) and those concerning living with diabetes. Individual patients ascribe different degrees of importance to each target. A role of the physician is to facilitate individual goals, with the aim of improving glycaemic control and psychological adjustment to life with diabetes.

Diabetes and quality of life

'Quality of life', measured by scales including QALYs (Quality Adjusted Life Years), is seen by some as a scientific measure of the value of daily life, others regard it as a nebulous assessment of general well-being. Diabetes-related quality of life (QoL) includes measures of symptom distress, emotional well-being, functional limitations, cognitive performance and social relationships. Tools for the measurement of QoL include generic scales (such as the SF-36, GHQ-12, NHP and SIP) and those measuring diabetes-specific variables:

- 1. DQOL (psychosocial impact of disease)
- 2. DHP (distress, barriers and eating restraints)

- 3. PAID (diabetes-related distress)
- 4. DSC-2 (diabetes-related symptoms)

QoL is a useful surrogate marker for a multitude of factors and gives some idea of how well a patient is coping with diabetes overall. Few would deny that a patient whose life is dominated by diabetes-related restrictions is not a 'diabetes success', no matter how good their metabolic control. Conceptualising the disease from the patient's perspective requires an understanding of the factors that contribute to quality of life.

Achieving good metabolic control: hurdles and hoops

The maintenance of blood glucose concentration within recommended limits is a task that requires patients to juggle numerous responsibilities. Barriers to achieving the desired outcome can be divided into the following categories, between which there is significant overlap:

- Lifestyle
- Educational
- Psychological
- Environmental

Lifestyle barriers. The lifestyle limitations imposed on diabetic patients originate partly from the disease itself and partly from the treatment. Traditional longacting oral hypoglycaemic agents (OHAs) including the sulphonylureas stimulate pancreatic insulin production from their onset of action until renal or hepatic metabolism inactivates the drug, which is then excreted. Duration of action of drugs is variable, but for the longer-acting preparations there are periods when a relative excess of insulin increases the danger of hypoglycaemia. The commonest risk periods are between meals and during the night, when insulin action is unopposed by food intake.

Between-meal snacks are recommended to offset the long duration of action of some OHAs and reduce the chance of hypoglycaemia. Since one-third of all hypoglycaemic episodes requiring hospital admission in people taking sulphonylureas are caused by missed meals (as discussed by Nattrass and Lauritzen¹¹ in this publication), and missed or delayed meals are common,¹² the risk of hypoglycaemia is not insignificant. Over-zealous glucose reduction following mismatched treatment/exercise and food intake also compromise stable glycaemic control. Fear of precipitating hypoglycaemia results for many in a fearful tightrope walk between hyper- and hypoglycaemia. Despite warnings from the physician about the dangers of elevated glucose, many patients would rather err on the side of hyperglycaemia than risk the consequences of an unexpected 'hypo'. Anxiety from family members intensifies the patient's fears, increasing the pressure to walk this fine line precisely.

Although frequent snacking helps avoid hypoglycaemia, it increases the potential for weight gain in a

torv

advice.

To a newly diagnosed patient the grave long-term risks seem far removed from real life. Added to this, difficulty carrying out practical self-management

skills such as diet planning, home blood glucose monitoring, record-keeping and injection technique foster resentment and can reduce compliance and commitment to learning. One skill with important implications is the detection of incipient hypo-

physiological warning mechanisms.

glycaemia, a task made more difficult by defective

Hypoglycaemic unawareness: a specific physiological barrier. Tight glycaemic control is constrained by the requirement to protect the brain from the devastating effects of hypoglycaemia at all costs. Type 1 diabetic patients are at greater risk of hypoglycaemia, but Type 2 diabetic patients experience it with increasing frequency and severity over time and duration of illness.^{7,13} Were the sequelae not so severe, current treatments could lower blood glucose levels and reduce long-term complications extremely effectively. However, hypoglycaemic episodes are dangerous: in the short-term, seizures, coma and death are possible, and mortality rate from sulphonylurea-induced hypoglycaemia is as high as 10%. In the longer term, 5% of survivors of a sulphonylureainduced hypoglycaemic episode will have permanent neurological sequelae.14 Treatment strategies must anticipate and minimise these risks.

