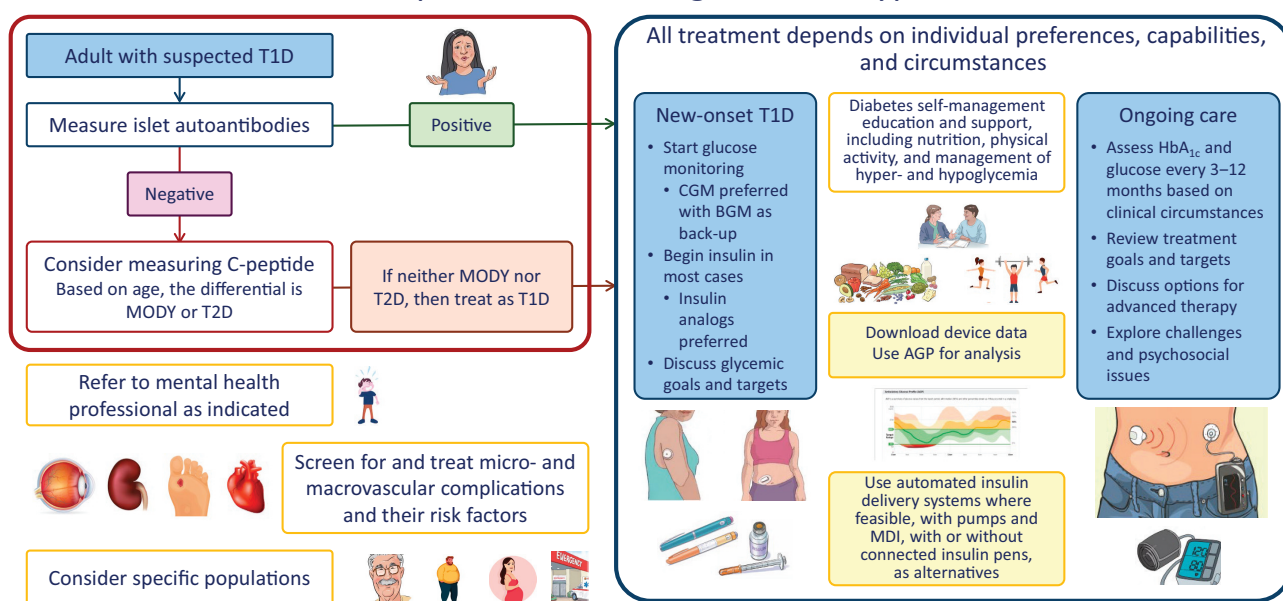


The Management of Type 1 Diabetes in Adults. The Updated 2026 Consensus Report by the American Diabetes Association (ADA) and the European Association for the Study of Diabetes (EASD)

Richard I.G. Holt, J. Hans DeVries, Amy Hess-Fischl, Irl B. Hirsch, M. Sue Kirkman, Tomasz Klupa, Barbara Ludwig, Kirsten Nørgaard, Jeremy Pettus, Eric Renard, Jay S. Skyler, Frank J. Snoek, Ruth S. Weinstock, and Anne L. Peters

Diabetes Care 2026;49(00):1–42 | <https://doi.org/10.2337/doi26-0122>

EASD–ADA Consensus Report on the Management of Type 1 Diabetes in Adults



AGP, ambulatory glucose profile; BGM, blood glucose monitoring; CGM, continuous glucose monitoring; MDI, multiple daily injections; MODY, maturity-onset diabetes of the young; T1D, type 1 diabetes; T2D, type 2 diabetes. Graphical elements were adapted from the 2021 EASD/ADA consensus report (<https://doi.org/10.1007/s00125-021-05568-3>). Original sources could not be definitively identified.



The Management of Type 1 Diabetes in Adults. The Updated 2026 Consensus Report by the American Diabetes Association (ADA) and the European Association for the Study of Diabetes (EASD)

<https://doi.org/10.2337/dci26-0122>

Richard I.G. Holt,^{1,2} J. Hans DeVries,^{3,4,5}
Amy Hess-Fischl,⁶ Irl B. Hirsch,⁷
M. Sue Kirkman,⁸ Tomasz Klupa,⁹
Barbara Ludwig,¹⁰ Kirsten Nørgaard,^{11,12}
Jeremy Pettus,¹³ Eric Renard,^{14,15}
Jay S. Skyler,^{16,17} Frank J. Snoek,¹⁸
Ruth S. Weinstock,¹⁹ and
Anne L. Peters^{20,21}

This 2026 consensus report from the European Association for the Study of Diabetes (EASD) and the American Diabetes Association (ADA) builds on the 2021 consensus report to provide guidance for managing type 1 diabetes in adults. Reflecting the rapid advances in the field, this update places particular emphasis on the integration of new technologies, while all sections have been revised to incorporate new evidence, advances in clinical practice, and novel therapies, including interventions to delay the onset of stage 3 type 1 diabetes. It also broadens its scope to screening for long-term diabetes complications, and the management of obesity and cardiovascular risk factors. Psychosocial care and diabetes self-management education and support (DSMES) remain key elements.

The report was developed using the Accurate Consensus Reporting Document (ACCORD) framework. The guidance aligns with the current ADA “Standards of Care in Diabetes” (Standards of Care) and relevant EASD and ADA documents and aims to support clinicians globally in delivering high-quality, personalized care for adults with type 1 diabetes, with recommendations that can be adapted across diverse health care systems and resource settings.

SECTION 1: INTRODUCTION AND RATIONALE FOR THE CONSENSUS REPORT

Type 1 diabetes is an autoimmune condition characterized by destruction of insulin-producing β -cells, resulting in profound insulin deficiency. Globally, type 1 diabetes affects an estimated 9 million people worldwide and accounts for 1–2% of all diabetes cases, while in Europe and North America it represents around 5% of diabetes diagnoses (1). Although incidence peaks in adolescence and early adulthood, it can develop at any age; in the U.S., the median age at onset is 24 years, with over half of new cases occurring in adults (2). Owing to long survival after diagnosis, the prevalence is higher among adults than children (3).

Since insulin’s discovery over a century ago, advances in insulin formulations, delivery systems, and glucose monitoring technologies have transformed care. However, many individuals remain above the glycemic targets for preventing complications, contributing to the persistent physical and emotional burden of living with type 1 diabetes.



View Online Article

¹Human Development and Health, Faculty of Medicine, University of Southampton, Southampton, U.K.

²National Institute for Health Research Southampton Biomedical Research Centre, University Hospital Southampton NHS Foundation Trust, Southampton, U.K.

³Amsterdam UMC, Internal Medicine, University of Amsterdam, Amsterdam, the Netherlands

⁴Diabeter Center Amsterdam, Amsterdam, the Netherlands

⁵Profil, Neuss, Germany

⁶Kovler Diabetes Center, University of Chicago Medicine, Chicago, IL

⁷UW Medicine Diabetes Institute, Seattle, WA

⁸University of North Carolina School of Medicine, Chapel Hill, NC

⁹Department of Metabolic Diseases, Center for Advanced Technologies in Diabetes, Jagiellonian University Medical College, Kraków, Poland

¹⁰University Hospital Carl Gustav Carus, Technische Universität Dresden, Dresden, Germany

¹¹Diabetes Technology Research, Steno Diabetes Center Copenhagen, Herlev, Denmark

¹²Department of Clinical Medicine, Faculty of Health and Medical Sciences, University of Copenhagen, Copenhagen, Denmark

¹³Division of Endocrinology, University of California, San Diego, La Jolla, CA

¹⁴Montpellier University Hospital, Montpellier, France

In response to these challenges, and the rapid pace of development, the European Association for the Study of Diabetes (EASD) and American Diabetes Association (ADA) published a consensus report in 2021 on the management of type 1 diabetes in adults focusing on glycemic management and acute complications (4,5).

This updated version addresses recent developments, including interventions to delay the onset of diabetes, emerging therapies, and new technologies. The 2021 consensus report included a detailed section on starting and adjusting subcutaneous insulin regimens (4,5). We elected not to include these details in this update, as the principles have not changed significantly and other resources are available (4–8). The current report broadens the scope to screening for long-term diabetes complications, and management of obesity and cardiovascular risk factors, while psychosocial care and diabetes self-management education and support (DSMES) remain integral components. In addition, we have added key points at the beginning of each section. Although developed by authors from Europe and the U.S., both EASD and ADA serve an international membership, and our aim is to support clinicians in optimizing care for people with type 1 diabetes, wherever they work.

This report follows the Accurate Consensus Reporting Document (ACCORD) principles (9). The 14-member writing group was appointed by the EASD Committee on Clinical Affairs on behalf of the EASD Board, and the ADA's scientific leadership on behalf of the ADA Board, based on their clinical and research expertise in type 1 diabetes. It included equal representation from both organizations across different clinical disciplines, with attention

to gender and geographical balance. All members of the original writing group participated in this update, which was co-chaired by Richard Holt (EASD) and Anne Peters (ADA). Two or four members of the writing group were assigned to be the primary authors of each section. These individuals had specific knowledge of the area and were tasked with reviewing and summarizing the available literature.

Before the evidence review and writing began, the writing group met twice online to agree on goals, content, methodology, and the writing teams to lead the report subsections. The section leads conducted literature searches to identify additional English-language studies published between January 2021 and July 2025. Evidence is drawn from observational and interventional studies, reflecting the limited availability of high-quality randomized controlled trials (RCTs) in many areas. Questions on clinical practice and interprofessional team collaboration for the management of type 1 diabetes provided the core of this report.

Recorded monthly virtual meetings (January 2025–August 2025) and ongoing email and web-based collaboration supported development. Meetings were held under the auspices of EASD, and a member of the ADA scientific team was present for all discussions. Writing group members collaboratively identified the topic areas and sections that needed to be updated during open discussions. Topic areas and questions were posed to the full group by the chairs and other writing group members during meetings. Discussions were carried out in detail to clarify meaning, resolve questions and bring forward new ideas. Qualitative meeting summaries were shared with writing group members and allowed reflection and

opportunities to air discussion points throughout the development of the report until consensus was reached. All writing group members collectively reviewed all sections to verify scientific rigor, language, and utility to the intended readership. Each section, in turn, was revised and approved by the entire writing group.

The guidance aligns, where possible, with the current ADA Standards of Care (6,10–20) and relevant EASD and ADA guidance documents (21–25). The draft report was presented at the EASD Annual Meeting in Vienna in 2025, after which public comments were invited from health care professionals and people with lived experience of type 1 diabetes. Several revisions were made in response to this input. The revised consensus report was peer reviewed externally and by EASD and ADA, and further changes were incorporated where appropriate.

The report represents the consensus of the writing group, acknowledging limitations in the evidence base.

SECTION 2: DIAGNOSIS OF TYPE 1 DIABETES

Key Points

- Adults with new-onset type 1 diabetes are frequently misdiagnosed as having type 2 diabetes, leading to delays in insulin treatment and an increased risk of diabetic
- Accurate classification of diabetes type has important clinical implications, influencing education, treatment strategy, and access to both technologies and adjunctive therapies.
- A diagnostic algorithm that integrates clinical features with islet autoantibody and C-peptide measurement can guide classification, particularly when the diagnosis is not self-evident.

¹⁵Institute of Functional Genomics, University of Montpellier, CNRS, Inserm, Montpellier, France

¹⁶Division of Endocrinology, Diabetes, and Metabolism, University of Miami Miller School of Medicine, Miami, FL

¹⁷Diabetes Research Institute, University of Miami Miller School of Medicine, Miami, FL

¹⁸Amsterdam UMC, Medical Psychology, Vrije Universiteit, Amsterdam, the Netherlands

¹⁹Department of Medicine, SUNY Upstate Medical University, Syracuse, NY

²⁰Keck School of Medicine, University of Southern California, Los Angeles, CA

²¹Leonard D. Schaeffer Institute for Public Policy & Government Service, University of Southern California, Los Angeles, CA

Corresponding author: Richard I.G. Holt, rich@ucl.ac.uk

Received 18 November 2025 and accepted 25 June 2026

This article contains supplementary material online at <https://doi.org/10.2337/figshare.33084311>.

This article is being simultaneously published in *Diabetologia* (<https://doi.org/10.1007/s00125-026-06833-z>) and *Diabetes Care* (<https://doi.org/10.2337/dci26-0122>) by the European Association for the Study of Diabetes (EASD) and the American Diabetes Association (ADA).

This article underwent external peer review. Following this, the report was assessed for EASD

by its Committee on Clinical Affairs and approved by its Executive Board and for ADA by its Professional Practice Committee.

© 2026 by the American Diabetes Association and the European Association for the Study of Diabetes. Readers may use this work for educational, noncommercial purposes if properly cited and unaltered. This publication and its contents may not be reproduced, distributed, or used for text or data mining, machine learning, or other similar technologies without prior written permission. More information is available at <https://diabetesjournals.org/journals/pages/license>.

- No single clinical feature or biomarker confirms type 1 diabetes in isolation, and diagnosis may require integration of clinical presentation with immunological and biochemical data and reassessment over time when uncertainty persists.
- Islet autoantibodies are highly sensitive markers, with approximately 90–95% of people presenting with clinical features of type 1 diabetes testing positive.
- C-peptide provides complementary information on endogenous insulin secretion and is particularly valuable when islet autoantibodies are absent or the clinical course is atypical.
- Three stages of type 1 diabetes have been defined, reflecting disease progression from islet autoimmunity with normoglycemia to dysglycemia and overt diabetes. These stages may guide screening efforts and the development of therapeutics for early intervention in the disease process.

Adults with new-onset type 1 diabetes may present after a short illness lasting 1–4 weeks or with a more slowly evolving disease course that can be mistaken for type 2 diabetes. Several other types of diabetes, for example, ketosis-prone type 2 diabetes (26), can be misdiagnosed as type 1 diabetes. In older adults, pancreatic cancer may present with diabetes and weight loss.

Most diagnostic data are derived from White European populations and may not be representative of other ethnic groups. Furthermore, most studies of the pathophysiology and natural history of type 1 diabetes come from children, and the pathogenesis of adult-onset type 1 diabetes may differ (27). The clinical presentation may vary, but the classical triad of thirst and polydipsia, polyuria, and weight loss are common symptoms of type 1 diabetes. Accurate classification of the type of diabetes has implications beyond insulin treatment. Education, insulin regimen, use of adjuvant therapies, access to newer technologies, need for psychosocial support, and concurrent disease screening may all depend on the diagnosis an individual receives. Furthermore, accurate diagnosis allows an assessment of the risk of diabetes in first-degree relatives and appropriate counseling. Although profound insulin deficiency is the hallmark of type 1 diabetes, some adults with type 1 diabetes retain some insulin secretion for

years postdiagnosis and may not require insulin treatment initially (28). This can create diagnostic ambiguity about the diabetes type and its optimal management. The use of a diagnostic algorithm for the investigation of adults with suspected type 1 diabetes can help mitigate this uncertainty (Fig. 1).

Differentiating Type 1 Diabetes From Type 2 Diabetes

Identifying type 1 diabetes in adults with newly diagnosed diabetes may be challenging, particularly when clinical features overlap with those of type 2 diabetes, such as older age with low or normal BMI or younger age with elevated BMI. Diabetic ketoacidosis (DKA), once considered pathognomonic of type 1 diabetes, may occur in ketosis-prone type 2 diabetes and in other types of diabetes, including that precipitated by use of SGLT2 inhibitors. Ketosis-prone type 2 diabetes is typically characterized by overweight or obesity, absence of islet autoantibodies, and measurable C-peptide shortly after resolution of the initial ketoacidosis (29).

Misclassification of type 1 diabetes in adults is common; over 40% of individuals diagnosed after age 30 years are initially treated as having type 2 diabetes (30–32). This misdiagnosis of type 2 diabetes is particularly likely in those with overweight or obesity and may cause confusion and distress. No single clinical feature confirms type 1 diabetes in isolation (33). The most discriminative feature is younger age at diagnosis (<35 years), with lower BMI (<25 kg/m²), unintentional weight loss, DKA, and glucose >20 mmol/L (>360 mg/dL) at presentation also being informative. Other features classically associated with type 1 diabetes, such as ketosis without acidosis, osmotic symptoms, family history, or a history of autoimmune diseases, are weak discriminators (32,33).

The strong relationship between age and type 2 diabetes incidence means that even “classical” features of type 1 diabetes may have a limited predictive value in older adults, in whom type 2 diabetes is far more common (34). Most older adults with low BMI will have type 2 diabetes (33,35,36), especially in ethnic groups with a high risk of type 2 diabetes (37). Rapid progression to insulin treatment (<3 years) is highly suggestive of type 1 diabetes at any age (30,32,38).

The term “latent autoimmune diabetes in adults” (LADA) is used to describe adults with diabetes who have islet autoantibodies but do not initially require insulin therapy. However, LADA is a controversial concept with no universally accepted diagnostic criteria and considerable heterogeneity in clinical presentation and progression (39,40). Proposed definitions vary by age, autoantibody titer, and duration of insulin independence, resulting in a diverse group of individuals. Earlier studies are likely to have been confounded by including mixed populations of individuals with either type 1 diabetes or type 2 diabetes (41). LADA represents a spectrum of type 1 diabetes, rather than a distinct subtype, characterized by slower β -cell loss than in children with type 1 diabetes and variable progression to insulin dependence. Use of the LADA label may be misleading and risks delaying insulin initiation. Management should focus on autoimmune status, insulin secretion, and clinical course.

Some individuals have features of both type 1 diabetes and type 2 diabetes, for example, islet autoimmunity as well as obesity or insulin resistance, as indicated by high insulin requirements. There are no diagnostic criteria enabling a later diagnosis of type 2 diabetes in people with established type 1 diabetes. Nevertheless, recognizing a second diabetes diagnosis may be important to facilitate access to noninsulin therapies (see section 8).

Investigation of Adults With Suspected Type 1 Diabetes

Although an initial diagnosis of type 1 diabetes is generally made on clinical grounds in adults presenting with hyperglycemia, the measurement of islet autoantibodies and C-peptide can help distinguish type 1 diabetes from other types of diabetes.

Islet Autoantibodies

Assessment of islet autoantibodies at diagnosis is recommended as the primary investigation in adults with suspected type 1 diabetes, where available. GAD antibody testing should be performed first; if negative, follow-up testing for IA-2 and/or ZNT8 antibodies should be performed, where available, as this can reduce the false-negative rate. Islet cell antibody (ICA) measurement is no longer recommended because of its imprecision and replacement by direct single antibody assays (42,43).