Unfortunately, asymptomatic hypoglycaemic episodes are common. Unawareness of the warning signs of hypoglycaemia results from reduced secretion of insulin, glucagon and adrenaline seen in Type 1 diabetes¹⁵ and impaired response to adrenaline.¹⁶ Diminished adrenaline response to reduced glucose concentration, a phenomenon termed 'defective glucose counter-regulation', is responsible for loss of the autonomic warning signs of hypoglycaemia known as 'hypoglycaemic unawareness'.¹⁷ The increased incidence of severe hypoglycaemia during intensive treatment of Type 1 diabetic patients¹⁶ is also attributable to this. It is suggested, but not yet confirmed, that the same mechanisms contribute to these phenomena in Type 2 diabetes.

Both hypoglycaemic unawareness and the adrenaline deficit in defective glucose counter-regulation follow recent iatrogenic hypoglycaemia,^{15,18–20} which itself further reduces autonomic warning signs of low blood glucose.^{15,16} Physiological defences against developing hypoglycaemia are impaired and a vicious circle of recurrent hypoglycaemia is set up (Figure 1). This can be broken, with restoration of full awareness, if hypoglycaemia is avoided for just two weeks.^{18,19} Training in recognition of early warning signs of reduced blood glucose can reduce fear and increase confidence, and can go some way towards reducing this particular barrier to good glycaemic control.

Psychological barriers. Every chronic disease challenges the patient with a new identity over which they

population already struggling to counteract the anabolic

effects of insulin produced by the secret gogue action of

OHAs. The UKPDS study group investigated weight

gain with antidiabetic therapies and found increases

varied from 2.5 kg with diet treatment alone to 7.5 kg with chlorpropamide, over the 15-y study period.⁷ The

weight gain induced by glibenclamide and insulin fell

between these levels. In all cases, weight gain began soon after initiation of treatment. In no cases did weight remain unchanged and no weight loss was observed. As

weight increases, patients lose the physiological oppor-

tunity to improve glycaemic control by weight modera-

tion. Feelings of failure and despondency can

jeopardize further attempts to lose weight, resulting in

The reality of daily obstacles to good metabolic

control was investigated in an observational study

of 123 non-insulin-dependent patients.¹² Quantit-

ative data on lifestyle, eating habits and diabetes

information resources were gathered from focus

groups of 8-10 members drawn from Europe and the

USA. Qualitative data were gathered from questions

and questionnaires at screening and during focus group

discussions. Results showed widespread dissatis-

faction with the lack of information provided to

diabetic patients: only 20% were satisfied with the

amount they received. There was uniform disappoint-

ment at the emphasis on restrictive diet and lifestyle

changes rather than diabetes as a disease. US patients

tended to be more satisfied with the information

received, praising the quality of resources provided

by the American Diabetes Association, whereas 91%

of Spanish patients were frustrated by lack of

resources. Food recommendations were felt to be

restrictive and poorly adapted to individual needs,

and compliance with dietary advice was difficult for

all. Despite recognising the importance of regular

meals, only 28% were able to comply with this (18%

in the USA). In addition, healthcare providers

were frequently experienced as unsupportive when

discussing difficulties in adhering to meal-related

Educational barriers. Educational barriers to good

metabolic control include lack of knowledge and

practical skills. Inadequate information about basic

diabetes pathophysiology precludes an understanding

of the rationale for treatment; improved glycaemic

control is then thwarted by misconceptions and inap-

propriate behavioural modifications.¹² One of the

commonest misconceptions is that eating regularly is

a part of diabetes *treatment*, rather than a compensa-

acting OHAs.12 This can work against attempted

by

the

long-

behaviour necessitated

a spiral of deteriorating glycaemic control.



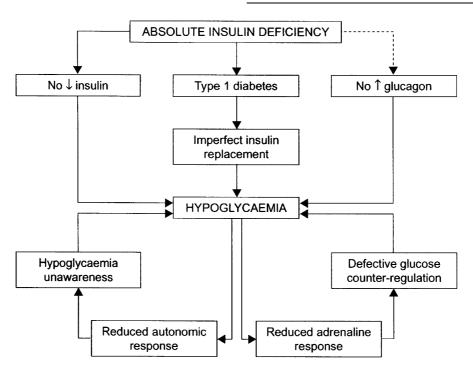


Figure 1 Schematic diagram of the concept of hypoglycaemia-associated autonomic failure in diabetes, and the role of iatrogenic hypoglycaemia in the pathogenesis of hypoglycaemic unawareness and defective glucose counter-regulation.²⁰

have no control, but by which their lives are defined in some way. Despite statements that equal opportunities guide employment decisions and that we should look beyond the disease to the individual, chronic diseases still carry stigma. Fear, anger, frustration and countless additional emotions can afflict the patient with a lifelong disease. The psychological demands can seem overwhelming and psychological aspects of diabetes care are important in improving quality of life.