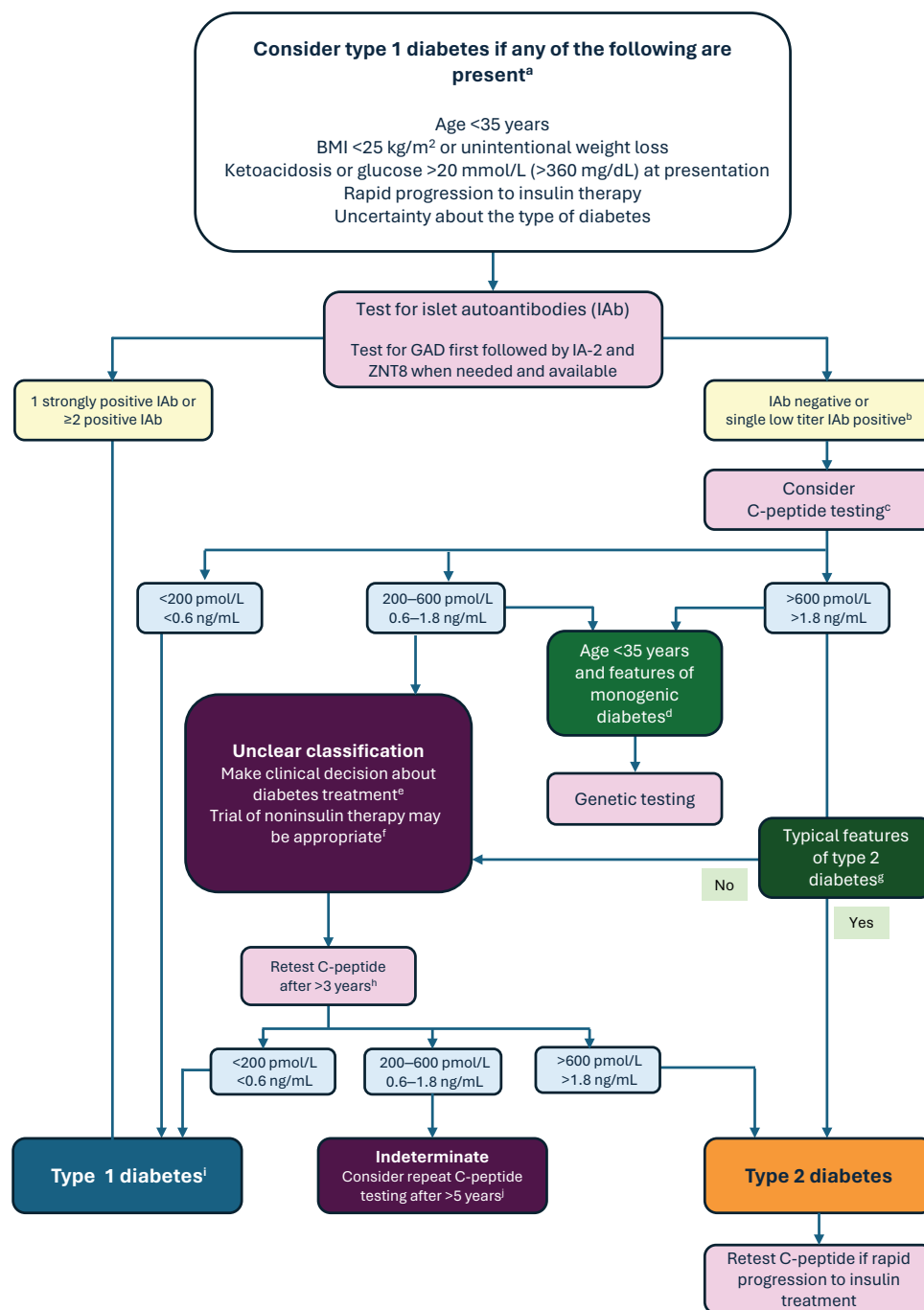


Figure 1—Flow chart for the investigation of suspected type 1 diabetes in adults. ^aNo single clinical feature confirms type 1 diabetes in isolation. Each of the listed factors should prompt consideration of type 1 diabetes but must be interpreted alongside other clinical and biochemical findings. No single clinical feature alone should automatically trigger islet autoantibody testing. Other weaker discriminators include the presence of autoimmunity, osmotic symptoms, ketosis without acidosis, and first-degree relatives with type 1 diabetes (32,33). ^bThe definition of low titer is dependent on the assay and laboratory used (43,45). Please refer to local advice. ^cA random C-peptide test should be performed with concurrent glucose measurement within 5 h of eating (28,32,53). If the result is ≥ 600 pmol/L (≥ 1.8 mg/mL), the circumstances of testing do not matter. If the result is < 600 pmol/L (< 1.8 mg/mL) and the concurrent glucose level is < 4 mmol/L (< 72 mg/dL) or the person may have been fasting, consider repeating the test (53). Tests showing very low C-peptide levels (< 80 pmol/L [< 0.24 mg/mL]) do not need to be repeated. In individuals treated with insulin, C-peptide must be measured prior to insulin discontinuation to exclude severe insulin deficiency. Do not test C-peptide within 2 weeks of a hyperglycemic emergency, within 12 h of a severe hypoglycemic episode, or in individuals with end-stage renal failure (due to altered clearance). ^dMonogenic diabetes should be considered when there are clinical features that are unusual in type 1 or type 2 diabetes (59). A high probability on a monogenic diabetes prediction model (www.diabetesgenes.org/exeter-diabetes-app/ModyCalculator; accessed 22 April 2026) and the presence of one or more of the following clinical features is suggestive of monogenic diabetes: $HbA_{1c} < 58$ mmol/mol ($< 7.5\%$) at diagnosis, one parent with diabetes, or features of a specific monogenic cause (e.g., renal cysts, partial lipodystrophy, maternally inherited deafness, severe insulin resistance in the absence of obesity). ^eType 2 diabetes should be strongly considered in older individuals. In some cases, investigation for pancreatic cancer or other types of diabetes may be appropriate. ^fIndividuals with suspected type 1 diabetes who are not treated with insulin will require careful monitoring and education so that insulin can be rapidly initiated in the event of glycemic deterioration. ^gFeatures of type 2 diabetes include increased BMI

The sensitivity of islet autoantibodies for diagnosis of type 1 diabetes is higher in younger adults and in those with rapid progression to insulin dependence. In White European and North American cohorts, reported sensitivity is approximately 70–80% for GAD alone, 85–90% for GAD plus IA-2 and 90–95% when all three antibodies are measured (44,45). Sensitivity is greatest when antibodies are measured close to diagnosis, as antibody positivity declines with increasing duration of diabetes (46).

The specificity of islet autoantibodies increases markedly with higher titers, multiple antibodies, and clinical features consistent with insulin deficiency. Positivity for two or more islet autoantibodies is strongly predictive of rapid progression and severe insulin deficiency; such individuals should be considered to have type 1 diabetes, even if they do not require insulin at diagnosis (44,45).

Low-titer GAD antibodies are detectable in approximately 1–2% of the general adult population and may lead to false-positive results. Isolated low-titer GAD antibody positivity requires cautious interpretation, and GAD should be measured only when type 1 diabetes is clinically suspected (45).

The absence of islet autoantibodies does not exclude type 1 diabetes. Approximately 5–10% of White European individuals with new-onset type 1 diabetes test negative for islet autoantibodies (32,33,47), and further diagnostic consideration is warranted. In those below 35 years of age, type 1 diabetes remains the most likely diagnosis, particularly in the absence of clinical features of type 2 diabetes or monogenic diabetes. In those aged over 35 years, the likelihood of type 2 diabetes increases with advancing age in the absence of islet autoantibodies. However, differentiating between type 1 diabetes and type 2 diabetes based solely on age and clinical features is not always accurate, particularly in people from African, Asian, Indigenous, Hispanic, and other diverse ancestral backgrounds.

A clinical decision regarding treatment is essential. Regardless of features suggestive of type 2 diabetes or the absence

of islet autoantibodies, individuals with suspected type 1 diabetes should be offered insulin treatment. However, in some individuals, when the clinical course aligns more closely with type 2 diabetes, a trial of noninsulin therapy may be appropriate. Those managed without insulin will require close monitoring and education to ensure prompt initiation of insulin if glucose levels deteriorate.

C-Peptide Measurement

Plasma C-peptide measurement assesses endogenous insulin secretion and should be considered when there is uncertainty about the type of diabetes in individuals without islet autoantibodies (48,49). It should not be measured within 2 weeks of a hyperglycemic emergency or within 12 h of a severe hypoglycemic episode. C-peptide may be falsely elevated in people with end-stage renal failure due to altered clearance. C-peptide should be measured alongside a concurrent glucose concentration within 5 h of eating. If the concurrent glucose level is <4 mmol/L (<72 mg/dL) and C-peptide level is <600 pmol/L (<1.8 mg/mL), the test should be repeated (32). At present, different commercially available C-peptide assays show significant variability, which limits the clinical utility of these defined cut-off values (50). In response, there have been calls to standardize C-peptide measurements to improve diagnostic accuracy and ensure patient safety, and a process for standardization has been established (51,52).

C-peptide levels fall progressively in people with type 1 diabetes, albeit more slowly in adults than in children, and are usually low or undetectable by 3 years after diagnosis. Consequently, the discriminative value of C-peptide for distinguishing type 1 diabetes increases with time since initial diagnosis. Beyond 3 years after diagnosis, a random C-peptide test is recommended if the diabetes type remains uncertain. Measurement of C-peptide should also be considered in those with clinically diagnosed type 1 diabetes of at least 3 years' duration, as routine C-peptide testing has led to reclassification in 11% of

such individuals (53). In individuals treated with insulin, this test should be performed prior to insulin tapering or discontinuation to exclude severe insulin deficiency. A persistent nonfasting C-peptide >600 pmol/L (>1.8 mg/mL) strongly suggests type 2 diabetes, and people with C-peptide in this range are often able to transition to noninsulin therapies (54–57). By contrast, except in rare circumstances, low or absent C-peptide confirms the diagnosis of type 1 diabetes. Low C-peptide concentrations may occur in some types of secondary diabetes and very long-standing type 2 diabetes, but such cases are rarely mistaken for type 1 diabetes; however, investigation of other types of diabetes may be considered when appropriate. Conditions in which C-peptide is low, such as postpancreatectomy or immune checkpoint inhibitor-induced diabetes, require similar treatment to type 1 diabetes.

A C-peptide measurement may have clinical utility earlier than 3 years after diagnosis. However, clinicians should be aware that normal C-peptide values may be seen in people with type 1 diabetes shortly after diagnosis. This scenario is more likely in adults than children because C-peptide values are typically higher at diagnosis in adults and fall more slowly (58). Consequently, while a low C-peptide concentration confirms insulin deficiency, a normal C-peptide concentration does not exclude type 1 diabetes.

Differentiating Type 1 Diabetes From Monogenic Diabetes

Depending on the population, monogenic diabetes accounts for up to 4% of those diagnosed with diabetes before 30 years of age. The likelihood of monogenic diabetes rises to 20% where islet autoantibodies are negative and C-peptide secretion is maintained (59). Monogenic diabetes is commonly mistaken for type 1 diabetes because of the young age at onset. Accurate diagnosis of monogenic diabetes enables tailored treatment, often allowing discontinuation of insulin, and carries important implications for screening for concurrent

(≥ 25 kg/m² [≥ 23 kg/m² in people of South Asian ethnicity]), absence of weight loss, absence of ketoacidosis and less marked hyperglycemia. ^hConsider earlier testing if clinically indicated. C-peptide <200 pmol/L (<0.6 ng/mL) will confirm type 1 diabetes (32). ⁱSome types of secondary diabetes may be associated with profound insulin deficiency, e.g., pancreatitis, and should be considered if clinical features indicate this. ^jC-peptide values 200–600 pmol/L (0.6–1.8 ng/mL) are usually consistent with type 1 diabetes but may occur in insulin-treated type 2 diabetes, particularly in people with normal or low BMI or after a long duration of illness (32). As C-peptide may decline more slowly in adults than children, C-peptide levels can remain indeterminate for longer periods of time and so retesting may be required.

Table 1—Stages of type 1 diabetes in people with two or more islet autoantibodies

Stage 1	Stage 2	Stage 3
• No IFG, no IGT, no increase in HbA_{1c}	• IFG: FPG 5.6–6.9 mmol/L (100–125 mg/dL) or • IGT: 2-h PG 7.8–11.0 mmol/L (140–199 mg/dL) or • HbA_{1c}: 39–47 mmol/mol (5.7–6.4%) or $\geq 10\%$ increase in HbA_{1c}^a	• Diabetes meeting standard diagnostic criteria (11)

^aA $\geq 10\%$ increase in HbA_{1c} refers to a sustained relative rise from a prior baseline measurement, confirmed on repeat testing over a period of usually 6–12 months, and interpreted in the context of other measures of dysglycemia. Small absolute changes within the range of biological or analytical variability should be interpreted with caution. FPG, fasting plasma glucose; IFG, impaired fasting glycemia; IGT, impaired glucose tolerance; PG, plasma glucose.

conditions and for genetic counseling in family members (60,61).

Genetic Testing

Molecular genetic testing for neonatal diabetes should be considered in all individuals diagnosed with diabetes under 6 months of age, regardless of current age, as over 80% will have monogenic neonatal diabetes, and the 30–50% with ATP-sensitive potassium (K_{ATP}) channel mutations can replace insulin with sulfonylureas (62,63).

Monogenic diabetes should be considered in those with one or more of the following features: diagnosis of diabetes before 35 years of age, HbA_{1c} <58 mmol/mol (<7.5%) at diagnosis, one parent with diabetes, and features of a specific monogenic cause (e.g., renal cysts, partial lipodystrophy, maternally inherited deafness, severe insulin resistance in the absence of obesity) (64). A monogenic diabetes prediction model is available at www.diabetesgenes.org/mody-probability-calculator (accessed 5 August 2025) to help identify individuals diagnosed between 6 months and 35 years who are at increased risk of monogenic diabetes (65). For people with diabetes who are of Hispanic, African, South Asian, East Asian, or other ethnicities in which young-onset type 2 diabetes is more prevalent, the probability of monogenic diabetes is lower. Low BMI and age at diagnosis are the most important discriminators for monogenic diabetes versus type 2 diabetes in these groups (66). Those at increased risk should undergo islet autoantibody and C-peptide testing. Molecular genetic testing, which is not universally available, should be considered only if islet autoantibodies are negative and nonfasting C-peptide is >200 pmol/L (>0.6 ng/mL) (67–69).

Stages of Type 1 Diabetes

Three stages of type 1 diabetes have been defined, characterized by the presence of

multiple islet autoantibodies (which may disappear in stage 3) and differentiated by the presence of normoglycemia (stage 1), dysglycemia (stage 2), or clinical diabetes (stage 3) (Table 1) (70).

Screening programs for type 1 diabetes are being implemented, although their risks and benefits remain unclear. Potential benefits of screening and early diagnosis during stage 1 or stage 2 type 1 diabetes include a lower risk of DKA at clinical diagnosis, avoidance of hospital admission, time to develop glycemic management skills through presymptomatic education and counseling, and time to psychologically adjust leading to reductions in both parental and child stress at diagnosis (71). Potential harms of screening for type 1 diabetes include psychological impacts (including anxiety, distress, and altered illness perception), overdiagnosis and medicalization, false-positive results and diagnostic uncertainty, potential inequities in access and outcomes, and financial and resource implications.

Several recent consensus reports have addressed the early identification of type 1 diabetes and have proposed different screening and monitoring strategies, ranging from screening of first-degree relatives of people with type 1 diabetes or high genetic risk scores to population screening (21,72,73). The first two approaches lead to higher rates of positive screening but miss the majority of potential cases, as only approximately 10% of individuals with type 1 diabetes have an affected first-degree relative (74–77). Conversely, screening for antibodies among the general population yields very low positivity rates (~0.3%). Furthermore, the optimal screening time remains to be determined and the cost-effectiveness and psychosocial impact of screening need further evaluation (73). Interventions to potentially delay or prevent progression to stage 3 are being

evaluated in stage 1 and stage 2 type 1 diabetes (see section 10). Individuals with stage 2 type 1 diabetes may benefit from teplizumab, an anti-CD3 antibody, which has been approved in North America, Europe, China, Asia, and parts of the Middle East and Latin America (78).

SECTION 3: OVERVIEW OF THE MANAGEMENT OF TYPE 1 DIABETES

Key Points

- The overarching aim of diabetes care is to support people with type 1 diabetes to optimize their health and quality of life.
- Care should be person centered and tailored to the needs, priorities, and circumstances of the individual with type 1 diabetes.
- Optimizing glycemic levels remains central to preventing acute and long-term complications of type 1 diabetes but must be balanced against the risk of hypoglycemia.
- Cardiovascular risk management is an integral part of type 1 diabetes care.
- Effective management of type 1 diabetes requires ongoing education, psychosocial support, and engagement with self-management.
- Care should be delivered by appropriately trained multidisciplinary teams using flexible models of care.

Type 1 diabetes is a complex and demanding condition that requires ongoing medical, educational, and psychosocial support from health care professionals with the appropriate skills, training, and resources. Care may differ at particular times of life, such as at the point of diagnosis, during concomitant illness or pregnancy, at the onset of complications, and later in life. The aim of diabetes care and management is to support people with type 1 diabetes to live a long and healthy

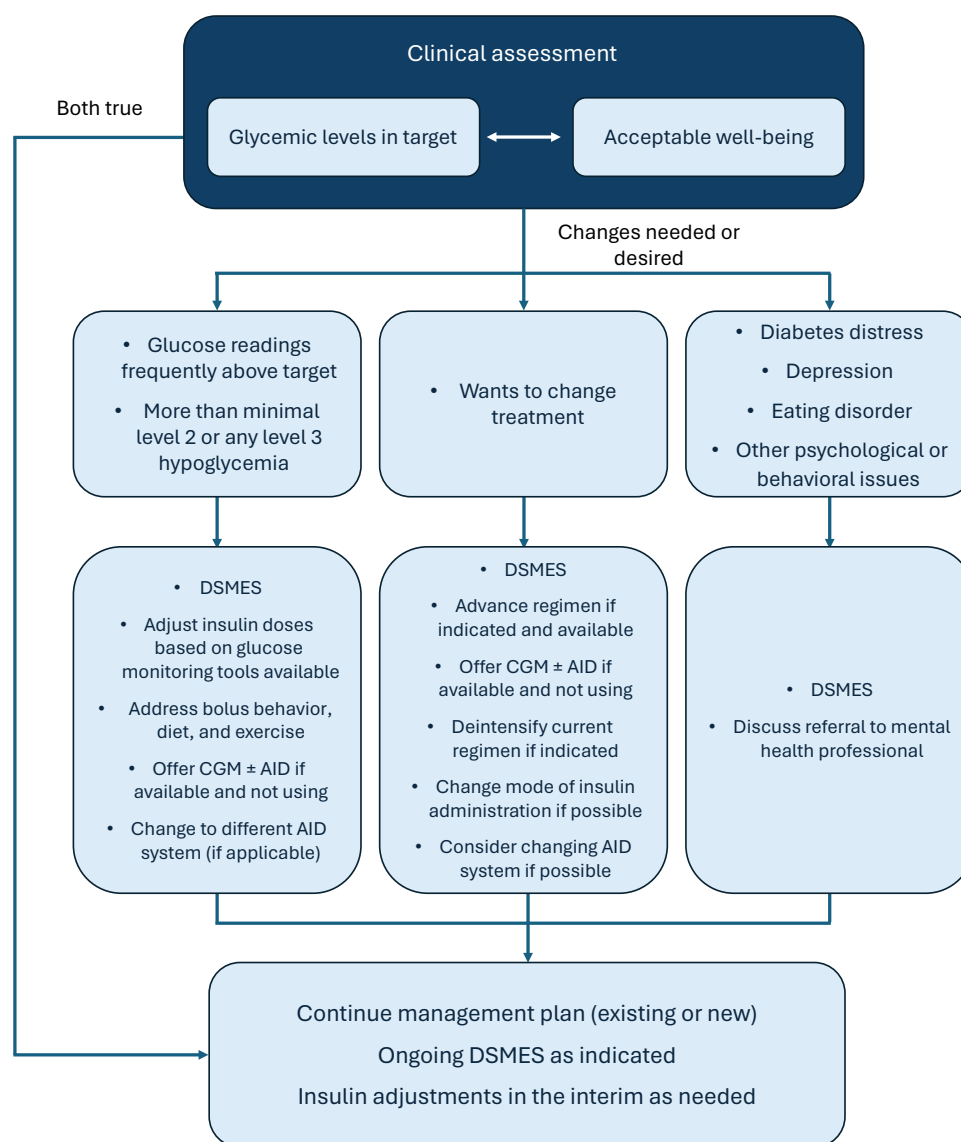


Figure 2—A framework for the management of individuals with type 1 diabetes, considering both glycemic levels and other psychosocial issues. AID, automated insulin delivery; CGM, continuous glucose monitoring.

life. The management strategies to achieve this aim broadly include:

- effectively delivering exogenous insulin to maintain glucose levels as close to an individual's target range as is safely possible to prevent the development and progression of diabetes complications while minimizing episodes of hypoglycemia;
- effectively managing cardiovascular risk factors;
- providing approaches, treatments, and devices that minimize the psychosocial burden of living with type 1 diabetes, while promoting engagement in self-care and psychological well-being.

The Diabetes Control and Complications Trial (DCCT) and its observational follow-up, the Epidemiology of Diabetes Interventions

and Complications (EDIC) study, demonstrated the importance of glycemic management in preventing diabetes complications (79,80). Intensive insulin therapy lowered HbA_{1c} by 22 mmol/mol (2%) over 7 years compared with standard care, resulting in a 60% reduction in retinopathy, 30% reduction in nephropathy, and 20% reduction in neuropathy. These benefits persisted over the long term despite the convergence of HbA_{1c} levels after the trial, with reductions in cardiovascular disease (CVD) emerging over time. These landmark findings underpin current glycemic targets worldwide. HbA_{1c} levels in adults with type 1 diabetes often reflect glycemic trajectories that were established earlier in life. Longitudinal cohort and registry data show that HbA_{1c}

typically rises during adolescence, peaks in late adolescence and young adulthood, and only partially improves thereafter (81,82). Consequently, many individuals enter adult services with elevated HbA_{1c} and require additional support from their health care teams.

Although optimizing glycemic levels is critical for the prevention of acute and long-term complications, interactions with the health care team should not solely focus on glycemic management but also include consideration of well-being and treatment satisfaction. Figure 2 provides a framework for such interactions, integrating glucose-focused interventions with assessments of diabetes distress, other psychosocial issues, and satisfaction with the current treatment regimen.

Management begins with a detailed evaluation at the initial consultation, followed by more targeted interval contacts with a focus on person-centered care (Table 2). A personalized approach to visit frequency is recommended, but visits should occur at least annually. For some people, an annual in-person consultation with the primary care team is sufficient. More frequent contact, however, may be needed for many individuals, for example, those who have been recently diagnosed, those who are not achieving their diabetes goals, those who require cardiovascular risk management, and those who would benefit from additional self-management education and psychosocial support. Additional visits may also be useful when the therapeutic regimen changes, for example, when the insulin regimen is modified or when a new device is started. Troubleshooting technology, preparation for and back-up plans in the event of system failure, and treatment of hypoglycemia should be reviewed at least annually.

Visits can be conducted in person or via telemedicine (83); however, despite the value of telemedicine, people should have the option of scheduling an in-person visit where possible (84). An annual in-person visit permits a physical examination to be undertaken, including of the feet and insulin injection or infusion sites. The use of telemedicine should be personalized and will vary depending on individual needs, computer literacy, and access to care (85). Additionally, asynchronous remote monitoring systems are being developed that can identify issues that may occur between regularly scheduled visits (86).

SECTION 4: DIABETES SELF-MANAGEMENT EDUCATION AND SUPPORT

Key Points

- Diabetes self-management education and support (DSMES) is a core component of care and is recommended for all people with type 1 diabetes.
- DSMES should be offered at key time points across the life course: at diagnosis, annually or when not meeting targets, if complications develop, and when transitions occur.
- Educational content and delivery should be personalized to each individual's needs, preferences, and circumstances.

- DSMES should address both practical self-management skills and psychosocial aspects of living with type 1 diabetes.
- Structured, evidence-based DSMES programs are effective and should be delivered as an ongoing, collaborative process integrated into routine clinical care and complemented by flexible, ongoing support.

DSMES is an essential component of type 1 diabetes care that allows other diabetes interventions to work optimally. The objective of DSMES is to provide those living with type 1 diabetes (and their caregivers, if applicable) with the knowledge, skills, and confidence to successfully self-manage their diabetes on a daily basis and, thereby, reduce the risks of acute and long-term diabetes complications while maintaining quality of life (87). DSMES aims to empower people with type 1 diabetes, with an emphasis on shared decision-making and active collaboration with the health care team. Where possible, DSMES programs should be evidence based and conform to local and national standards to ensure effectiveness.

DSMES Delivery Methods and Content

DSMES delivery methods and content should be guided by a comprehensive assessment, tailored to each individual's unique needs. This includes consideration of the time of diagnosis, prior education, psychosocial and cognitive status, literacy level, family history, and comorbidities, as well as ethnic, sociocultural, financial, geographical, and lifestyle factors (88). A structured, periodic assessment of educational needs and barriers should be an integral part of ongoing diabetes care (see Text box, "Needs assessment for DSMES"). As not all topic areas are useful or necessary for every person with diabetes every time DSMES is offered, the purpose of the assessment is to closely examine and discover current knowledge and self-management needs unbiased by the opinions of health care professionals.

DSMES can be delivered by various methods ranging from the provision of diabetes information and one-to-one advice to ongoing learning, which may be informal (e.g., through peer groups), and structured education that meets nationally agreed criteria, including evidence-based curricula, quality assurance

Needs assessment for DSMES

Key assessment features:

- Health history
- Cognition, functional health literacy and numeracy
- Health beliefs and attitudes
- Emotional health and support systems
- Religious and cultural influences
- Physical limitations
- Social determinants of health (e.g., financial status)
- Barriers

of teaching standards, and regular audit (89). These programs are guided by learning and behavior change theories and apply effective behavior change techniques (90). While DSMES has traditionally been delivered in person, the emergence of digital platforms and creative approaches, such as online education and the use of visual storytelling through comics, has expanded the ways in which individuals with type 1 diabetes can engage with and learn about their condition (91,92).

Structured programs for adults with type 1 diabetes are effective in improving both glycemic outcomes and psychosocial outcomes (89). Structured DSMES programs most often include multiple components and cover a broad range of topics, from pathophysiology to medical technology and healthy coping (Table 3).

DSMES should not be delivered as a one-off invention but should be offered on an ongoing basis and tailored to an individual's changing educational needs. People with type 1 diabetes may be diagnosed at a young age or during adulthood, and many live with type 1 diabetes throughout different life stages. There are four key moments when DSMES is especially important: 1) at the time of diagnosis, 2) when glycemic targets are not being met, 3) during periods of transition, and 4) on the development of diabetes-related complications (Fig. 3) (93). DSMES should be revisited when a child transitions to adult diabetes services, as there may be significant knowledge gaps in someone

Component of care	Details of evaluation
Medical and family history	
Diabetes history	Date of diagnosis Islet autoantibodies (date measured) C-peptide with concurrent glucose measurement (date measured) Episodes of DKA Episodes of level 3 hypoglycemia Hypoglycemia awareness
Family history	Type 1 diabetes or type 2 diabetes in first-degree relatives Other autoimmune disorders
Common comorbidities	Autoimmune disorders: thyroid, celiac, other ^a Hypertension Dyslipidemia Overweight and obesity Eating disorders Mild cognitive impairment and dementia Hearing loss Sleep disorder Dermopathy Fractures Joint and soft tissue disorders: cheiroarthropathy, trigger finger, capsulitis, carpal tunnel syndrome Dental and periodontal disease, which may contribute to altered taste
Other	Pregnancy and contraception history Vaccination history if applicable
Additional behavioral/lifestyle factors	Diet and nutrition: use of carbohydrate counting, weight history Physical activity Smoking, alcohol consumption, and substance use Sleep Occupation
Diabetes management	
Glycemic targets	HbA _{1c} CGM metrics including time in range, time above range, time below range, glycemic variability, median glucose, and adequate data capture
Current insulin regimen	MDI: pens, including connected pens; syringes; needles Insulin pump or AID system (type/model, settings, backup injection plan)
Glucose monitoring	CGM: type/model, data sharing (if applicable) Capillary glucose monitoring: type of meter/strips, frequency of use, mean (SD), range, and patterns
Other	Noninsulin diabetes medications Glucagon prescribed when indicated (remind person with diabetes to check expiration date) Ketone testing supplies prescribed when indicated
Psychosocial issues	Use of diabetes software or mobile applications Monitor psychological well-being including diabetes distress, depressive symptoms, and anxiety Consider fear of hypoglycemia and disordered eating Consider the impact of social determinants of health, including finance, and arrange social support as appropriate and available
DSMES	Assess cognitive status Assess individual needs and plan appropriate DSMES Dietary review Consider contraception and pregnancy planning in women of child-bearing age
Physical examination and complications screening	
General	Height Weight, BMI: every visit if not stable, otherwise annually Skin, including injection/infusion sites: every visit if skin complaints or erratic glucose readings are present, otherwise annually
Retinopathy	Initial examination within 5 years after diagnosis, followed by examinations every 1–4 years depending on prior findings, glycemic levels, blood pressure, and comorbidities Performed using validated approaches, most commonly retinal photography

Downloaded from diabetesjournals.org/care/article-pdf/doi/10.2337/dci26-0122/894606/dci260122.pdf by guest on 26 September 2026

Table 2—Continued

Component of care	Details of evaluation
Nephropathy	Initial examination within 5 years of diagnosis, followed by examination every 1–2 years Urinary albumin/creatinine ratio (uACR) measured in a random spot urine sample (annually) with confirmatory testing if elevated (repeat spot urine or 24 h collection) Creatinine and eGFR measured annually; may be more often if kidney disease present
Peripheral neuropathy	Initial examination within 5 years of diagnosis followed by annual examination Initial assessment with 10-g monofilament Further assessment may include either temperature or pinprick sensation (small-fiber function) and vibration sensation using a 128-Hz tuning fork (large-fiber function) If neuropathy is present, assess TSH and vitamin B ₁₂ . Consider alternative causes where clinically indicated Foot ulceration or deformity
Autonomic neuropathy	Screening is initially based on history and physical examination History-taking should assess for symptoms such as orthostatic hypotension, syncope, early satiety, erectile dysfunction, altered sweating patterns (particularly gustatory sweating), and dry, cracked skin on the extremities On examination, resting or fixed tachycardia (after ruling out hyperthyroidism), orthostatic hypotension, or evidence of peripheral dryness or cracking of the skin may be present More detailed tests include heart rate variability on electrocardiography, heart rate and blood pressure response to standing, and heart rate response to a Valsalva maneuver
Macrovascular disease	Annual assessment but more often if abnormalities or symptoms are present. Includes blood pressure and pulse, cardiovascular examination, and assessment of lower limb pulses Measurement of lipid profile: frequency dependent on the presence of previous lipid abnormalities or treatment
Laboratory testing	HbA _{1c} every 3–12 months ALT and AST: at least once and thereafter as indicated clinically Serum potassium: if taking ACE-I, ARB, or diuretics TSH, celiac screen: at least once and as indicated clinically ^a
Goal setting	Personalized, attainable, realistic: behavioral considerations (diet and nutrition, activity, smoking cessation) Glycemic: HbA _{1c} , time in range, hypoglycemia
Treatment plan	Formulate treatment plan with shared decision-making
Referrals	As needed: podiatry, cardiology, nephrology, ophthalmology, vascular surgery, gynecology, urology, orthopedic surgery, mental health professional, dentist, others

^aIndividuals with type 1 diabetes are at increased risk of other autoimmune diseases, including autoimmune thyroid disorders, pernicious anemia, celiac disease, collagen vascular diseases, and Addison disease (382). The optimal frequency of screening for these conditions in adults has not been established. Where indicated in the table, screening frequencies are in line with the ADA Standards of Care (6,10,12–17,19). ACE-I, ACE inhibitor; AID, automated insulin device; ALT, alanine aminotransferase; ARB, angiotensin II receptor blocker; AST, aspartate aminotransferase; CGM, continuous glucose monitoring; MDI, multiple daily injections; TSH, thyroid stimulating hormone.

diagnosed early in life, when education is directed at parents and caregivers. DSMES should also be revisited when there is a transfer of health care services or when a caregiver takes over any aspect of diabetes management to identify any new educational requirements.

An emerging area is the increase in number of individuals diagnosed with pre-stage 3 type 1 diabetes, as a result of type 1 diabetes screening efforts, and DSMES is needed for this group of people and their families (21). DSMES should be provided based on current needs and include the practical and psychosocial implications of their antibody status, the benefits of regular monitoring, and symptom awareness.

A wide range of smartphone and web-based applications (apps) are available to support people with type 1 diabetes in managing the complexities of daily self-care. While these tools are increasingly popular, the evidence supporting their safety, accuracy, and clinical effectiveness remains limited. Key concerns include insufficient validation of app algorithms, lack of standardized training for users, poor interoperability with other devices and systems, and inadequate data privacy protections (25).

Effective use of diabetes health apps requires ongoing dialogue between the health care team and the individual with diabetes (94). This includes assessing the person's understanding of the

app's content, their digital literacy, and the influence of social determinants of health on their ability to engage with the technology. It is equally important that health care professionals maintain up-to-date knowledge and competency regarding the apps their patients use. This enables informed evaluation of whether a given app is appropriate, safe, and beneficial for an individual's care needs.

SECTION 5: HEALTH-RELATED BEHAVIORS

Health-related behaviors, such as eating patterns, physical activity, sleep, and stress management, play a critical role in optimizing glycemic levels, reducing

Table 3—Key content areas of DSMES

Content area	Examples that focus on type 1 diabetes
Diabetes pathophysiology and treatment options	Immunology of β -cell destruction
Healthy eating	Basic and advanced carbohydrate counting vs. intuitive dosing Impact of meal composition (fat, protein, glycemic index, fiber, sugar, alcohol) on glucose levels Use of technology to enhance dosing recommendations
Physical activity	Impact on glucose and insulin dose recommendations
Medication usage	Types of insulin Methods of insulin delivery
Monitoring and using patient-generated health data	Technology and its ability to provide more frequent remote communication between people with type 1 diabetes and their health care professionals Review of CGM, pump, and connected insulin pen downloads and applications
Acute complications (hypoglycemia, hyperglycemia, and DKA), sick-day advice, and preparedness for emergencies	Signs and symptoms of DKA, including euglycemic DKA Problem solving insulin pump issues or malfunctions, including unexplained hyperglycemia, setting requirements, and infusion set and site issues, such as occlusions Back-up plan for pump/CGM failure (e.g., availability of insulin for injection and doses, meter and test strips) Glucagon use Ketone testing
Preventing, detecting, and treating chronic complications, including immunizations and preventive eye, foot, dental, and renal examinations, as indicated by an individual's duration of diabetes and health status	Understanding individual risk for complications in type 1 diabetes Preventing the development and progression of complications in the future
Healthy coping with psychosocial issues and concerns	Discussing strategies to help reduce diabetes distress and prevent "diabetes burnout"
Problem solving	Goal setting Developing personal strategies to promote health and behavior change Identifying problems and solutions Identifying and accessing resources Sick-day advice Management of pump failure or pump holidays Planning for procedures or surgery Preconception planning Pregnancy and diabetes

CGM, continuous glucose monitoring; DKA, diabetic ketoacidosis; MDI, multiple daily injection.

the risk of complications and enhancing overall quality of life (95). These behaviors often cluster, such that individuals may simultaneously engage in multiple healthy or unhealthy habits, which can amplify their impact on health outcomes. Religious and other cultural considerations, such as fasting, may also impact self-management of type 1 diabetes, and health care professionals should provide appropriate guidance and support to accommodate this (96). Health care professionals should adopt an integrated, person-centered approach to type 1 diabetes care that considers individual behavioral profiles and tailors interventions accordingly.

Nutrition Therapy

Key Points

- Nutrition therapy is personalized and aligned with an individual's preferences, culture, health literacy, and access to food.
- Carbohydrate intake is the primary dietary determinant of postprandial glucose and should guide prandial insulin dosing.
- Carbohydrate counting is effective, but alternative strategies (e.g., consistency or intuitive dosing) may be appropriate.
- Meal composition (fat, protein, fiber, alcohol) influences glycemic responses and insulin needs.

- Low-carbohydrate and very-low-carbohydrate eating patterns may be used safely with appropriate education and monitoring.