In addition to coping with the psychological difficulties presented by any chronic disease, attaining specific diabetes-related targets is a daily struggle. The perceived inability to alter outcome in any obvious way can remove any sense of potency. For some, the changes required and the difficulty in achieving set goals becomes more handicapping than having diabetes in the first place.

Up to 33% of people with diabetes suffer an episode of major depression during their lifetime, a figure significantly greater in statistical terms than that for the general population.²¹ Major depression has significant impact on morbidity, all-cause mortality, functional disability, absenteeism from work and increased healthcare expenditure. Depression in diabetes is generally severe and recurrent, and $\leq 10\%$ who achieve remission remain symptom-free for the subsequent five years.²¹ Women are affected more frequently, as in the non-diabetic population, but no difference in incidence between Type 1 and Type 2 diabetes has been shown.

Depression has a complex physiological relationship with diabetes and *vice versa* (Figure 2). Prospective and cross-sectional studies have linked depression to glucose dysregulation, with a significant 1.8% increase in HbA_{1c} being directly attributable to depression.²² Depression worsens glucose control directly as well as via obesity, physical inactivity and treatment noncompliance. In turn, animal studies have shown hyperglycaemia to alter mood via disruption of hypothalamic-pituitary-adrenocortical (HPA) axis activity²³ and 5-HT (serotonin) sensitivity.²³ 5-HT deficiency is a known risk factor for depressive disorder and disrupted HPA responsiveness that is reported in diabetes may be responsible for the increased reactivity to stress seen in diabetic subjects.²⁴

Depression is an independent risk factor for increased diabetic complications, particularly macrovascular disease^{22,26} and retinopathy.²⁷ Other independent correlates such as time spent in poor glycaemic control and duration of diabetes are additive risks.²⁷ Predictors of response to treatment for

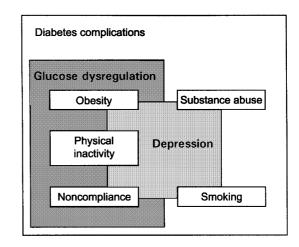


Figure 2 The relationship between behavioural and medical factors in diabetes. All intersecting boxes are statistically significant associations.²⁵

Barriers to good glycaemic control FJ Snoek

depression were tested in a randomised clinical trial of cognitive behavioural therapy (CBT) in which a total of 42 Type 2 diabetic patients took part.²¹ Non-remission of depression was associated with poorer compliance with blood glucose monitoring, higher HbA_{1c} and weight, and previous treatment for depression. In the group receiving CBT, diabetic complications and poor compliance with blood glucose monitoring were significant independent predictors of poor response. Such evidence suggests that factors relating to the medical illness impact negatively on prognosis for recovery from depression.

Sadly, many patients and physicians subscribe to the belief that depression is the natural response to a diagnosis of diabetes and to the effects of hyperglycaemia. Certainly hyperglycaemia can produce similar symptoms to depression (headaches, exhaustion). It can also worsen the prognosis for recovery from depression by masking symptoms, improvement and adverse drug interactions. However, treatment that focuses purely on glycaemic control fails to target depression. The composite evidence suggests that depression should be treated as part of the medical illness, with psychoactive medication accompanying efforts to improve glycaemic control.

If an antidepressant is required, the choice should take into account potential interactions with any aspect of diabetes or its treatment. Many psychotropic agents directly elevate or reduce blood glucose levels. A double-blind, placebo-controlled trial of nortriptyline in depressed, poorly-controlled diabetic patients demonstrated significant, sustained elevations in blood glucose that were not accounted for by the tendency of tricyclic antidepressants to induce weight gain.²⁸ Patients with generalised anxiety disorder (GAD) have benefited from the direct hypoglycaemic effect of alprazolam on glucose levels, independent from the effect on GAD.²⁹ These studies amongst others highlight the need to consider the effects on blood glucose of any psychoactive agent being considered.

Environmental barriers. Good glycaemic control relies on patient commitment to self-management, but this can and should be strengthened by external support. Even devout efforts to improve metabolic control can be defeated by negative judgements and lack of practical and emotional support from family, friends, employers and healthcare providers. Scarce personal finances or lack of institutional resources can also compromise outcome. Provision of high-quality, patient-centred care requires support in all these arenas: personal, financial and organisational.

Overcoming the barriers

Education can improve outcome

In order to provide useful, pertinent information to patients it is necessary to know the degree of understanding the patient has, and where there is room for improvement. Presenting information in an acceptable format that shows sensitivity to cultural and dietary customs is likely to improve the chance of agreeing a mutually acceptable treatment plan.

Successful educational interventions take various forms, from information leaflets and weekly lectures to intensive inpatient workshops. It has been suggested that four main outcomes should be used in evaluating the success of an educational intervention:³⁰

- Patient knowledge (about diabetes)
- Patient behaviour (including self-management skills)
- Patient attitude (emotional aspects of living with diabetes)
- Patient control of blood glucose

To investigate the effectiveness of patient education in improving diabetic outcomes, Brown performed a meta-analysis of 82 studies evaluating the effects of adult diabetes education on knowledge, self-care behaviours, psychological outcomes and metabolic control (Figure 3).³¹

Diabetes patient education was found to have a moderate to large effect on improving both general and diet-specific knowledge. Despite a three-fold increase in self-reported dietary compliance, weight was only minimally improved. This may be explained by the lag time between diet alteration and weight change, and by the inevitable inaccuracy of selfreported data. Practical self-care skills improved minimally to moderately overall, with urine-testing skill showing greater improvement than insulin injection technique. Measures of metabolic control benefited moderately. Glycosylated haemoglobin showed the greatest improvement, and insulin dose (an indirect measure of metabolic control) the least. Psychological outcome benefited little from educational interventions, probably reflecting the lack of specific counselling or therapy among most of the studies included.

The theory behind educating patients is that longterm prognosis can be improved if patients understand the need for, and methods involved in, maintaining tight glucose control. However, knowledge and behaviour are notoriously poorly correlated32 and improved knowledge following educational intervention does not always translate into improved metabolic control.33 Korhonen and colleagues34 demonstrated highly significant improvement in diabetic control following intensive patient education, but on 18month follow-up no difference from the control group could be observed. Good control correlated not with education, but with a low score for anxiety and depression at intake, and a high score for selfconfidence. However, in the Diabetes Education Study (DIABEDS),^{35,36} a randomised controlled trial of patient and physician education, greater and statistically significant reductions in fasting blood glucose, HbA_{1c}, body weight and blood pressure were seen in the patient education group 11-14 months after inter-

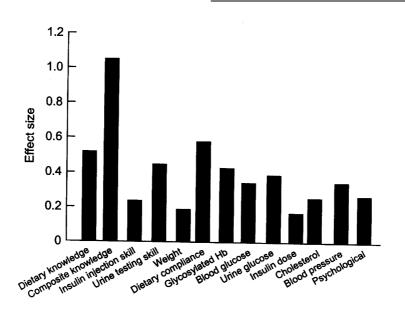


Figure 3 Results of an 82-study meta-analysis. The effect of patient education on knowledge, self-care behaviours, metabolic control and psychological outcomes. (Data are weighted mean effect sizes: 0.2 = small effect size; 0.5 = medium; 0.8 = large).³¹

vention, despite poor knowledge retention. The identical intervention for clinicians resulted in significant reductions in patient fasting glucose, HbA_{1c} and body weight. The greatest improvements in overall outcome followed combined physician and patient education.

The specific needs of the elderly. The greater challenge of altering learning and behaviour in the elderly is highlighted by the inverse relationship between age and acquisition of new knowledge.³¹ This underlines the need for age-appropriate education, but also reveals a more general theme concerning specific barriers to improving metabolic control in the elderly. Psychological and physiological changes threaten good control in this population. While increasing ill health necessitates poly-pharmacy, declining hepatic and renal function favour drug accumulation and toxicity. Prescribing certain medications becomes unwise. The requirement for multiple drugs complicates the patient's daily routine and the physician's attempts to simplify treatment.

Many measures of cognitive performance are slowed or reduced as age increases. Remembering to take tablets, inject insulin or make an appointment to see the chiropodist become a supreme effort, sometimes compounded by anxiety not to burden younger relatives with requests for help. Good glycaemic control is more elusive as age advances and the motivation to change is undermined by thoughts such as, 'What's the point? I may not live to see the benefits anyway'. In an era when sophisticated medical treatments allow increasing longevity, it is crucial that the specific needs of the elderly are addressed.