Nutrition, in particular carbohydrate intake, has a major effect on blood glucose levels, and people with type 1 diabetes need to understand the effect of food on their diabetes and plan meals accordingly (see Text box, "Goal of nutrition therapy for type 1 diabetes"). People with type 1 diabetes should be referred for personalized medical nutrition therapy provided by a registered dietitian who is knowledgeable and skilled in providing diabetes-specific

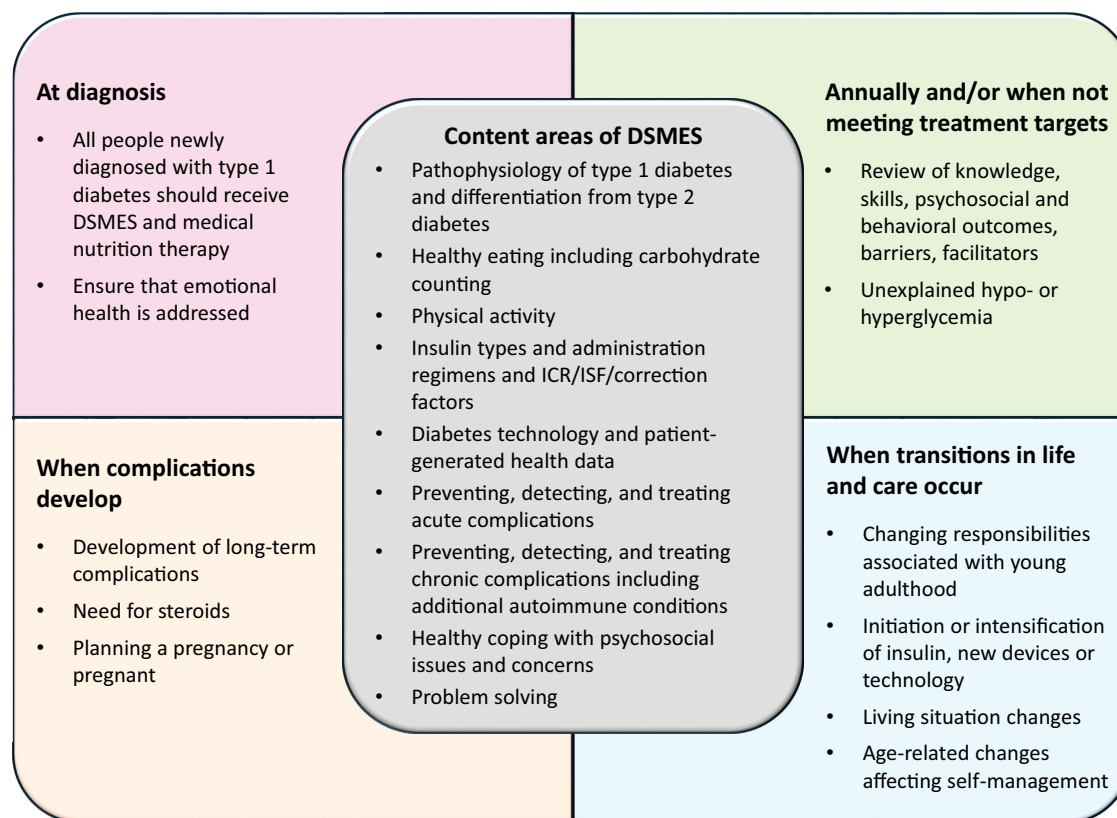


Figure 3—The four critical times when DSMES is particularly needed for people with diabetes (and their caregivers, when applicable). ICR, insulin/carbohydrate ratio; ISF, insulin sensitivity factor.

nutritional advice in conjunction with the diabetes technology being used. When integrated into an overall management program, medical nutrition therapy delivered by a registered dietitian is associated with a reduction in HbA_{1c} of 11–21 mmol/mol (1.0–1.9%) in people with suboptimally managed type 1 diabetes (12).

Nutritional recommendations for people with type 1 diabetes are based on personal preferences, socioeconomic status, cultural background, and comorbidities. Carbohydrate counting is the most common meal planning approach in type 1 diabetes. In conjunction with promoting healthy eating patterns, carbohydrate counting and the use of insulin/carbohydrate ratios can be useful methods for adjusting mealtime insulin dosing for optimal glycemic outcomes (97). When carbohydrate counting is not possible, teaching carbohydrate consistency is an alternative approach. While low-carbohydrate and very-low-carbohydrate eating patterns are increasingly popular, reduce HbA_{1c} levels, and increase time in range in the

Goal of nutrition therapy for type 1 diabetes (12)

- Promote healthy eating patterns, emphasizing a variety of nutrient-dense foods in appropriate amounts to improve overall health, HbA_{1c}, blood pressure, and cholesterol and aid maintenance of weight or achievement of weight goal
- Personalize nutrition advice based on individual and cultural preferences, health literacy, and access to healthy food choices
- Provide practical tools for day-to-day meal planning
- Focus on helping people dose their prandial insulin based on their ability to master carbohydrate-counting skills

short term, it is important to incorporate these alongside healthy eating guidelines.

Additional meal components, including high fat and/or high protein, may contribute to delayed hyperglycemia and the need for insulin dose adjustments. As the glycemic impact of a mixed meal varies substantially between individuals, and between people using multiple daily injections (MDI) and those using automated insulin delivery (AID) systems, postprandial glucose monitoring for up to 3 h or longer may be needed to determine initial and subsequent correction doses (12).

New interactive technologies using mobile phones to provide information, insulin bolus calculations based on insulin/carbohydrate ratios, and telemedicine communications with care providers may aid in reducing both weight gain and the time required for education (25). Artificial intelligence (AI) is increasingly being used to evaluate meal composition and is being integrated into various devices.

In the case of extreme low weight, unhealthy eating habits should be reviewed, including the possibility of insulin omission. Disordered eating is discussed further in section 6.

Physical Activity

Key Points

- Regular physical activity, including sports, should be encouraged for most people with type 1 diabetes, given its cardiovascular, metabolic, and psychosocial benefits.
- Different types, intensities, and durations of activity have variable effects on glucose levels and insulin requirements.
- Education should focus on balancing insulin dosing, carbohydrate intake, and glucose monitoring to reduce hypoglycemia and hyperglycemia.
- Diabetes-related technologies, such as AID systems, may support safe participation in physical activity when adjusted appropriately.
- Individual contraindications and diabetes complications should be considered when advising on activity type and intensity.

A combination of aerobic and resistance exercise on most days is associated with improved fitness, increased insulin sensitivity leading to reduced insulin requirements, improved cardiovascular health with better lipid profiles and endothelial function, and decreased mortality risk (98–100). Although regular exercise confers glycemic benefits, including improvement in HbA_{1c}, evidence for an independent beneficial effect on the preservation of β -cell function in early type 1 diabetes remains limited and inconclusive (101,102). Regular physical activity is also associated with reduced risks of microvascular complications and osteoporosis (100,103). Exercise helps maintain a healthy BMI and promotes sleep quality and psychological well-being.

Although most people with type 1 diabetes should be encouraged to undertake physical activity safely, it is important to consider cardiovascular and lower extremity comorbid conditions. Advice should be given regarding appropriate footwear and foot inspection for those with peripheral neuropathy to avoid the risk of ulceration. Walking does not increase the risk of ulceration in people with peripheral neuropathy, but weight-bearing exercise should be avoided in active foot disease (104). In individuals with proliferative or severe nonproliferative diabetic retinopathy, vigorous aerobic or resistance exercise involving straining

should be avoided because of the risk of vitreous hemorrhage or retinal detachment (105). These individuals should consult an ophthalmologist before beginning any high-intensity exercise program.

Education regarding physical activity should focus on the acute effects of exercise on glucose concentrations, which depend on several factors, including the individual's baseline fitness level; the type, intensity, and duration of activity; the amount of insulin in the circulation; the blood glucose concentration before exercise; and the composition of the last meal or snack. People with type 1 diabetes should learn about the effects of exercise on glucose levels and how to balance exogenous insulin delivery and carbohydrate intake for different types and intensities of

Considerations for glycemic management during exercise

- Type, duration, and intensity of exercise
- Time of day
- Fasting or nonfasting
- Acute and delayed effects of exercise on glucose concentrations
- Previous episodes of hypoglycemia
- Glucose monitoring before, during, and after physical activity
- Insulin dose adjustments up to 2–3 h before and after physical activity
- Insulin on board
- Management of hypoglycemia and hyperglycemia during and after physical activity
- Safety measures such as use of medical ID, availability of carbohydrates to consume before, during, and after exercise, exercising with others, glucagon, adequate hydration, backup supplies in case AID/CGM fails for longer duration or more remote exercise

Further details are provided in Supplementary Material 1.

exercise to minimize the risks of hypoglycemia and hyperglycemia (see Text box, “Considerations for glycemic management during exercise”). Post-exercise hypoglycemia can occur up to 24 h after the end of physical activity, and its occurrence is higher in untrained individuals and those engaging in physical activity irregularly (106). While general recommendations can be made (see Supplementary Material 1), the glycemic effects of physical activity and exercise can differ within and between individuals, highlighting the importance of experiential learning. Nevertheless, diabetes health care professionals can guide and support individuals with type 1 diabetes to reduce anxiety and build confidence around physical activity.

The introduction of continuous glucose monitoring (CGM) and AID systems has enabled more nuanced and proactive strategies to support safe participation in physical activity, including approaches to minimize hypoglycemia and hyperglycemia during and after exercise (107). Detailed guidance is beyond the scope of this report but the consensus statement for management of exercise in type 1 diabetes provides detailed suggestions regarding the use of CGM trend arrows and adjustment of insulin doses and carbohydrate intake (105). EASD and ISPAD have also developed a position statement on the use of AID systems during exercise (24).

Sleep

Key Points

- Many people with type 1 diabetes have disrupted sleep that may directly and indirectly adversely affect glucose levels.
- Health care professionals should ask about and support the management of diabetes-related factors that may impact sleep or sleep disorders.

Sleep patterns may be disrupted in people with type 1 diabetes as a result of both behavioral and physiological aspects of diabetes and its management, including, but not limited to, technology alerts and fear of overnight hypoglycemia (108–110). Many individuals with type 1 diabetes sleep less than current recommendations and have an increased risk of sleep-disordered breathing, which is associated with an increased risk of long-term diabetes complications (109–111).

There are mixed reports about the effect of sleep duration on glycemic management (109–111). Reviewing the frequency and appropriateness of technology alerts and alarms is an important first step in reducing unnecessary sleep disruption. It is also essential to address the fear of nocturnal hypoglycemia and to identify strategies to minimize its risk.

Alcohol and Recreational Drug Use

Key Points

- The use of alcohol and recreational drugs should be discussed to assess the potential risks and impact on glucose levels and diabetes self-management.
- Clinicians should be alert to features suggestive of substance use and ensure appropriate assessment and referral, given its association with increased risks of acute diabetes complications.

Many individuals with type 1 diabetes consume alcohol, although its effects on glycemic management are not always adequately considered. Increased alcohol consumption is associated with a higher risk of glycemic variability, typically with hyperglycemia initially and the potential for hypoglycemia hours later (112). Factors in alcoholic beverages that impact glucose levels include the amount of carbohydrate and glucose, the amount of alcohol by volume, and the volume of beverage consumed in conjunction with food. Excessive alcohol consumption impairs cognitive function and symptom awareness, leading to a diminished ability to self-manage diabetes (112). Alcohol consumption also appears to increase the risks of DKA and lactic acidosis, particularly in those with suboptimal glycemic management and in the context of reduced endogenous insulin. Alcohol inhibits hepatic gluconeogenesis, leading to an increased risk of hypoglycemia for up to 24 h after the last drink (112). Hypoglycemia is particularly hazardous because of the potential to confuse hypoglycemic symptoms with alcohol intoxication (112). Some of this glycemic variability may occur through the association with other risk-taking behaviors.

An association between recent recreational cannabis consumption and a more than twofold increased risk of DKA has been reported from countries where cannabis has been legalized, possibly related to the emergence of higher potency formulations of

cannabis and other synthetic cannabinoids (113,114). Heavy users can develop cannabis hyperemesis syndrome, which mimics gastroparesis (115). Use of cocaine and other stimulant-like drugs increases glucose production and inhibits glucose clearance, which increases DKA risk. A diagnosis of a substance use disorder is associated with an increase in all-cause mortality in populations with diabetes across a range of substances, including cocaine, opioids, and cannabis (114).

Health care professionals should ask about substance use and inform people with type 1 diabetes about the effects of drugs and alcohol on diabetes and related risks to avoid them seeking information elsewhere that may be incorrect and misleading (116).

Discussions regarding the impact of substance use on self-management should also be included in the education provided. Brief interventions to reduce substance use have been well validated in various populations and have the potential to improve diabetes medication taking and outcomes (117). However, targeted research specifically in people with type 1 diabetes remains limited, and more tailored interventions are needed. In the case of substance use disorder, referral to a specialized clinic is warranted.

Smoking

Key Points

- Smoking status should be assessed during routine consultations.
- Smoking cessation should be promoted and supported in all individuals with type 1 diabetes.

Smoking is a risk factor for both macrovascular and microvascular complications in individuals with type 1 diabetes (118–120). People who smoke tend to have suboptimal glycemic management, with reduced time in range, increased time in hyperglycemia, greater glucose variability, and a higher incidence of morning hypoglycemia (121). These findings emphasize the importance of smoking cessation as a crucial component of diabetes management.

Travel and Driving

Key Points

- Planning ahead is the key to safe travel for individuals with type 1 diabetes.
- Safe driving practices should be discussed regularly with people who drive.

Travel considerations for people with type 1 diabetes

- **Preparation:** Always carry diabetes-related and emergency supplies. Carry tools to keep insulin at the correct temperature
- **Insulin adjustment:** Plan insulin dosing to minimize glucose fluctuations, especially when crossing time zones. This should be supported by frequent glucose monitoring
- **Environmental factors:** Be mindful of changes in routine, climate, stress levels and physical activity that can affect glucose levels
- **Diet:** Research local foods to estimate carbohydrate content for better insulin management
- **Communication:** Carry note cards in the local language indicating that the individual has type 1 diabetes and may need urgent help in case of hypoglycemia. Smart phone apps can provide useful translation tools
- **Air travel:** Carry all supplies in carry-on/hand luggage. Consider bringing extra supplies. Declare medical equipment to security staff. Review pump and CGM manufacturer advice regarding flying. Check with national aviation security authorities regarding security screening rules

Individuals with type 1 diabetes should ensure they are well-prepared for travel and that all necessary diabetes-related and emergency supplies are easily accessible while traveling (see Text box, “Travel considerations for people with type 1 diabetes”) (122).

Unrecognized hypoglycemia and rapidly dropping glucose levels are the most relevant hazards for drivers with type 1 diabetes (123). These risks may be reduced by monitoring glucose prior to driving and at regular intervals during longer drives.

Local regulations and recommendations for driving with type 1 diabetes should be followed.

Employment

Key Points

- People with type 1 diabetes should have equal opportunities to pursue a wide range of careers.
- Employers should provide reasonable accommodation in accordance with local and national guidelines to enable people with type 1 diabetes to perform their roles effectively.
- Where the essential requirements of a role cannot be met without unacceptable risk to the person with diabetes or others around them, alternative roles may need to be considered.

People with type 1 diabetes are able to successfully undertake many different jobs; however, stigma and misconceptions about diabetes, particularly concerns about hypoglycemia and insulin access in challenging environments, can still limit employment opportunities (124). Chronic complications may also affect suitability for certain roles. While some occupations continue to restrict individuals with diabetes because of perceived safety risks, progress has been made in broadening employment opportunities, including to professions such as commercial airline pilot. It is not always necessary to disclose having type 1 diabetes, but some workplaces have specific protocols for safe employment for those with type 1 diabetes. People should coordinate with their human resources department if there are any issues. An updated ADA statement provides guidance for workers with diabetes, their health care teams, and employers navigating the legal and practical considerations for diabetes in the employment setting, including reasonable accommodations and the legally required interactive process between employers and workers (125).

To support safe and equitable employment, individuals with type 1 diabetes should be encouraged to pursue any role they are qualified for and can perform safely. Workplaces should provide reasonable accommodations, if possible, including access to insulin and glucose monitoring and time for self-management of blood glucose levels. Employment

laws differ between countries and sometimes between different regions within a country.

SECTION 6: PSYCHOSOCIAL CARE

Key Points

- Psychosocial factors should always be considered as part of routine adult diabetes care, which should be nonjudgmental, person centered, and responsive to changing life circumstances.
- Periodic screening of psychological well-being is recommended, at least annually, using validated questionnaires.
- Psychosocial problems are associated with increased risks of suboptimal self-management, higher HbA_{1c}, and diabetes-related complications.
- The diabetes team should preferably include a mental health professional who can advise the team and consult with people with diabetes in need of psychological support.

Type 1 diabetes is a psychologically challenging, chronic condition that impacts all domains of life, with treatment outcomes highly dependent on an individual's ongoing self-management. In this context, psychosocial factors play a significant role and encompass a person's cognitive functioning, beliefs, motivations, attitudes, coping strategies, feelings, and relationships with others (126). Poor mental health in type 1 diabetes is prevalent and associated with suboptimal glycemic levels and increased risks of complications (127). It is therefore imperative to adopt a biopsychosocial approach to type 1 diabetes, to achieve optimal outcomes in terms of both metabolic outcomes and psychosocial outcomes.

Social Determinants of Health

Life circumstances can significantly impact health in people with type 1 diabetes. In a review of social determinants of health and diabetes, the following domains were found to affect people with type 1 diabetes: 1) the neighborhood and physical environment (e.g., housing stability and interpersonal safety); 2) the built environment (e.g., walkability, access to green spaces, and access to transportation); 3) environmental exposures (e.g., pollution); 4) food access, availability, and affordability; and 5) health care access, affordability, and quality (128). Socioeconomic

challenges, particularly the inability to pay for food, insulin, other medications and supplies, and utilities, should be recognized and, where possible, addressed.

Psychological Problems

As in the general population, people living with type 1 diabetes experience chronic stress, such as stress related to work or financial hardship, as well as major stressful life events, such as unemployment or bereavement. Although not diabetes related, it is important to consider stress in type 1 diabetes because of its impact on quality of life and overall well-being and because it may also complicate diabetes self-management, thereby increasing the risk of not reaching glycemic targets (129).

Diabetes distress is common, affecting 20–40% of adults with type 1 diabetes, and can be experienced at any point from early adulthood to older age (130). “Critical” times are those following diagnosis, when complications develop, and when there is a loss of social support, for example, when an older adult loses their spouse or carer (131). Feeling powerless and overwhelmed by the daily self-care demands, fear of hypoglycemia, and worries about complications are among the most cited sources of distress by people with type 1 diabetes (132). Stigma, lack of social support, or feeling “policed” by family, friends, or coworkers can also evoke emotional distress in individuals with type 1 diabetes (133). Prolonged elevated diabetes distress can lead to “diabetes burnout” and is associated with an increased risk of depression, less engagement in self-care, and higher HbA_{1c} levels (134).

Depression and anxiety symptoms are twice as prevalent among adults with type 1 diabetes compared with adults without diabetes and negatively impact daily functioning and quality of life (135,136). Anxiety and depression often coexist and may partly overlap with symptoms of diabetes distress (137). Depression, at all levels of severity, is a risk factor for suboptimal self-care, hyperglycemia, long-term complications, and excess mortality (138). The association between generalized anxiety disorder and suboptimal blood glucose levels is less clear (139). Fear of hypoglycemia affects up to 10% of adults with type 1 diabetes, particularly among those experiencing

repeated episodes of level 3 hypoglycemia (140). Fear of hypoglycemia may translate into “phobic” avoidance behaviors aimed at keeping blood glucose at a “safe” level, resulting in persistent hyperglycemia. Although less common, lack of fear of hypoglycemia and/or fear of hyperglycemia may also be problematic, particularly in those with impaired hypoglycemia awareness and frequent severe hypoglycemic events (141,142).

Dysfunctional eating behaviors and eating disorders, including anorexia nervosa, bulimia nervosa, and binge eating, are overrepresented in people with type 1 diabetes, particularly in young women, but may also occur in men (143). However, people with type 1 diabetes are less likely to receive outpatient treatment for their eating disorders than their diabetes-free peers, despite their greater risk for major adverse health outcomes (144). Insulin omission may be used as a weight-loss strategy, particularly in girls and younger women, resulting in elevated HbA_{1c} levels (145).

Intellectual disabilities and neurodiversity warrant attention, as they may limit a person’s capacity to self-manage diabetes. Type 1 diabetes has been linked with attention deficit hyperactivity disorder (ADHD) and autism (146), with a Swedish nationwide cohort study reporting that comorbid neurodevelopmental disorders, primarily ADHD and intellectual disability, are associated with suboptimal glycemic levels and a higher risk of diabetes-related complications in childhood-onset type 1 diabetes (147).

As the life expectancy of people with type 1 diabetes increases, normal cognitive decline associated with aging may impact mental health and the capacity to self-manage diabetes and treatment outcomes. Importantly, in the 32-year follow-up of the DCCT/EDIC study, cognitive decline accelerated exponentially in participants with type 1 diabetes after 18 years of follow-up, and clinically significant cognitive impairment was found in up to 50% of older individuals with type 1 diabetes (148).

Psychosocial Screening and Monitoring

Monitoring of well-being and quality of life issues using person-reported outcome measures should always be considered in routine consultations and should not be restricted to those who report having

psychological difficulties (149,150). Assessment and periodic monitoring of an individual’s mental and social health status, at least on an annual basis, is recommended to identify individual needs and promote psychological well-being, engagement in self-management, and satisfaction with care (151,152). Given the prevalence of psychological issues in type 1 diabetes, screening can assist in case finding and timely referral for those in need of additional care, not least because psychological comorbidities can negatively affect diabetes outcomes and vice versa (153).

Implementing a standardized psychosocial evaluation is feasible but may require changes to current service provision, for example inviting people with diabetes to complete a set of questionnaires online or at the clinic prior to their visit (154). Practical, validated psychosocial screening tools are available for use in type 1 diabetes care, in multiple languages (see Supplementary Material 2).

There is no universally agreed core set of psychosocial measures for use across countries as part of clinical care for adults with type 1 diabetes, but there is consensus that well-being, diabetes distress (including worries around hypoglycemia, complications, and treatment burden), depressive symptoms, and social stress are among the most important domains to assess (150,155,156).

When disordered eating behaviors are reported, referral to a specialist mental health professional for a diagnostic interview for eating disorders should be considered (157). In case of suspected cognitive impairment, brief cognitive screening is recommended, followed by referral for more elaborate cognitive testing if needed.

Clinicians conducting psychosocial screening should possess a solid understanding of the psychosocial challenges commonly faced by individuals with type 1 diabetes, particularly those that may complicate diabetes management. They should also demonstrate strong communication skills, including active listening, a nonjudgmental approach to discussing sensitive issues, and the ability to constructively explore the option of referral to in-house or external specialized psychosocial services when appropriate.

Psychosocial Interventions

People living with type 1 diabetes may, at various stages, experience adjustment

problems and diabetes distress, which should be recognized as a normal response rather than a clinical condition (158). However, this does not imply that diabetes distress should be ignored. All members of the diabetes care team have a shared responsibility to provide supportive care as an integral component of diabetes management, helping people cope with the ongoing demands of the condition. Ideally, the diabetes care team should include a mental health professional (psychiatrist, clinical psychologist, and/or social worker) with diabetes expertise, who can offer guidance to the team and direct support to those requiring psychosocial care (159,160). Social needs may be addressed by social workers and community organizations. Social support from family and friends, peer-led initiatives, and digital self-help programs can play a valuable role in helping individuals cope effectively with the psychosocial demands of living with type 1 diabetes (161,162).

Psychological therapies, including time-limited in-person or online cognitive behavioral therapy, mindfulness, acceptance and commitment therapy and interpersonal therapies, are effective with regard to self-management and a range of psychological issues, including diabetes distress and depression (163–165). The effects of individual and group psychotherapy on glycemic levels are generally small but increase when DSMES is incorporated into the treatment (166). A small proportion of adults with type 1 diabetes have psychiatric conditions requiring treatment with psychotropic medications, which may impact glycemic management. In these cases close collaboration with a specialist mental health professional is warranted (167,168).

SECTION 7: INTERVENTIONS TO MANAGE GLUCOSE LEVELS

Key Points

- Lifelong insulin therapy is essential in type 1 diabetes and should be personalized to optimize glycemic levels.
- Continuous glucose monitoring (CGM) is the preferred method for monitoring glucose, as it protects against hypoglycemia and provides a complete view of glucose levels, both in real time and retrospectively, for optimizing treatment decisions.

- Regardless of CGM use, all individuals require capillary blood glucose testing supplies and a means of testing blood or urine ketones, with clear instructions on when and how to use these methods.
- HbA_{1c} remains the traditional measure of chronic glucose levels because of its prognostic value.
- Glycemic targets should be personalized.
- Analog insulins are preferred for subcutaneous insulin replacement.
- Automated insulin delivery (AID) systems are the optimal method of insulin delivery if used consistently.
- Diabetes self-management education and support strongly contributes to the effectiveness of multiple daily injection regimens.
- When using multiple daily injections, pen devices are preferred to vials and syringes.

Monitoring Glycemia

People with type 1 diabetes should have real-time access to information about their glucose levels and trends, including warning alarms and alerts, to help them maintain glucose within the target range and respond promptly to hypoglycemia and hyperglycemia. People with type 1 diabetes and their health care teams should review glucose data as often as needed to maintain blood glucose within recommended ranges.

Continuous Glucose Monitoring

CGM is the standard of care for glucose monitoring in adults with type 1 diabetes. CGM supports the optimization of glycemic levels, reduces rates of hypoglycemia, and improves quality of life (14). A meta-analysis of 24 studies comparing CGM with blood glucose monitoring (BGM) found that mean HbA_{1c} declined by 3 mmol/mol (0.24%), time in range (TIR) increased by 5.6%, and time below range (TBR) decreased by 2.4% (169). CGM has also proven beneficial in reducing the burden of hypoglycemia in older adults with type 1 diabetes (170), as well as severe hypoglycemia in those with impaired awareness of hypoglycemia (IAH) (171).

Historically, two types of CGM devices were available: real-time CGM (rt-CGM), which provides continuous sensor values based on current interstitial glucose levels and trends to a receiver, smartphone,

or smartwatch and/or insulin pump, and intermittently scanned CGM, in which the sensor glucose levels are determined by scanning a small reader or smartphone across the transmitter. The latter, however, has been superseded by rt-CGM. CGM devices allow both real-time interaction by the user and retrospective analysis by the user and the health care team. The devices offer predictive and threshold alerts that notify users when they are predicted to reach, or have reached, certain hyperglycemic or hypoglycemic thresholds, facilitating actions that prevent hypoglycemia or significant hyperglycemia, or immediate safety measures such as treatment of hypoglycemia. Additionally, users can respond to trends in glucose, enabling proactive treatment decisions (e.g., to avoid hypoglycemia) rather than reactive responses. Most CGM devices allow users to share their data in real time with a carer or family member, enabling them to receive alerts about dangerous glucose levels.

For retrospective analysis, data from devices can be uploaded to cloud-based programs, allowing users and health care professionals to easily view the data at or between clinic visits, enhancing therapeutic decision-making, understanding and engagement, and behavior change. However, this requires a smartphone and/or a computer with internet connectivity, which may not be readily available in under-resourced settings. CGM downloads should be performed and reviewed at each diabetes management visit. People using CGM should be involved in determining and understanding their personalized treatment goals (mean glucose, TIR, TBR, and other metrics) and, if they are not meeting these goals, establishing what changes could be implemented to help reach them. Examples of CGM target goals are shown in Fig. 4. Standardized glucose reports, such as the ambulatory glucose profile (AGP) and daily tracings, can facilitate these discussions (14).

The accuracy of CGM devices has improved over time and most sensors no longer require confirmatory capillary BGM. The U.S. Food and Drug Administration (FDA) requires CGM systems to meet standards for accuracy and precision prior to approval (172). However, in the EU the requirements for European Conformity (CE) marking are more vague, and proposals have been issued

for strengthening them (173). Access to BGM testing may be required, however, when there are concerns that CGM readings may be unreliable, such as when symptoms do not match concurrent CGM glucose values (e.g., falsely low readings due to sensor compression), when CGM devices are warming up or otherwise unavailable, and during correction of hypoglycemia due to the physiological lag between interstitial and blood glucose levels (174).

People with type 1 diabetes should be encouraged to review their own CGM reports regularly and follow their progress over time, contacting their health care team as needed for worsening or changing trends. Users can, for example, set alerts on some CGM devices to inform them of their weekly TIR and how it has changed compared with previous weeks. Such alerts can be helpful motivational tools and prompt the person with diabetes to contact their health care team when needed.

CGM is increasingly used in research and specialist settings in the presymptomatic stage of type 1 diabetes (stages 1 and stage 2) to detect early dysglycemia and refine risk stratification. CGM can identify subtle glucose abnormalities that may precede stage 3 type 1 diabetes and is being explored as a tool to monitor progression and as an outcome measure in prevention trials (21). Routine clinical use of CGM at these stages is not currently recommended, pending further research.

Capillary Blood Glucose Monitoring

Capillary BGM involves the use of a handheld meter that provides a measurement of plasma-calibrated capillary glucose. Frequent BGM monitoring should be advised for individuals not using CGM. In such cases, regular blood glucose measurements are an integral part of diabetes management, helping to guide insulin dosing, food intake, and the prevention of hypoglycemia. All those with type 1 diabetes should have the necessary equipment to undertake BGM, regardless of whether they are using CGM (14). Capillary BGM targets are shown in Table 4.

Ketone Measurement

Ketone bodies are produced when insulin concentrations are too low and/or counterregulatory hormone levels are

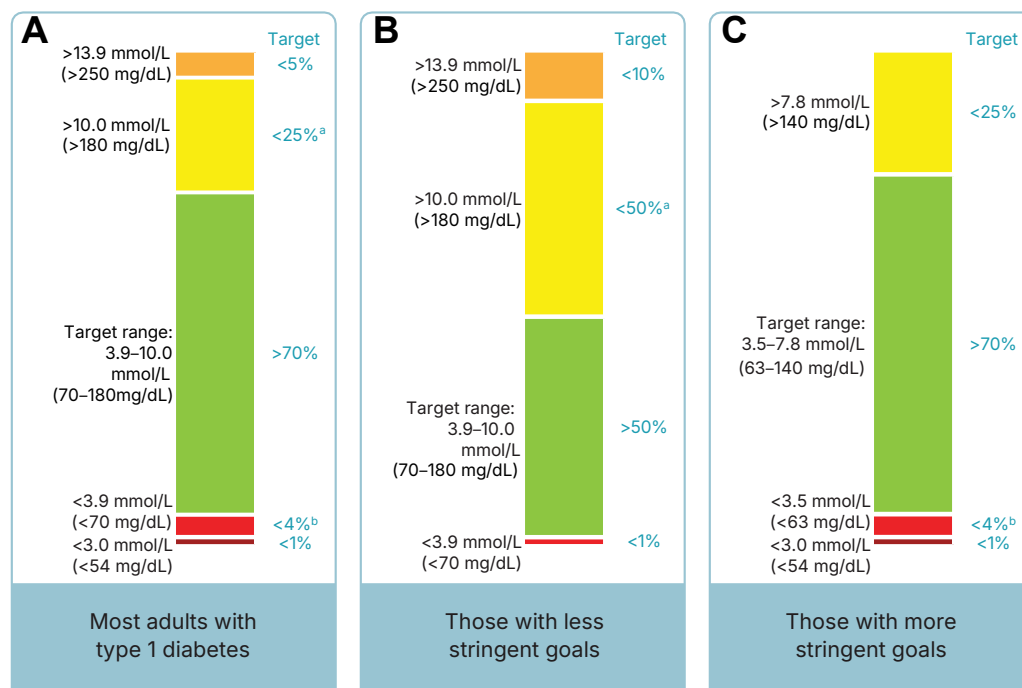


Figure 4—A framework for CGM targets for adults with type 1 diabetes. **A:** Most adults with type 1 diabetes. For most adults, the Glucose Management Indicator (GMI) should be <53 mmol/mol (<7.0%) and glycemic variability (%CV) should be <36%, although some studies suggest that lower %CV targets (<33%) provide additional protection against hypoglycemia. **B:** Adults for whom less stringent glycemic targets may be indicated or desired (frailty, limited life expectancy, problematic hypoglycemia). **C:** Adults who are pregnant or for whom more stringent glycemic targets are indicated or desired. In pregnancy, glycemic target ranges are lower than those recommended outside pregnancy. ^aIncludes percentage of values >13.9 mmol/L (>250 mg/dL); ^bincludes percentage of values <3.0 mmol/L (<54 mg/dL). Adapted and reprinted with permission from Battelino et al. (353).

too high to prevent lipolysis. If left untreated, ketosis can lead to progressive dehydration and DKA. Ketone measurement is important during periods of illness or hyperglycemia to facilitate the management of the hyperglycemia and prevent and/or treat DKA.

Adults with type 1 diabetes should have the ability to undertake ketone measurement at home and be instructed on when to test and how to respond to concerning

levels. Measurement can be performed using either blood or urine. Blood measurements are preferred as they are easy to perform, provide quantitative values, and reflect current ketone concentrations, whereas urine ketone measurements lag behind real-time ketone levels (175). However, blood ketone testing is more costly and may not be accessible to all. Methods for continuous ketone measurement are being actively developed (176,177).

HbA_{1c} Measurement

Monitoring of glucose levels over time has traditionally relied on HbA_{1c}, which has been used in most studies demonstrating that improved glycemic management reduces the risks of developing diabetes-related complications (79). There is a strong correlation ($r > 0.9$) between HbA_{1c} and mean blood glucose levels during the preceding 3 months if glucose levels have been stable. However, in some situations, for example when erythrocyte turnover is altered or in the presence of some hemoglobinopathies, HbA_{1c} does not reflect mean glucose, with mismatches of >2.2 mmol/L (>40 mg/dL) between mean glucose determined by CGM and that determined by HbA_{1c} occurring more than 5% of the time (13,175,178). Although there is variability in the relationship between mean glucose and HbA_{1c} between individuals, the relationship within an individual tends to be stable over time (179).

HbA_{1c} is an indicator of mean glucose but does not inform about glycemic variability and hypoglycemia and, therefore, should not be the only method used to evaluate glucose levels in type 1 diabetes

Table 4—Capillary blood glucose and HbA_{1c} targets for most adults with type 1 diabetes

Variable	Target value
Outside pregnancy	
HbA _{1c}	<53 mmol/mol (<7.0%)
Preprandial glucose	4.4–7.2 mmol/L (80–130 mg/dL)
1–2 h postprandial glucose ^a	<10.0 mmol/L (<180 mg/dL)
During pregnancy	
Fasting	<5.3 mmol/L (<95 mg/dL)
1 h after meals	<7.8 mmol/L (<140 mg/dL)
2 h after meals	<6.7 mmol/L (<120 mg/dL)

^aA postprandial glucose target of <7.8 mmol/L (<140 mg/dL) may be recommended if this can be achieved safely. Higher targets are recommended in those with limited life expectancy or where the harms of treatment are greater than the benefits. In some individuals at notably higher risk for level 3 hypoglycemia, it may be necessary to increase the glucose target range to decrease the TBR.

(179,180). With the widespread adoption of CGM, which provides Glucose Management Indicator (GMI) estimates alongside other detailed metrics, the need for frequent HbA_{1c} testing is being reviewed. There is emerging evidence linking CGM-derived metrics, especially TIR, to the risk of long-term diabetes complications after adjustment for HbA_{1c}, supporting a role for CGM in risk assessment beyond HbA_{1c} (181). It has been argued that HbA_{1c} testing could be performed less frequently than the current ADA recommendation of two to four times per year if CGM data are consistently shared with health care providers (13). An HbA_{1c} target of <53 mmol/mol (<7.0%) is appropriate for most adults with type 1 diabetes. Variability in HbA_{1c} has been associated with an increased risk of vascular complications and mortality, regardless of mean HbA_{1c} (182).

Insulin Therapy

Barring successful β -cell replacement, exogenous insulin is the primary treatment for type 1 diabetes. The ideal insulin replacement regimen maintains blood glucose within recommended ranges while allowing flexibility in terms of mealtimes, carbohydrate consumption, and activity levels. Insulin regimens all comprise two key elements: a basal insulin to restrain gluconeogenesis and ketogenesis in the fasting or postabsorptive state, and a mealtime or bolus insulin to cover food intake and hyperglycemia. These basal and mealtime components can come from either MDI regimens or insulin pump therapy. Regardless of the regimen chosen, glucose monitoring remains a critical and consistent component to guide insulin administration.

MDI Regimens

MDI regimens comprise a subcutaneous basal insulin analog, usually given as a single daily injection, together with separate injections of a mealtime rapid-acting or ultra-rapid-acting insulin analog. Ultra-rapid analogs have a slightly earlier time of onset and peak action than rapid-acting analogs. Compared with rapid-acting analogs, they reduce postprandial hyperglycemia but do not reduce HbA_{1c} or hypoglycemia further (183). Hence, both analog forms are usually referred to as rapid-acting analogs. Basal insulins have evolved from animal-derived and subsequently recombinant

human insulin preparations, including intermediate-acting formulations such as neutral protamine Hagedorn (NPH) insulin, to relatively “peakless” basal insulin analogs and newer second-generation analogs that have an extended duration of action, lower pharmacokinetic variability, and more consistent blood insulin levels (183). Compared with the original basal insulin analog glargine U-100, newer basal insulin analogs (insulin glargine U-300, insulin degludec) are associated with fewer episodes of hypoglycemia, a longer duration of action, and more flexible dosing schedules (6). As such, second-generation basal insulin analogs may be preferred in MDI regimens, particularly in those with problematic hypoglycemia.

Once-weekly basal insulin preparations have been developed, with two (insulin icodec and insulin efsitora) having completed phase 3 trials in adults with diabetes. In type 1 diabetes trials, both weekly insulins (compared with degludec as the basal insulin) demonstrated noninferiority for HbA_{1c} lowering but significantly higher rates of level 2 and level 3 hypoglycemia clustering at days in the middle of the dosing interval (184,185). Insulin icodec has been approved by the European Medicines Agency (EMA) for use in the EU, but the U.S. FDA has not approved either icodec or efsitora for people with type 1 diabetes, in part due to concerns about safety. Once-weekly basal insulins are currently not recommended for routine use in people with type 1 diabetes.

The 2021 consensus report (4,5) included a comprehensive section on the management of type 1 diabetes using MDI regimens. No substantive updates were identified and this guidance remains relevant for many individuals with type 1 diabetes, particularly in less well-resourced health care settings. It is therefore not repeated here, and readers seeking detailed information on MDI regimens are encouraged to consult the 2021 consensus report (4,5) and the current ADA Standards of Care (6), which provide practical algorithms for the initiation and titration of multiple daily insulin injections. Other sources of practical advice are available on the IDF (7) and National Institute for Health and Care Excellence (NICE) (8) websites.