Structure of care

Integration of education and clinical care facilitates high-quality diabetes management across the age spectrum. In an integrated care structure: teaching and counselling support clinical aspects of treatment; newly learned skills and attitudes are explored in a supportive environment; and nurse educators, dieticians, psychologists and chiropodists work alongside physicians. In this setting significant decreases in rates of hospitalisation and acute complications are possible.³⁷

The benefits of integrated care were demonstrated in Ipswich, UK, following the opening of a purposebuilt diabetes centre in which the role of the diabetes specialist nurse was strengthened.³⁸ A progressive, sustained and statistically significant improvement in glycaemic control was seen in all patients attending the clinic, including the elderly. Annual admission rates for ketoacidosis and hypoglycaemia fell in a highly significant manner. In addition, fewer patients failed to attend appointments, suggesting increased patient satisfaction, possibly relating to reduced waiting times. It was possible to cut waiting times because nurses agreed to see those patients who did not require physician input. Purpose-built diabetes inpatient and day centres, in which diabetes specialist nurses play a key role, have become increasingly common in recent years. They are a valuable resource for a population in which the prevalence of disease is increasing¹ and routine hospital-based care for all is not a practical option.

Current practice in many countries is for diabetes care to be either general practitioner (GP)/community-based, hospital-based or shared between the two. Inefficient or inaccessible care provision is undoubtedly a barrier to patients who seek to improve their metabolic control. General measures that improve efficiency and patient outcome in all care settings include the 3 Rs-registration, recall and regular review (audit).³⁹ The success of shared care schemes has been shown to depend on a centralised prompting Barriers to good glycaemic control FJ Snoek

system for patient appointments and a structured checklist for the GP. Structured GP care comprising systematic recall of patients, and adherence to a standard protocol carried out by a GP with a special interest in diabetes, can provide comparable, sometimes superior care to hospital practice.³⁹ Once a satisfactory care structure has been set up, patient knowledge, skills and confidence can be enhanced and targets achieved more easily.

Empowering patients

It has been said that 95% of diabetes management is carried out by the patient.⁴⁰ In recent years 'empowerment', 'the process by which people gain mastery over their own affairs',⁴¹ has become an increasingly applied alternative to the physician-led, patient-compliance model.⁴² The empowerment model, applicable in many healthcare situations including diabetes, emphasises personal strengths, shared goals, shared decision-making and self-generated solutions to problems. The patient identifies individual problems and needs, and learns strategies to allow maladaptive behaviours to be challenged and newer, healthier ones implemented. As confidence increases, a feeling of mastery boosts the motivation to succeed. The empowerment model offers advantages for the healthcare provider too, reducing the burden of responsibility and power, and shifting the focus towards shared care.

The value of empowerment in diabetes care is highlighted by the superior outcomes achieved over traditional, more paternalistic approaches. When taught how to negotiate with and ask questions of their physician, patients become more assertive and able to elicit information. They experience fewer functional limitations, including days lost from work, and can achieve significant reductions in HbA_{1c} (Figure 4).⁴³ Physician support for patient autonomy increases feelings of competence and motivation to take prescribed medication⁴⁴ and correlates with improved HbA_{1c}.^{31,43,45} In contrast, nurses perceived as controlling and directive in their communication are associated with patients with poorer metabolic control (P < 0.01).⁴⁵

Empowerment, like any model of care, can have limitations. Patients who find it difficult to take responsibility for other areas of their lives may find it no easier to do so with their health. While empowerment respects individual decisions concerning approaches to treatment, it does not prohibit physicians from questioning or challenging seemingly irrational decisions. Nor does it remove from them the responsibility to give constructive, supportive advice and prescribe the treatment that their professional training indicates is appropriate. Patient decisions should be discussed within the framework of goals and values, needs, health beliefs and culture, which the patient brings to the consultation. Exploring difficulties in this manner can reveal socio-cultural atti-

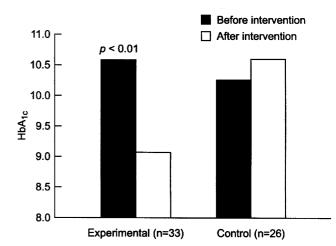


Figure 4 Effects of patient participation in care on glycosylated haemoglobin.⁴³

tudes that fully explain a choice that seemed foolish or illogical at first glance.