Method of Injection. Insulin was traditionally administered using vials and insulin

syringes, but since the 1980s insulin pens that use cartridges or that are prefilled and disposable have become the most common mode of insulin administration in high-income countries. Insulin pens administer insulin to the nearest 1 unit (in some cases 0.5 units) increment and have the benefits of increased convenience, easier use, and, perhaps, increased accuracy compared with vial and syringe use (186). Some newer basal insulin analogs no longer come in vials. In lower resource settings or for people without health insurance, insulin pens are more costly and therefore vials and syringes may be used. Whether using pens or syringes, smaller gauge and shorter needles provide almost painless injections and reduce the risk of intramuscular (vs. subcutaneous) injection. Skin thickness is not significantly increased in individuals with overweight or obesity, contrary to common assumptions. Needles as short as 4 mm, injected at a 90° angle, enter the subcutaneous space with minimal risk of intramuscular injection in most adults (187).

Connected Pens and Bolus Calculators

Technology is emerging to help MDI users with the management of insulin doses and timing. Connected, or “smart,” insulin pens are either reusable injector pens with embedded electronics or standard pens with an add-on cap. Connected pens record each insulin dose and timestamp and transmit the data wirelessly to a smartphone or cloud-based platform. The systems may include dose reminders, bolus calculators and insulin-on-board tracking, and logging of insulin type and other variables. Some also integrate with CGM. There is a significant negative impact of missed basal and bolus insulin injections on glycemic levels, while greater interaction with smart pen data is associated with improved glucose self-management (188–190). An observational study showed an increase in TIR by up to 2 h per day by switching to a smart pen system (191). However, RCTs of smart pens have not shown significant improvements in glycemic levels. Several smart pens are approved by the U.S. FDA and/or carry the CE mark. A developing area is the incorporation of AI into connected pen platforms that analyze recent data and provide users with optimized insulin dosing advice (192).

Insulin Pumps

Insulin pumps deliver a continuous supply of rapid-acting analog insulin throughout a 24-h period. Most pumps are worn externally and are connected via tubing to a thin cannula inserted under the skin, although some systems deliver insulin via a pod connected to the cannula without external tubing. Insulin pumps were designed to mimic endogenous insulin kinetics more closely, delivering both basal and bolus components of insulin therapy. The basal component is a low-level infusion of insulin that may vary by time of day based on insulin needs. Bolus doses are larger amounts of insulin given over short periods of time before meals and/or to treat hyperglycemia. Users can manually adjust bolus doses based on carbohydrate intake, current blood glucose levels, and physical activity. Historically, insulin pumps did not integrate with CGM systems and thus users had to monitor glucose with BGM or CGM and perform most actions of the pump manually. Integration of pump functions with CGM data initially led to “threshold low glucose suspend” or “predictive low glucose suspend” features to avert hypoglycemia. Today’s pumps integrate more fully with CGM systems to form “hybrid” closed-loop systems (automated basal delivery while most bolus functions are controlled by the user), referred to herein as AID systems (14).

AID Systems

AID systems comprise three components: 1) an insulin pump to deliver insulin, 2) a continuous glucose monitor, and 3) an algorithm that interprets integrated CGM and pump data to determine whether changes in insulin delivery are needed (either more or less insulin).

Continuous glucose monitor data are continuously interpreted by the system to modify basal insulin delivery or trigger the delivery of a correction bolus (in some systems). These systems are generally capable of handling slow drifts in glucose values (up or down) but still require users to inform the pump when they are eating either by entering the carbohydrate count or announcing a meal (depending on the system). Physical exercise also needs to be announced, triggering the algorithm to set a higher glucose target to reduce the risk of hypoglycemia. AID systems have rapidly become the preferred mode of insulin

delivery for many people with type 1 diabetes, as studies with different AID systems have consistently shown improved glucose levels (lower HbA_{1c} and/or improved TIR) with reductions or no increases in CGM-detected hypoglycemia (14,193). Furthermore, quality of life is often improved (14).

This is a rapidly evolving area and several AID systems are available in different countries, with varying CGM system compatibility (Table 5). Each system has unique attributes and selection must therefore be personalized. Individuals should be encouraged to explore each option and consider key elements such as the capability of the CGM system and pump, whether the pump has external tubing, the size of the pump, the reservoir size (how much insulin the pump holds), whether the system requires batteries or is rechargeable, whether the system is waterproof, and the ease of self-management. Some systems require a compatible smartphone, which may present a barrier for individuals without access to suitable devices. For the latest information, readers may wish to refer to the “Diabetes Technology” section of the ADA Standards of Care, a frequently updated living document (14), and the PANTHER Program (<https://www.pantherprogram.org>) and Diabetotech (<https://www.diabetotech.com>) websites. It is important for both users and clinicians to be familiar with how the algorithms in different AID systems work, and what parameters can and cannot be adjusted to change glycemic outcomes.

Some people with type 1 diabetes employ user-driven open-source or do-it-yourself (DIY) AID systems. These systems use commercially available CGM devices and insulin pumps with open-source software algorithms that integrate CGM device and pump data to control basal insulin delivery and correction doses (194). These systems originated in response to initial delays in the commercial development of AID systems and the perceived limitations of the early commercial systems. In the U.S., the DIY Loop app has been approved by the FDA and is now part of a new AID system (twiist). Regulatory bodies do not allow health care professionals to prescribe algorithms other than Loop, but the hardware for the systems can be prescribed and managed. Health care

teams should respect an individual’s right to make informed choices about their care and continue to offer support to people using these systems, although support may be limited by a lack of knowledge of the control algorithm settings.

Fully closed-loop AID systems, which require almost no user input even for meals or exercise, are currently being evaluated in clinical trials in both North America and Europe (195). The expectation is that some of these will receive regulatory approval in the next few years. This could allow people with type 1 diabetes to spend more time in the target range, with a minimal risk of hypoglycemia and a reduction in the day-to-day burden of diabetes management.

Alternative Routes of Administration

Although subcutaneous insulin therapy has been the mainstay of treatment for over a century, this mode does not mimic physiological insulin secretion well. Healthy β -cells secrete insulin into the portal circulation at the onset of glucose intake, with approximately 70% of the insulin cleared by the liver and not entering the systemic circulation. Peak blood levels of endogenous insulin occur within 15–30 min after the start of a meal. Conversely, injected subcutaneous rapid-acting insulin enters the systemic circulation with some delay, shows a peak action at around 120 min, and is removed relatively slowly (196,197). Other modes of insulin administration have been designed to achieve pharmacokinetic and pharmacodynamic profiles that more closely resemble the action of endogenous insulin.

An inhaled human insulin is available that consists of powdered human insulin coated onto microparticles for oral inhalation; however, this formulation is only approved in a few countries (198). This delivery mechanism provides a very rapid onset and short duration of action. Compared with mealtime rapid-acting insulin analog injections, inhaled mealtime insulin results in lower post-meal glucose levels and equivalent TIR, hypoglycemia, and time above range (TAR). Cough occurs in about 25% of users, with slight but not significant reductions in forced expiratory volume (FEV₁) seen on spirometry (198,199). People using inhaled insulin should be monitored with periodic spirometry because of possible effects on lung function (200).

Implantable intraperitoneal programmable pumps are available in some areas

Table 5—Examples of widely used AID systems showing key characteristics and adjustable settings that can be used to optimize glycemic outcomes while in automated mode

Characteristic	MiniMed 780G	Omnipod 5	Tandem Control-IQ (t:slim X2 or Mobi)	myLife CamAPS FX (Ypsopump)	iLet ACE closed-loop insulin pump	Sequel Med Tech twiist	Diabeloop
Infusion mode	Tubing	Tubeless pod	Tubing	Tubing	Tubing	Tubing	Tubing
CGM compatibility	MiniMed Guardian 4 Instinct	Dexcom G6 Dexcom G7 Abbott Libre 2+	Dexcom G6 Dexcom G7 Abbott Libre 2+	Dexcom G6 Dexcom G7 Abbott Libre 3+	Dexcom G6 Dexcom G7 Abbott Libre 3+	Freestyle Libre 3 Eversense 365	Dexcom G6 Dexcom G7
Control algorithm	SmartGuard	SmartAdjust	Control-IQ	CamDia APS FX	iLet Dosing Decision	Tidepool Loop	DBLG1, DBLG2
Automation mode	Proportional Integral Derivative controller with insulin feedback + correction bolus	Model Predictive Control and Adaptive	Model Predictive Control + correction bolus	Model Predictive Control and Adaptive	Model Predictive Control and Adaptive	Model Predictive Control	Machine learning + correction bolus
Specific features	7-day infusion set; correction bolus every 5 min	Bolus calculator including basal active insulin	Extended bolus option; multiple profiles for basal, ISF, and I:C ratio; correction bolus every 60 min; simplified bolus	Broad glucose target (4.4–11.1 mmol/L [80–200 mg/dL]); Boost and Ease-Off modes	Carbohydrate counting not required	Emoji to estimate carbohydrate absorption time; maximum basal rate setting; Pre-Meal range for up to 1 h before meals, 3.7–7.2 mmol/L [67–130 mg/dL], typically set lower than the general correction range)	Dana pump or Kaleido pump; Zen mode for temporary target in between exercise mode and standard mode
Compatible smart phone required	No	No	Tandem T-slim: no; Tandem Mobi: yes	Yes	No	Yes	No
Settings that can be adjusted to optimize glycemic outcomes in automated mode							
Insulin/carbohydrate ratio (I:C ratio)	Yes	Yes	Yes	Yes	No	Yes	Yes
Insulin sensitivity factor (ISF)	No	Yes	Yes	Yes	No	Yes	Yes
Active insulin time	Yes	Yes	No	No	No	No	No
Glucose target	Yes	Yes	No	Yes	Yes	Yes	Yes
Basal rate	No	No	Yes	No	No	No	No
Exercise mode ^a	Yes	Yes	Yes	Yes	No	Yes	Yes
Sleep mode ^a	No	No	Yes	No	No	No	No

The table shows systems available at the time of publication but is not an exhaustive list of all currently available devices. ^aTarget can be changed with sleep mode (but not independently). Targets always changed with exercise mode. I:C, insulin/carbohydrate; ISF, insulin sensitivity factor.

in Europe. The pump is embedded in a subcutaneous pocket of the abdominal wall and infuses short-acting (regular) insulin with a stabilizing agent through the peritoneal route (201). Despite not being connected to a CGM device, these systems improve glycemic outcomes compared with subcutaneous insulin pumps, including sustained reductions in HbA_{1c} levels and fewer severe hypoglycemic events (202). Integration with CGM devices may allow fully automated insulin delivery via the peritoneal route (203).

Adverse Effects of Insulin

The main adverse effect associated with insulin therapy is hypoglycemia, which is discussed in section 9. Insulin may cause body weight gain, which can lead to some people with type 1 diabetes reducing their insulin doses. Sections 13 and 6 on management of overweight and obesity and psychosocial care, respectively, provide more information.

Skin reactions to subcutaneous insulin therapy include local inflammation (often due to either the pH of the insulin solution or additives to the insulin), insulin-induced lipohypertrophy, insulin-induced lipoatrophy, and allergy. Lipohypertrophy is common, typically resulting from repeated use of the same injection or pump site. It contributes to increased insulin requirements and causes glycemic variability, leading to both hyperglycemia and hypoglycemia (204,205). The condition is frequently underdiagnosed, but this can be improved through structured DSMES. In the single-arm Lipohypertrophy Monitoring Study (LIMO), 171 individuals with insulin-treated diabetes were followed prospectively; approximately two-thirds had lipohypertrophy at baseline, with many injecting into affected areas, rotating injection sites incorrectly, and reusing needles. An intervention combining education on injection technique and site rotation with the provision of single-use 4-mm pen needles reduced severe and unexplained hypoglycemia and glycemic variability, although HbA_{1c} did not improve significantly (206). People with type 1 diabetes should receive instruction about proper injection and pump site insertion techniques, including regular site rotation, at insulin initiation and periodically thereafter. Clinicians should inspect and palpate injection and infusion sites at least annually. While ultrasound can detect earlier and smaller lesions than

physical examination (204), its added clinical value remains uncertain.

Insulin-induced lipoatrophy has become rare due to improvements in the purity of human and analog insulin. True insulin allergy is rare and typically presents as recurrent local or systemic immediate- or delayed-type hypersensitivity reactions elicited by each injection, with symptoms centered on injection sites. In some cases, switching to a different insulin preparation or changing from MDI to insulin pump therapy may alleviate allergic responses (207).

SECTION 8: ADJUNCTIVE THERAPIES FOR GLYCEMIC MANAGEMENT

Key Points

- Evidence supporting the use of adjunctive therapies in type 1 diabetes to improve glycemic management is limited and none are routinely recommended.
- Off-label use of adjunctive therapies is increasing but requires a full understanding of the risks and benefits in an individual with type 1 diabetes.

While insulin therapy is essential for people with type 1 diabetes, maintaining blood glucose within recommended ranges with insulin alone is difficult because of the risks of hypoglycemia, the slow action of “rapid-acting” insulins, insulin resistance, difficulty with carbohydrate counting, and many additional factors. Furthermore, insulin therapy is often associated with undesirable weight gain, which may worsen insulin resistance. Insulin therapy does not address other pathophysiological abnormalities that are present in people with type 1 diabetes, such as α -cell dysfunction, and does not fully protect individuals from an increased risk of CVD. Thus, the potential role of “adjunctive therapies” is to improve glycemic management without increasing hypoglycemia and body weight.

The evidence supporting the use of adjunctive therapies is increasing but is too limited to allow for general recommendations about their use. However, these therapies can be considered in individual cases (Table 6); however, before these drugs are prescribed, insulin therapy should be optimized. The use of incretin-based drugs and sodium–glucose cotransporter 2 (SGLT2) inhibitors for obesity and cardiovascular risk

management is described in sections 12 and 13.

Metformin

The effect of metformin on glucose levels has been evaluated in numerous small trials in people with type 1 diabetes. A recently updated meta-analysis reported modest benefits for HbA_{1c} and reductions in total daily insulin dose, BMI, and total cholesterol (208). However, the largest and longest study to date, involving 428 people with type 1 diabetes treated for 3 years, found no difference in the primary end point of changes in mean carotid intima–media thickness (a marker of CVD risk). Effects on HbA_{1c} were minimal and not sustained, weight reduction was modest (~1 kg) and there was no significant change in total daily insulin dose (209). Thus, metformin therapy is generally not recommended for people with type 1 diabetes, although it may be considered on an individual basis.

Pramlintide

Pramlintide, an amylin analog, is the only approved adjunctive therapy to insulin in the U.S.; however, it is not approved in Europe. Injection prior to meals suppresses glucagon secretion, delays gastric emptying, and promotes satiety (210–213). Clinical trials have shown a reduction in HbA_{1c} (3–4 mmol/mol [0.3–0.4%]) and modest (~1 kg) weight loss with pramlintide (214–217). Because of its adverse effects and the need for additional injections, clinical uptake of pramlintide has been limited. It was withdrawn from the market in the mid-2010s and is no longer clinically available. However, coformulations of amylin with insulin are currently in development, and the use of pramlintide in pumps or AID systems is also being explored.

GLP-1 Receptor Agonists and Dual GLP-1 Receptor/GIP Receptor Agonists

Glucagon-like peptide 1 (GLP-1) receptor agonists have been explored in people with type 1 diabetes for two indications. The first aimed to ameliorate β -cell decline at the time of diagnosis, and there are ongoing trials of this approach. In one study of 308 people with recently

Table 6—Adjunctive therapies for glycemic management in type 1 diabetes

Variable	Metformin	Pramlintide	GLP-1 ± GIP receptor agonists	SGLT2 or SGLT1/2 inhibitors
HbA _{1c} reduction	1–7 mmol/mol (0.1–0.7%)	3–4 mmol/mol (0.3–0.4%)	2–4 mmol/mol (0.2–0.4%)	2–4 mmol/mol (0.2–0.4%)
Fasting glucose	Minimal effect	No effect	Minimal effect	Modest decrease (0.8 mmol/L [15 mg/dL])
Postprandial glucose	Minimal effect	Significant decrease	Modest decrease	Modest decrease
TIR	No data	No data	No data	Increased (~12% at higher doses)
Insulin dose	Mean reduction of $-0.44 \text{ U kg}^{-1} \text{ day}^{-1}$	Mealtime reductions	Predominantly mealtime reductions	Mealtime and basal reductions (~10% total reduction)
Body weight or BMI	BMI reduction (-0.71 kg/m^2)	Modest weight decrease (~1 kg)	Significant weight decrease (~5–15 kg)	Moderate weight decrease (2–3 kg)
Systolic blood pressure	No change	No change	4 mmHg decrease, increase in heart rate	3–4 mmHg decrease
Hypoglycemia	Low risk	Potential increase in level 3 hypoglycemia if prandial insulin doses are not decreased	Increase in hypoglycemia	Low risk
Side effects	GI side effects	GI side effects	GI side effects; increase in ketosis	Genital mycotic infections; 2.5- to 6.0-fold increased risk of DKA; dehydration
Approval status for type 1 diabetes in EU/U.S.	Not approved	U.S. approved	Not approved	Not approved
Cardiovascular benefits in other populations, including type 2 diabetes	Yes	No	Yes	Yes

EU, European Union; GI, gastrointestinal; GIP, glucose-dependent insulinotropic polypeptide; GLP-1, glucagon-like peptide 1.

diagnosed type 1 diabetes, liraglutide, when used in combination with anti-IL-21 antibody, preserved β -cell function (218). The second indication is as an adjunctive therapy in established type 1 diabetes by blunting glucagon secretion, slowing gastric emptying, and promoting satiety and thereby weight loss (219). The largest clinical trials in people with type 1 diabetes have been conducted with liraglutide and showed, at daily doses of 1.8 mg, decreases in HbA_{1c} (2–4 mmol/mol [0.2–0.4%]), weight (~5 kg), and insulin doses (220,221). However, increased rates of hypoglycemia and ketosis were also found. A recent study found that semaglutide is safe in people using AID systems, lowering HbA_{1c} by 3 mmol/mol (0.3%) and body weight by 8.8 kg (222), while two studies reported that the dual GLP-1/glucose-dependent insulinotropic polypeptide (GIP) receptor agonist tirzepatide significantly reduced HbA_{1c} and

body weight in adults with type 1 diabetes (223,224). Given the effect on insulin requirements, users should be provided with careful advice about insulin dose adjustment when GLP-1 receptor agonists or dual GLP-1/GIP receptor agonists are started or stopped.