The relevance of rational drug treatment

Whichever model of care an individual's treatment is based on, drug therapy is likely to be an important aspect at some stage. By reducing the extent to which diabetic treatments impose undesirable routines and side-effects the ideal drug would reduce several important barriers to good glycaemic control.

Ideally the patient's meal pattern would dictate treatment and not vice versa. As well as increasing lifestyle flexibility this would make it easier to remember to take medication. If a drug could recreate a physiological pattern of insulin release, tailoring availability to need, the risk of hypoglycaemia would be reduced. The avoidance of iatrogenic hypoglycaemia would then lessen hypoglycaemic unawareness and reverse defective glucose counter-regulation. By limiting insulin availability to times of need, the body would be exposed to the anabolic effects of insulin for less time, making weight control easier to achieve. A short-acting prandial insulin profile would also improve metabolic control by restoring insulin sensitivity through weight loss achieved by the reduced need for snacking.

The challenge of achieving good glycaemic control with minimal risk of hypoglycaemia and weight gain may be met by a new class of antidiabetic drug: the prandial glucose regulators. Studies of flexible meal-time dosing with one such drug, repaglinide, demonstrate improved metabolic control with non-significant or absent weight gain over 16 weeks.⁴⁶ Variable meal patterns (two, three or four meals/day) are also consistent with improved metabolic control and weight loss of up to 2.4 kg (P < 0.03).⁴⁶ Four one-y comparative double-blind studies showed hypoglycaemia with repaglinide to be reduced compared with sulphony-lureas (P < 0.03). Omission of the lunchtime meal caused no severe hypoglycaemic episodes with repa-

glinide, but glibenclamide was associated with 24% severe hypoglycaemic events.⁴⁷

Prandial glucose regulation is a promising new approach to diabetic drug treatment and has produced a valuable addition to the existing formulary. Perhaps repaglinide will be the first of many contenders for the coveted title, 'The patient's choice of therapy in diabetes care'.

Conclusions

The barriers to achieving tight control of blood glucose are numerous but not insurmountable. Factors relating to lifestyle, education, physiology, psychology and environment all contribute to the way in which an individual copes with disease. In trying to reduce the obstacles met by patients, it is effective to integrate appropriate education into structured clinical care, prescribe thoughtfully and rationally, and strive to empower patients by supporting their autonomy.

Quality of life is an important endpoint in itself but also plays a part in improving glycaemic control. If diabetes does not restrict the lives of patients but becomes a challenge met with enthusiasm and determination, both patient and physician will reap the benefits.

References

- 1 Amos A, McCarty D, Zimmet P. The rising global burden of diabetes and its complications: estimates and projections to the year 2010. *Diabet Med* 1997; **14**: S1–85.
- 2 American Diabetes Association. Ray N, Thamer M, Gardner E, Chan J. Economic consequences of diabetes mellitus in the US in 1997: report from the American Diabetes Association. *Diabetes Care* 1998; **21**: 296–309.
- 3 Ratner R. Type 2 diabetes mellitus: the grand overview. *Diabetic Med* 1998; **15**(Suppl 4): S4–S7.
- 4 Diabetes Control and Complications Trial Research Group. The effect of intensive treatment of diabetes on the development and progression of long-term complications in insulindependent diabetes mellitus. *New Engl J Med* 1993; **329**: 977–986.
- 5 Reichard P, Berglund B, Britz A, Cars I, Nilsson BY, Rosenqvist U. Intensified conventional insulin treatment retards the microvascular complications of insulin-dependent diabetes mellitus (IDDM): The Stockholm Diabetes Intervention Study (SDIS) after 5 y. J Intern Med 1991; 230: 101–108.
- 6 UK Prospective Diabetes Study (UKPDS) Group. UK Prospective Diabetes Study 6. Complications in newly diagnosed Type 2 diabetic patients and their association with different clinical and biochemical risk factors. *Diabetes Res* 1991; 13: 1-11.
- 7 UK Prospective Diabetes Study (UKPDS) Group. Intensive blood glucose control with sulphonylureas or insulin compared with conventional treatment and risk of complications in patients with Type 2 diabetes (UKPDS 33). *Lancet* 1998; **352**: 837–853.
- 8 Nattrass M. The theory of treating Type 2 diabetes. *Int J Obes Relat Metab Disord* 2000; **24**(Suppl 3): S2–S5.
- 9 Ary DV, Toobert D, Wilson W, Glasgow RE. Patient perspective on factors contributing to nonadherence to diabetes regime. *Diabetes Care* 1986; **9**: 168–172.

- 10 Day J, Bodmer C, Dunn O. Development of a questionnaire identifying factors responsible for successful self-management of insulin-treated diabetes. *Diabetic Med* 1996; 13: 564–573.
- 11 Nattrass M, Lauritzen T. Review of prandial glucose regulation with repaglinide: a solution to the problem of hypoglycaemia in the treatment of Type 2 diabetes? *Int J Obes Relat Metab Disord* 2000; **24**(Suppl 3): S21–S31.
- 12 Brown Frandsen K, Smedegaard Kristensen J. Compliance with, and understanding of, mealtime advice in patients with Type 2 diabetes. *Diabetes* 2000; **49**(Suppl 1): A176.
- 13 UK Prospective Diabetes Study (UKPDS) Group. United Kingdom Prospective Diabetes Study 24: a 6-year randomised, controlled trial comparing sulfonylurea, insulin and metformin therapy in patients with newly diagnosed Type 2 diabetes that could not be controlled with diet therapy. *Ann Intern Med* 1998; **128**: 165–175.
- 14 Gerich J. Oral hypoglycaemic agents. New Engl J Med 1989;321: 1231-1245.
- 15 Cryer P. Hypoglycaemia in diabetes mellitus. *Hypoglycaemia: Pathophysiology, Diagnosis and Treatment*. Oxford University Press: New York, 1997, pp 91–125.
- 16 Dagogo-Jack S, Craft S, Cryer P. Hypoglycaemia-associated autonomic failure in insulin-dependent diabetes mellitus. Recent antecedent hypoglycaemia reduces autonomic responses to, symptoms of, and defence against subsequent hypoglycaemia. J Clin Invest 1993; 91: 819–828.
- 17 Gold A, MacLeod K, Frier B. Frequency of severe hypoglycaemia in patients with Type 1 diabetes with impaired awareness of hypoglycaemia. *Diabetes Care* 1994; 17: 697–703.
- 18 Fanelli CG, Epifano L, Rambotti AM, Pampanelli S, Di Vincenzo A, Modarelli F, Lepore M, Annibale B, Ciofetta M, Bottini P *et al.* Meticulous prevention of hypoglycaemia normalizes the glycaemic thresholds and magnitude of most neuroendocrine responses to, symptoms of, and cognitive function during hypoglycaemia in intensively treated patients with short-term IDDM. *Diabetes* 1993; **42**: 1683–1689.
- 19 Fanelli C, Pampanelli S, Epifano L, Rambotti AM, Di Vincenzo A, Modarelli F, Ciofetta M, Lepore M, Annibale B, Torlone E *et al.* Long-term recovery from unawareness, deficient counter-regulation and lack of cognitive dysfunction during hypoglycaemia following institution of rational intensive therapy in IDDM. *Diabetologia* 1994; **37**: 1265–1276.
- 20 Cryer P. Managing Diabetes: Lessons from Type 1 diabetes mellitus. *Diabetic Med* 1998; **15**(Suppl 4): S8–S12.
- 21 Lustman P, Clouse R, Freedland K. Management of major depression in adults with diabetes. Implications of recent clinical trials. *Semin Clin Neuropsychiatry* 1998; 3: 102–114.
- 22 Mazze R, Lucido D, Shamoon H. Psychological and social correlates of glycaemic control. *Diabetes Care* 1984; 7: 360–366.
- 23 Bellush L, Reid S. Metabolic and neurochemical profiles in insulin-treated diabetic rats. Am J Physiol 1994; 266: R87–94.
- 24 Lustman PJ, Skor DA, Carney RM, Santiago JV, Cryer PE. Stress and diabetic control. *Lancet* 1983; 1(8234): 588.
- 25 Lustman PJ. Depression in adults with diabetes. *Diabetes Spectrum* (Special Issue) 1994; 7: 161–189.
- 26 Carney R, Freedland K, Lustman P. Depression and coronary disease in diabetic patients: a 10-year follow-up. *Psychosom Med* 1994; 56: 149.
- 27 Kovacs M, Mukerji P, Drash A, Iyengar S. Biomedical and psychiatric risk factors for retinopathy among children with IDDM. *Diabetes Care* 1995; 18: 1592–1599.
- 28 Lustman PJ, Griffiths LS, Clouse RE, Freedland KE, Eisen SA, Rubin EH, Carney RM, McGill JB. Effects of nortriptyline on depression and glucose regulation in diabetes: results of a double-blind placebo-controlled trial. *Psychosom Med* 1997; **59**: 241–250.
- 29 Lustman PJ, Griffiths LS, Clouse RE, Freedland KE, Eisen SA, Rubin EH, Carney RM, McGill JB. Effects of alprazolam on glucose regulation in adult diabetic patients: results of a double-blind, placebo-controlled trial. *Diabetes Care* 1995; 18: 1133–1139.