SGLT Inhibitors

In several phase 3 programs in people with type 1 diabetes, the use of SGLT1 or SGLT1/2 inhibitors has been shown to reduce HbA_{1c}, improve TIR, reduce body weight, and improve blood pressure (225). However, a meta-analysis of 20 studies including 7,183 participants reported that individuals treated with SGLT1 or dual SGLT1/2 inhibitors had a 2.57- to 5.96-fold higher incidence of DKA than those receiving placebo (226). This increased rate of DKA led to the U.S. FDA's rejection of market authorization for their use in type 1 diabetes. Although the EMA previously approved

low-dose dapagliflozin (5 mg) and sotagliflozin (200 mg) as an adjunctive therapy for type 1 diabetes in those with a BMI $\geq 27 \text{ kg/m}^2$, their market authorization was subsequently withdrawn on the request of the market authorization holders (227). While no risk mitigation strategies have been proven to lower the risk of DKA, a consensus statement on SGLT2 inhibitors and DKA suggested careful patient selection, appropriate insulin dose adjustment to avoid insulinopenia, starting with a low dose of SGLT2 inhibitors, and regular ketone measurements with prompt action to address elevated values as sensible precautions aimed at preventing DKA (228). The development and clinical use of continuous ketone monitors may help inform future approaches to reduce the risk of DKA associated with SGLT inhibitors in people with type 1 diabetes (177); however, this potential role is speculative and would require formal evaluation.

SECTION 9: ACUTE METABOLIC EMERGENCIES

Key Points

- Clinicians should proactively identify impaired awareness of hypoglycemia (IAH) using validated questionnaires or simple questions such as “Can you always feel when your blood glucose is low?” and “At what blood glucose level do you feel symptoms?”
- IAH is a significant risk factor for severe hypoglycemia, but the risk can be reduced through use of continuous glucose monitoring, titrated insulin regimens that mimic physiological endogenous insulin secretion, and structured or personalized education.
- Risk factors for diabetic ketoacidosis (DKA) include younger age, lower socioeconomic status, infection, intercurrent illness, psychological stress, omission of or under-dosing of insulin, and adjunctive use of sodium–glucose cotransporter 2 (SGLT2) inhibitors.
- DKA prevention strategies include diabetes self-management education and support and awareness of sick-day advice, including intensified glucose and ketone monitoring.

Hypoglycemia

Hypoglycemia, the most important limiting factor in the glycemic management of type 1 diabetes (13), is classified into three levels:

- Level 1: plasma or interstitial glucose concentrations <3.9 mmol/L (<70 mg/dL) and ≥ 3.0 mmol/L (≥ 54 mg/dL). This is an alert threshold at which people on insulin should take action.
- Level 2: plasma or interstitial glucose concentrations <3.0 mmol/L (<54 mg/dL). Level 2 hypoglycemia is considered clinically important hypoglycemia.
- Level 3: hypoglycemia characterized by an altered mental state and/or physical status needing the intervention of a third party for recovery. Level 3 hypoglycemia is also called severe hypoglycemia.

Epidemiology of and Risk Factors for Hypoglycemia in Type 1 Diabetes

Rates of hypoglycemia are generally determined from self-report, CGM data (for level 1 or level 2 hypoglycemia), and data from hospitals and emergency

departments. CGM data can include false-positive hypoglycemia, including compression lows, and should be validated. Hospital and emergency department data primarily describe, but likely underestimate, severe hypoglycemia. Hypoglycemia, including severe episodes, remains troublingly common in people with type 1 diabetes. In a study of approximately 67,000 adults with type 1 diabetes, adjusted rates of severe hypoglycemic emergencies increased from 25.7 to 32.9/1,000 person-years between 2011 and 2019 and then decreased to 25.6/1,000 person-years in 2020. Although the trends were not statistically significant, these analyses suggest that rates of severe hypoglycemia are not declining despite advances in therapies (229). In a cross-sectional survey of adults with type 1 diabetes (92% using CGM and almost half using AID systems), 20% reported having had at least one episode of severe hypoglycemia in the past year, with 12% reporting at least two episodes. Although a reported history of severe hypoglycemia was lower with use of technology (e.g., 34% of nonusers of CGM vs. 18% of CGM users), 16.6% of those using AID systems still reported an episode within the past year. However, the timing of initiation of AID use was not ascertained (230).

Risks for hypoglycemia, particularly level 3 hypoglycemia, include a longer duration of diabetes, older age, a history of recent level 2 or level 3 hypoglycemia, high glycemic variability, alcohol ingestion, exercise, a lower education level, a lower household income, chronic kidney disease, and IAH (13). IAH is the reduced ability to recognize low blood glucose levels, typically due to loss of counter-regulatory hormone responses and their associated symptoms (231). Confusion or loss of consciousness may be the first sign of hypoglycemia, pre-empting appropriate corrective behaviors. The prevalence of IAH is estimated to be 25–30% in people with type 1 diabetes but is likely to be underestimated based on CGM data (232). IAH markedly increases the risk of severe hypoglycemia. The subsequent fear of hypoglycemia may lead people with diabetes to intentionally omit insulin doses or relax their glycemic targets in an effort to prevent the recurrence of hypoglycemia. IAH is associated with, but not fully explained by, autonomic neuropathy (233). It can be

induced or worsened by recurrent hypoglycemia and alcohol use (13). Although most hypoglycemic episodes in people with type 1 diabetes result from insulin mismatch, clinicians should be aware that autoimmune thyroid and adrenal disorders and undiagnosed celiac disease, which are more common in this population, are potential causes of unexplained hypoglycemia.

Diabetes health care professionals should proactively ask people with type 1 diabetes whether, and at what glucose level, they feel hypoglycemia in order to identify IAH and adjust individual glucose targets to reduce the risk of severe hypoglycemia. Validated questionnaires, such as the Clarke (234), Pedersen-Bjergaard (235), and Gold (236) questionnaires and the Hypoglycemia Awareness Questionnaire (HypoA-Q) (237) can also identify IAH. However, simple questions such as “Can you always feel when your blood glucose is low?” and “At what blood glucose level do you feel symptoms?” can be used to identify IAH and the risk of severe hypoglycemia (13).

Significant discrepancies have been shown between continuous glucose monitor-detected and patient-reported hypoglycemia (174); hence, both modes of capturing hypoglycemia occurrence should be considered, especially because only patient-reported hypoglycemia has been shown to have a significant impact on daily functioning (238).

Consequences and Prevention of Hypoglycemia

Hypoglycemia has significant impacts on quality of life and psychological well-being. Severe hypoglycemia may lead to injury to the person with diabetes or to others, for example when driving. Hypoglycemia in type 1 diabetes is estimated to account for more than 8% of deaths among those aged <56 years (239). The long-term association of severe hypoglycemia with subsequent cognitive function has been examined in the DCCT-EDIC cohort. In these analyses, severe hypoglycemia in younger or middle-aged adults did not appear to affect neurocognitive function after 18 years of follow-up (240). However, after 32 years of follow-up (median age 59 years), more episodes of severe hypoglycemia were independently associated with greater decrements in psychomotor and mental efficiency (148).

Several strategies can be used to reduce the risk of clinically significant or severe hypoglycemia. Structured education

programs, such as Dose Adjustment For Normal Eating (DAFNE) and Blood Glucose Awareness Training (BGAT), which provide informed support for appropriate glucose monitoring and active insulin dose self-adjustment, have been shown to reduce severe hypoglycemia rates in those at high risk (241). As these specific programs are not widely available, periodic DSMES that addresses modifiable risks of hypoglycemia and optimization of insulin therapy is indicated in all adults with type 1 diabetes, with enhanced education and support provided for those with problematic episodes.

The use of insulin analogs and carefully titrated basal-bolus regimens is standard of care in people with type 1 diabetes, due to their closer mimicry of physiological endogenous insulin secretion and reduced risks of hypoglycemia. Compared with BGM, CGM has been shown to reduce TBR and episodes of severe hypoglycemia in those at high risk of hypoglycemia (170,171,241). AID systems have potential to detect impending or early hypoglycemia and to protect against more severe events. AID systems are associated with further reductions in CGM-detected hypoglycemia compared with use of CGM and insulin pumps in open-loop mode, including in populations with high rates of severe hypoglycemia (242). Despite evidence that AID technology reduces the risk of nonsevere hypoglycemia, randomized and real-world studies have not generally demonstrated reductions in episodes of severe hypoglycemia, with occasional exceptions (243). This may be due to statistical power issues, participant selection, or lack of sensitivity for the detection of severe hypoglycemia.

Early clinical physiology studies suggested that strict avoidance of hypoglycemia helped restore hypoglycemia awareness (233). However, several clinical trials have not shown reductions in rates of IAH with CGM use despite a reduced incidence of hypoglycemia (241). In a survey of adults with type 1 diabetes, rates of IAH (measured using the Gold questionnaire [236]) were about 30% in all subgroups (CGM users vs. BGM users, AID vs. open-loop pump or MDI use), despite differences in rates of reported severe hypoglycemia episodes (230).

In some situations, it may be necessary to increase the glucose target range if hypoglycemia cannot be rectified

through other means. Intractable severe hypoglycemia is an indication for islet or pancreas transplantation (see section 10).

Treatment of Hypoglycemia

The standard recommendation for correcting level 1 or level 2 hypoglycemia is the oral intake of approximately 15 g of glucose or equivalent simple carbohydrate when the capillary or interstitial blood glucose concentration is <3.9 mmol/L (<70 mg/dL). This should be repeated every 15 min until any symptoms have resolved and the blood glucose level is >3.9 mmol/L (>70 mg/dL). As there may be a 5- to 15-min lag between changes in capillary blood glucose and interstitial glucose, CGM may show delayed detection of the restoration of normoglycemia. For those with a tendency toward over-treatment, the use of BGM is recommended to determine when hypoglycemia has resolved (13). Less glucose (5–10 g) may be needed to correct hypoglycemia while using AID, because the delivery system should have already reduced or stopped basal insulin delivery, and because overcorrection may trigger additional insulin delivery (244,245).

When there is a reduced level of consciousness, oral glucose intake is contraindicated because of the risk of aspiration. Instead, caregivers or bystanders should administer glucagon via subcutaneous or intramuscular injection or via nasal delivery. Intravenous glucose injection is a possible alternative for health care professionals in cases of level 3 hypoglycemia. For many years, glucagon was dispensed as a powder and separate diluent that required mixing prior to administration. Newer forms of glucagon that are stable in solution and ready to inject, or that can be given intranasally, are easier for bystanders to use and may lead to more rapid correction of severe hypoglycemia (13). However, these preparations may be significantly more expensive and may not be available in all locations.

After the acute symptoms have resolved, a further 20 g of carbohydrate as part of a snack or meal may need to be ingested if there is still significant insulin on board or if exercise was the cause of the hypoglycemic episode. If possible, the cause of the episode should be ascertained to determine preventive actions for future episodes.

Diabetic Ketoacidosis

DKA is a life-threatening but preventable acute complication of type 1 diabetes, characterized by hyperglycemia, metabolic acidosis, and ketosis. Occasionally, DKA can be present when glucose levels are normal or only minimally elevated (<11.1 mmol/L [<200 mg/dL]); this “euglycemic” DKA is more common with pregnancy, insulin pump use, or adjunctive use of SGLT2 inhibitors. The underlying cause of DKA is insulin deficiency, either absolute (new diagnosis of type 1 diabetes or omission of insulin in those with diagnosed disease) or relative (increased counterregulatory hormones due to infection or other stressors without an adequate increase in insulin doses) (22). Omission, or inadequate doses, of insulin may be iatrogenic, such as when clinicians mischaracterize adult type 1 diabetes as the more prevalent type 2 diabetes.

The prevalence of DKA and risk factors for its development have been less well studied in adults with type 1 diabetes than in children. In the U.S., national surveillance of emergency department visits and hospital admissions suggests a rate of 28 cases per 1,000 adults with diabetes per year, with a worrying increase in emergency department visits and admissions for DKA since 2009 (246). In Denmark, the incidence of DKA in adults with type 1 diabetes doubled between 1996 and 2008, with a subsequent minor decrease from 2008 to 2020 (247). In a European (predominantly Germany and Austria) registry, adults with type 1 diabetes had DKA at a rate of 2.5 per 100 patient-years (248).

DKA is more common in younger adults (aged 18–44 years) with type 1 diabetes than in older individuals, and in people who are uninsured or of lower socioeconomic status. DKA can be the presenting manifestation of type 1 diabetes in 6–21% of adults at the time of diagnosis. In those with known diabetes, risk factors include infections, intercurrent illness, psychological stress, and omission or underdosing of insulin (22). A subset of individuals with type 1 diabetes has recurrent episodes of DKA, with some studies suggesting that 22% of adults admitted for DKA in the U.S. are readmitted for the same diagnosis within the following year, 14% of whom have four or more subsequent admissions.

Recurrent DKA is more common in younger adults, women, and those of low socioeconomic status (249). People with recurrent DKA frequently experience high levels of anxiety and diabetes-related distress, difficulties with emotion regulation, and personality dysfunction (250).

The use of SGLT inhibitors in adults with type 1 diabetes increases the risk of DKA by 2.57- to 5.96-fold, even in carefully selected and monitored participants in clinical trials (226). About a third of cases of DKA in the setting of SGLT inhibitor use are euglycemic DKA, suggesting that glucose monitoring alone is insufficient for detection (251). The integration of continuous ketone measurement into CGM systems may provide additional benefits in the future.

DSMES is an effective tool in reducing DKA risk. Additional medical and behavioral health interventions, including home ketone testing and psychosocial support, are often needed. Telemedicine has the potential to reach populations with decreased access to care, and 24-h access to advice about managing hyperglycemia and ketosis or ketonemia at home can reduce the risk of hospital admission (22).

A detailed description of the management of DKA is beyond the scope of this report, but the general principles of treatment are the replacement of fluid, insulin, and potassium. Addressing the underlying cause(s) and contributing factors is essential, as is a careful plan for outpatient follow-up. For further information regarding treatment of DKA, refer to the recent consensus report developed by ADA, EASD, and other organizations (22).

Sick-Day and Illness Management

Stressful events, including illness, may affect glucose levels and increase the risk of DKA. More frequent glucose and ketone measurements are necessary to guide insulin dose adjustments. Individuals should devise a sick-day management plan in consultation with their health care professional (20). Such protocols should specify the amounts and types of fluids and carbohydrates to consume, how often to monitor glucose and ketone levels, how and when to give supplemental insulin, and the circumstances under which a person with diabetes should seek urgent medical care (13). Those who use AID systems

should be aware that the underlying algorithms may not be able to adapt quickly to marked increases in insulin needs, such as following initiation of glucocorticoid therapy or during severe illness. Conversion to open-loop mode may be needed.

SECTION 10: PRESERVATION AND REPLACEMENT OF β -CELL FUNCTION

Key Points

- Current strategies aim to preserve endogenous β -cell function through immunomodulatory and metabolic therapies in early-stage type 1 diabetes.
- Clinical approaches to β -cell replacement include whole-organ pancreas and pancreatic islet transplantation.
- Evolving diabetes technologies are impacting patient eligibility and the role of transplantation in the current treatment landscape.
- Emerging alternatives to donor-derived islets include stem cell-based therapies and porcine islet xenotransplantation.

Prevention of Immune Destruction to Preserve β -Cell Function

Several immunotherapy approaches are being evaluated for their potential use in stage 1 or stage 2 type 1 diabetes to prevent stage 3 clinical type 1 diabetes, and for the preservation of β -cell function before and shortly after the onset of stage 3 clinical type 1 diabetes (252). To date, the most promising results from clinical trials have been seen with the anti-CD3 monoclonal antibody teplizumab (253), low-dose antithymocyte globulin (ATG) (254), anti-TNF- α drug golimumab (255), and JAK inhibitor baricitinib (256). These immunotherapies can preserve β -cell function in recent-onset type 1 diabetes, and, in addition, teplizumab delays the clinical onset of type 1 diabetes (257).

Teplizumab is the first U.S. FDA-approved disease-modifying therapy for type 1 diabetes that delays the onset of stage 3 type 1 diabetes. Although approval is for individuals aged ≥ 8 years who are in stage 2 of the disease, it should be noted that most study participants were young (median age 14 years) and that type 1 diabetes has a slower onset in older adults. Clinical trials have shown that a single 14-day course of teplizumab can delay the onset

of clinical type 1 diabetes by a median of 2–3 years in high-risk individuals (78).

Several trials are underway with the aim of not only preserving but also improving β -cell function, and of interrupting the type 1 diabetes disease process sufficiently to prevent the development of clinical disease (258,259). Family members of individuals with type 1 diabetes are being encouraged to undergo screening for islet autoantibodies. Many countries have established networks that facilitate screening and follow people with the potential to be enrolled in clinical trials (260–263).

Replacement of β -Cells

Whole-organ pancreas transplantation and pancreatic islet transplantation remain the primary clinical methods for β -cell replacement in individuals with type 1 diabetes. Both approaches have demonstrated efficacy in achieving normoglycemia, preventing hypoglycemia, and potentially stabilizing or reversing diabetes-related complications (264–269). However, the necessity for chronic systemic immunosuppression to prevent allogeneic rejection necessitates a careful assessment of the risk/benefit ratio, incorporating both medical and psychological factors (270). Both whole-organ pancreas and pancreatic islet transplantation continue to evolve as viable therapeutic options for individuals with type 1 diabetes. Recent developments have focused on enhancing the efficacy and accessibility of islet transplantation through the exploitation of alternative cell sources and strategies to reduce or avoid the need for immunosuppression (271,272).

Whole-Organ Pancreas Transplantation

The majority of pancreas transplants are performed simultaneously with kidney transplants (simultaneous pancreas and kidney [SPK] transplantation). This represents the standard treatment for individuals with type 1 diabetes and end-stage renal disease, provided there are no contraindications such as malignancies, chronic infections, inadequate self-management, or severe cardiovascular conditions. SPK transplants have demonstrated a 5-year pancreas graft survival rate of $>80\%$, surpassing the outcomes of pancreas transplants alone (PTA) or pancreas after kidney transplants

(264,266). Recipients of SPK transplants often experience significant amelioration of problematic hypoglycemia for extended periods (266,273,274).

PTA is typically considered for younger individuals (under 50 years of age) without obesity (BMI <30 kg/m²) or coronary artery disease (275). These criteria help minimize operative mortality to <1% and reduce early technical pancreas graft loss to <10% (264,275). The primary indications for PTA include a history of frequent, acute, and severe metabolic complications (e.g., hypoglycemia, hyperglycemia, DKA), significant clinical and emotional challenges with exogenous insulin therapy, and consistent failure of insulin-based management, including technological aids (276).

Pancreatic Islet Transplantation

Pancreatic islet transplantation, a less invasive procedure, is indicated for individuals with excessive glycemic lability and frequent severe hypoglycemia despite optimal medical therapy (277). This approach allows for the inclusion of older people and those with coronary artery disease who may not be suitable candidates for whole-pancreas transplantation (278–281). Advancements in patient selection and protocol optimization have led to substantial clinical improvements, leading to the maintenance of insulin independence for 5 years in approximately 50% of recipients (281,282). Beyond achieving insulin independence, recent multicenter clinical trials have emphasized the importance of attaining HbA_{1c} <53 mmol/mol (<7.0%) alongside the elimination of severe hypoglycemia as primary end points, reflecting clinically relevant goals. These outcomes have been associated with improved patient-reported outcomes (265,282–285).