- 30 Bush M. Compliance, education and diabetes control. *Mt Sinai J Med* 1987; **54**: 221–227.
- 31 Brown S. Studies of educational interventions and outcomes in diabetic adults: a meta-analysis revisited. *Patient Educ Couns* 1990; 16: 189–215.
- 32 Dunn S, Beeney L, Hoskins P, Turtle J. Knowledge and attitude change as predictors of metabolic improvement in diabetes education. *Soc Sci Med* 1990; **31**: 1135–1141.
- 33 Bloomgarden ZT, Karmally W, Metzger MJ, Brothers M, Nechemias C, Bookman J, Faierman D, Ginsberg-Fellner F, Rayfield E, Brown WV. Randomized, controlled trial of diabetic patient education: improved knowledge without improved metabolic status. *Diabetes Care* 1987; 10: 263–272.
- 34 Korhonen T, Huttunen JK, Aro A, Hentinen M, Ihalainen O, Majander H, Siitonen O, Uusitupa M, Pyorala K. A controlled trial on the effects of patient education in the treatment of insulin-dependent diabetes. *Diabetes Care* 1983; 6: 256–261.
- 35 Mazzuca SA, Moorman NH, Wheeler ML, Norton JA, Fineberg NS, Vinicor F, Cohen SJ, Clark CM Jr. The diabetes education study: a controlled trial of the effects of diabetes education. *Diabetes Care* 1986; **9**: 1-10.
- 36 Vinicor F, Cohen SJ, Mazzuca SA, Moorman N, Wheeler M, Kuebler T, Swanson S, Ours P, Fineberg SE, Gordon EE *et al.* DIABEDS: a randomized trial of the effects of physician and/or patient education on diabetes patient outcomes. *J Chronic Dis* 1987; **40**: 345–356.
- 37 Graber A, Christman B, Alogna M, Davidson J. Evaluation of diabetes patient-education programs. *Diabetes* 1977; **26**: 61–64.
- 38 Day J, Metcalfe J, Johnson P. Benefits provided by an integrated education and clinical diabetes centre: a follow-up study. *Diabetic Med* 1992; 9: 855–859.

- 39 Greenhalgh P. Shared care for diabetes. A systematic review. Occas Pap R Coll Gen Pract 1994; 67(i-viii): 1-35.
- 40 Anderson R. Is the problem of compliance all in our heads? *Diabetes Educ* 1985; **11**: 31–34.
- 41 Rappaport J. Terms of empowerment/exemplars of prevention: toward a theory for community psychology. Am J Community Psychol 1987; 15: 121–147.
- 42 Anderson R. Patient empowerment and the traditional medical model. A case of irreconcilable differences? *Diabetes Care* 1995; **18**: 412–415.
- 43 Greenfield S, Kaplan S, Ware JE Jr, Yano EM, Frank HJ. Patients' participation in medical care: effects on blood sugar control and quality of life in diabetes. *J Gen Intern Med* 1988; 3: 448–456.
- 44 Williams GC, Rodin GC, Ryan RM, Grolnick WS, Deci EL. Autonomous regulation: the motivational basis of adherence to medical regimes. *Health Psychol* 1998; 17: 269–276.
- 45 Street RL Jr, Piziak VK, Carpentier WS, Herzog J, Hejl J, Skinner G, McLellan L. Provider-patient communication and metabolic control. *Diabetes Care* 1993; **16**: 714– 721.
- 46 Damsbo P, Marbury TC, Hatorp V, Clauson P, Muller PG. Flexible prandial glucose regulation with repaglinide in patients with Type 2 diabetes. *Diabetes Res Clin Pract* 1999; **45**: 31–39.
- 47 Damsbo P, Clauson P, Marbury TC, Windfeld K. A double-blind randomized comparison of meal-related glycemic control by repaglinide and glibenclamide in well-controlled Type 2 diabetic patients. *Diabetes Care* 1999; **22**: 789–794.

(1) S20