Impact of Advancements in Diabetes Technology on Transplant Eligibility

The continuous advancements in diabetes technology, particularly AID systems, have significantly improved glycemic management. As a result, the number of individuals who meet the strict eligibility criteria for PTA or islet transplantation alone (ITA) has declined. These technological improvements offer a less invasive and lower-risk alternative to transplantation for many people, shifting the risk–benefit balance and reserving PTA and ITA for those with the most severe

glycemic instability or insulin resistance that cannot be managed with current diabetes technology (269,286).

Stem Cell Strategies

A major limiting factor for pancreas or islet transplantation is a limited supply of organs, given the need for cadaver donors. To solve the problem of availability, options under investigation include the use of stem cell–derived islets (287) and xenotransplantation with porcine islets (288,289). Stem cell strategies have used either patient-specific stem cells or universal allogeneic cells. In the former, the patient's own stem cells are reprogrammed or transdifferentiated to become β -cells (290). By contrast, generic allogeneic cells may be centrally produced from a bank of human embryonic stem cells (hESCs) or of induced pluripotent stem cells (iPSCs) and used for multiple recipients (291). One of the key issues is protecting the cells from immune attack, both rejection and recurrent autoimmunity. Three general strategies are being investigated: 1) the use of immunomodulatory drugs, 2) the use of a physical barrier (e.g., encapsulation) (292), and 3) gene editing for immune evasion and/or immune protection (293). Both academic and commercial groups are pursuing these approaches, and some are already in clinical trials. Nonetheless, for stem cell–based strategies, there remain both challenges and opportunities for achieving successful reversal of type 1 diabetes (272).

Regeneration of β -Cells

Several approaches have been studied with the aim of generating or regenerating β -cells, with most studies conducted in isolated cell systems or in animal models (294–296). These approaches include the use of DYRK1A inhibitors, menin inhibitors, and a combination of a GLP-1 receptor agonist, gastrin, and GABA. Clinical trials are expected to begin in the near future.

SECTION 11: SCREENING FOR MICROVASCULAR COMPLICATIONS

Key Points

- Screening for microvascular complications in type 1 diabetes does not need to be performed until 5 years after diagnosis.
- Regular assessment for retinopathy, nephropathy, and neuropathy enables early detection and intervention.

- Fundus photography, a 10-g monofilament test, urinary albumin/creatinine ratio, and eGFR testing are pivotal screening tests.
- Screening frequency should be personalized based on findings, glycemic levels, and comorbidities.
- Optimizing glycemic levels and blood pressure remains central to preventing the progression of complications.

As there are many guidelines available on the management of microvascular complications (16,17), this section focuses on the detection of microvascular complications.

Diabetic Retinopathy

In the modern era of type 1 diabetes management, the incidence and prevalence of diabetic retinopathy has decreased. In a comparison of the U.S. T1D Exchange ($n = 1,283$, mean diabetes duration 32 years) and the German/Austrian DPV ($n = 2014$, diabetes duration 29 years) registries, diabetic retinopathy was reported in 34% and 40% of participants, respectively (297). In the U.S., this compares with a prevalence of 75–82% in the early 2000s (298).

Retinal photography with remote reading by experts can enable screening in areas where qualified eye care professionals are not readily available (299,300) and increase efficiency with reduced costs where these professionals are available (17). Retinal photography can be performed within primary diabetes health care settings and is patient friendly, as pupil dilation is not always required. Images should be interpreted by a trained eye care professional or reading center technician or by approved AI programs. If sight-threatening diabetic retinopathy is noted on screening, referral to an ophthalmologist is recommended. Alternatively, a comprehensive eye examination can also be performed by an ophthalmologist or optometrist.

The initial retinal examination should be performed within 5 years of the onset of type 1 diabetes, with subsequent examinations generally recommended annually for those without or with mild diabetic retinopathy (17). Examinations every 1–2 years or even less frequently may be cost-effective after one or more normal eye examinations (301). More frequent examinations will

be required if retinopathy is progressing and risk factors, such as hyperglycemia and hypertension, are not adequately managed. Similarly, for advanced diabetic retinopathy or macular edema, more frequent examinations are recommended.

Treatment by primary health care providers should include optimizing glycemic and blood pressure management to reduce the risk or slow the progression of diabetic retinopathy. A recent meta-analysis of fibrates suggested that these drugs reduce the incidence and long-term progression of retinopathy; however, only one of the 17 studies was undertaken in people with type 1 diabetes (302). Given the limited evidence in people with type 1 diabetes, fibrates are not recommended as standard treatment for retinopathy (17).

Diabetic Kidney Disease

Diabetic kidney disease has historically impacted 30–40% of individuals with type 1 diabetes (303); however, with better treatment of glucose levels and blood pressure, the prevalence is decreasing in some (304), but not all (305,306), settings. One report from the U.S. noted a weighted estimate of 21.5%; however, there are large variations in the burden of diabetic kidney disease around the world, largely due to economic factors (307).

Screening for albuminuria should be performed by assessing the urinary albumin/creatinine ratio (uACR) in a random spot urine sample (17). In case of elevated uACR, confirmation by repeated examination is needed. A normal uACR is defined as <3 mg/mmol (<30 mg/g), moderately elevated albuminuria is defined as a uACR ≥ 3 – 30 mg/mmol (>30 – 300 mg/g), and severely elevated albuminuria is defined as a uACR ≥ 30 mg/mmol (≥ 300 mg/g). Because of high biological variability ($>20\%$) in urinary albumin excretion, a diagnosis of elevated albuminuria should be made only if at least two of three uACR measurements collected over a 3- to 6-month period are abnormal, after excluding other causes such as glomerular disease (by urinalysis) and orthostatic proteinuria, particularly in young adults (308). Furthermore transient elevations in uACR may occur due to exercise, intercurrent illness, urinary tract infection,

dehydration, menstruation, or collection and analytical factors.

The eGFR is usually calculated from serum creatinine using a validated formula (17). An eGFR persistently <60 mL/min per 1.73 m² and/or a uACR of >3 mg/mmol (>30 mg/mL) is considered abnormal.

The current recommendation is to assess uACR and eGFR in adults with type 1 diabetes yearly in those with a diabetes duration of ≥ 5 years (303), although less frequent measurement may be considered. Treatment of established moderately elevated albuminuria starts with renin–angiotensin–aldosterone system (RAAS) inhibition. Recent evidence indicates that finerenone produces a significantly greater reduction in uACR than placebo (309). Individuals should be referred to a nephrologist if urinary albumin levels increase progressively, there is a $>30\%$ decrease in eGFR following initiation of hemodynamically active therapy (RAAS inhibitors, mineralocorticoid receptor antagonists, or SGLT2 inhibitors,) or eGFR is <30 mL/min per 1.73 m² (310).

Neuropathy

Recent Danish data show a decreasing and low risk of diabetes-related foot complications in type 1 diabetes, with a cumulative 3-year risk of 0.2% in low-risk individuals (around 50%) and 3.9% in high-risk individuals (311). However, changes in prevalence in other countries are less well documented.

Yearly screening for peripheral neuropathy, also termed distal symmetrical polyneuropathy, is recommended from 5 years after diagnosis (17,312), although recent data strongly suggest that screening intervals can be prolonged in low-risk individuals (311). In low-risk individuals, screening for loss of protective sensibility should be performed using a 10-g monofilament. Full assessment for peripheral neuropathy includes a careful history and assessment of either temperature or pinprick sensation (for small-fiber function) and vibration sensation using a 128-Hz tuning fork (for large-fiber function) (17,312). The most common early symptoms are induced by the involvement of small fibers and include pain and dysesthesia (unpleasant sensations of burning and tingling). The involvement of large fibers may cause balance issues, numbness, and loss of protective sensation.

Importantly, up to 50% of diabetic peripheral neuropathy may be asymptomatic. If this is not recognized and preventive foot care is not implemented, there is a higher risk of diabetes-related foot ulcers and amputations.

While there are many etiologies of peripheral neuropathy, for those with type 1 diabetes it is important to consider hypothyroidism and vitamin B₁₂ deficiency, which are more often seen in this population (313). Other causes should be considered where clinically indicated.

Awareness of diabetic autonomic neuropathy is especially relevant if other microvascular complications are present, specifically diabetic retinopathy and peripheral neuropathy. History includes asking about dizziness, light-headedness or feeling faint on standing, syncope, early satiety, erectile dysfunction, changes in sweating patterns (especially gustatory sweating), or dry cracked skin of the extremities. On examination, resting tachycardia (after ruling out hyperthyroidism and other causes of tachycardia), orthostatic hypotension, or evidence of peripheral dryness or cracking of the skin may be found. While a resting heart rate greater than 100 bpm is generally noted early in those with autonomic neuropathy and can be used in early screening (314), more detailed testing such as measurement of heart rate variability using electrocardiography, heart rate response to standing, heart rate response to a Valsalva maneuver, and systolic response to standing are required for a definitive diagnosis (315).

SECTION 12: CARDIOVASCULAR RISK MANAGEMENT

Key Points

- Cardiovascular risk is substantially increased in adults with type 1 diabetes and requires proactive management.
- Cardiovascular risk management in type 1 diabetes includes striving for optimal glycemic, blood pressure, cholesterol, and body weight management and promoting smoking cessation.
- Studies undertaken in other populations including people with type 2 diabetes suggest that there may be a role for glucagon-like peptide 1 (GLP-1)-based drugs and sodium–glucose cotransporter 2 (SGLT2) inhibitors in preventing CVD in people with type 1 diabetes.

The prevention of CVD in people with type 1 diabetes extends beyond glycemic management to include the optimal management of blood pressure and use of lipid-lowering medication. There is an absence of cardiovascular outcome studies in people with type 1 diabetes, and so extrapolation from studies performed in other populations, mainly those with type 2 diabetes, is unavoidable (15,316). However, there is no reason to believe that preventive treatment in people with type 1 diabetes should be less needed or less effective.

Glycemia

In the EDIC cohort study following the DCCT randomized trial, 42% and 30% reductions in cardiovascular events at 17 years and after 30 years, respectively, were seen in those originally assigned to the intensive therapy cohort compared with those originally assigned to conventional therapy (80,317).

The benefits of GLP-1 receptor agonists and SGLT2 inhibitors in reducing heart failure, cardiovascular mortality, renal function decline, and other emerging outcomes have been shown in diverse populations, including people with type 2 diabetes. People with type 1 diabetes, however, were excluded from these studies because of an increased risk of DKA, especially with SGLT2 inhibitors. Given the effect size in the accumulating outcome studies in other populations, the benefit/risk ratio is likely to be positive, in the absence of recent or recurrent DKA. Currently, several studies are reassessing the use of both classes of drugs for type 1 diabetes.

Blood Pressure

Large RCTs in people without diabetes and Chinese people with type 2 diabetes have demonstrated that treatment of hypertension targeting a blood pressure <120/80 mmHg reduces cardiovascular events (318,319). Blood pressure targets should be personalized, with higher targets in the presence of, for example, orthostatic hypotension. If it can be safely attained, the on-treatment blood pressure goal is <130/80 mmHg, but a target of <120/80 mmHg is encouraged for those with higher cardiovascular or kidney disease risk (15).

ACE inhibitors, angiotensin receptor blockers, calcium antagonists, and

thiazides are recommended first-line therapies, with a preference for RAAS inhibition in case of moderately elevated albuminuria. β -Blockers are generally contraindicated, as they may diminish symptoms of hypoglycemia, and should only be used when clearly indicated and with caution (15).

Cholesterol

An observational study reported that lipid-lowering therapy is associated with a 22–44% reduction in the risk of CVD and death among individuals with type 1 diabetes without a prior history of CVD (320). Based on type 2 diabetes guidelines, statins should be considered for people aged over 40 years, and in those aged between 20 and 39 years when the 10-year cardiovascular risk estimated by one of the risk calculators suitable for people with type 1 diabetes exceeds 10% (119,321,322). Coronary artery calcium scores may be helpful for individuals who are at intermediate risk of atherosclerotic CVD, as a score of 0 substantially decreases the risk estimate. Coronary artery scores are not helpful for assessing the effect of statin therapy, as statins can increase the score by increasing plaque density and stabilization (323). The treatment goal is to aim for at least a 50% decrease from the initial LDL cholesterol level to a target of <1.8 mmol/L (<70 mg/dL) for primary prevention and <1.4 mmol/L (<55 mg/dL) for those with established CVD. Further information related to cholesterol lowering, for example secondary prevention, statin intolerance, and pregnancy, is available elsewhere (15).

Aspirin

Aspirin is indicated for all people with type 1 diabetes and existing CVD but is not generally recommended for primary prevention in type 1 diabetes and is associated with an increased risk of gastrointestinal bleeding (316).

Screening in Asymptomatic Individuals

Investigations for coronary artery disease should be considered in the presence of any of the following: signs or symptoms of cardiac or associated vascular disease: carotid bruits, transient ischemic attacks, stroke, claudication, or peripheral arterial disease or electrocardiographic abnormalities (e.g., Q waves) (15). However,

routine screening for coronary artery disease is not recommended, as it does not improve outcomes as long as atherosclerotic CVD risk factors are treated.

Heart Failure

Heart failure is becoming increasingly common in people with type 1 diabetes, especially women (324). NT-proBNP is recognized as a diagnostic and prognostic marker for heart failure, and its measurement may help identify those at risk (15). RAAS blockers and diuretics remain the preferred agents for early heart failure, but a cardiologist should be involved for those with more advanced disease. Given the improved outcomes seen in heart failure in those with type 2 diabetes and those without diabetes, people with heart failure and type 1 diabetes are likely to benefit from SGLT2 inhibitors. A decision to initiate treatment should be individualized, balancing the likely benefit against the 2.57- to 5.96-fold higher risk for DKA (226). If used, caution is required, with planned ketone monitoring, especially during intercurrent illness or surgical procedures, because of the risk of DKA, which is often euglycemic (228).

SECTION 13: MANAGEMENT OF OBESITY

Key Points

- Overweight and obesity are at least as common in people with type 1 diabetes as in the general population.
- Additional contributing factors include increased food intake to prevent hypoglycemia and the effects of hyperinsulinemia.
- Treatment should include behavioral interventions, pharmacotherapy using second-generation obesity management medication, and, if indicated, bariatric surgery.

Contrary to the common perception that people with type 1 diabetes are lean, obesity rates in type 1 diabetes mirror those in the general population and warrant clinical attention, not least because of the substantially higher risk of CVD (325,326). In addition to the usual etiological factors, recurrent food intake to prevent or correct hypoglycemia and the unphysiological subcutaneous administration of insulin further

contribute to weight gain in this population (327). Consequently, obesity management in type 1 diabetes presents a significant clinical challenge.

Behavioral Modification

Meal Planning

Currently, there is insufficient evidence to support one specific meal planning approach for weight loss in individuals with type 1 diabetes. Nutrition education and individual preferences should be prioritized to ensure long-term sustainability. A 3-month pilot study in young adults found no significant difference in weight loss (mean 2.7 kg) among participants randomized to a hypocaloric low-carbohydrate diet, a hypocaloric moderate low-fat diet, or a Mediterranean diet (328). Additionally, a review of eight studies investigating low-carbohydrate diets in type 1 diabetes reported mixed outcomes regarding changes in BMI and total daily insulin requirements. The limited sample size in these studies precluded a valid meta-analysis (329).

Any meal planning strategy aimed at reducing energy intake in type 1 diabetes must demonstrate safety, particularly concerning the risks of hypoglycemia and hyperglycemia. Very-low-carbohydrate (ketogenic) diets raise ongoing concerns, including increased risks of hypoglycemia, diminished glycogen stores that may blunt the glucagon response, and the potential for relative insulin deficiency leading to DKA (329,330).

Exercise

As in the general population, there is limited research specifically examining weight loss through exercise in individuals with type 1 diabetes. Although exercise offers numerous health benefits, a meta-analysis of 24 studies found no consistent reduction in BMI associated with physical activity alone (331). However, regular exercise when combined with strategies such as reducing total daily insulin doses and avoiding excessive energy intake to treat hypoglycemia may aid with weight loss and blunt loss of lean body mass (102).

Pharmacotherapy

First-Generation Obesity Management Medications

For individuals with diabetes, current guidelines recommend considering obesity medications for those with BMI

>27 kg/m². In the U.S. six medications are approved for chronic weight management, with three approved for short-term use only (phentermine, phendimetrazine, and diethylpropion) (332). None of these are approved in Europe, and none have been studied in individuals with type 1 diabetes. All are rarely used today in this population.

Metformin and SGLT2 inhibitors can cause small weight reductions when used as adjunctive agents but should not be used as treatments for weight loss (Table 6).

GLP-1 and Dual Receptor Agonists

There are currently six GLP-1 or dual receptor agonists available, all of which are approved for the treatment of type 2 diabetes. Among these, liraglutide, semaglutide, and tirzepatide are also approved for the treatment of obesity. While the prescribing information for these medications explicitly states that they are not approved for diabetes management in type 1 diabetes, the labeling for their obesity indications does not mention type 1 diabetes. As such, when prescribed for obesity, these agents are not contraindicated in individuals with type 1 diabetes and may be considered for selected people under appropriate clinical supervision.

Retrospective observational studies have demonstrated the potential effectiveness of GLP-1 receptor agonists in individuals with type 1 diabetes. In one study with a matched control group, participants receiving treatment experienced a mean weight loss of 7.2 kg over 12 months, compared with a 1.0-kg weight gain in the control group (333). Additionally, there was a modest, nonsignificant improvement in HbA_{1c} of 3 mmol/mol (0.3%), along with a statistically significant reduction in hypoglycemia as measured by CGM.

Another study reported a 10.1% reduction in body weight at 8 months among individuals with type 1 diabetes treated with tirzepatide (223). However, the use of these agents in this population requires support for insulin dose adjustments in response to the effects of incretin therapy, whether individuals are using MDI or insulin pump therapy, including AID systems (334). It is also important to consider the need for insulin dose adjustments in the context of weight loss. When initiating GLP-1 receptor agonist therapy, delayed gastric emptying,

typically a short-term effect, can increase the risk of immediate postprandial hypoglycemia, particularly during the early phase of treatment or in the presence of nausea or vomiting.

Bariatric Surgery

People with a BMI >35 kg/m² accompanied by weight-related complications, or a BMI >40 kg/m², meet the criteria for bariatric surgery (335). To date, bariatric surgery remains the most effective long-term treatment for severe obesity, although it carries the highest risk among available interventions; these include both short-term surgical risks and longer-term medical risks.

As with the use of obesity management medications, the evidence base for bariatric surgery is still emerging in individuals with type 1 diabetes. A systematic review and meta-analysis involving over 600 individuals with type 1 diabetes reported substantial benefits after just under 3 years, including a reduction in mean BMI from 42.6 kg/m² to 29 kg/m², decreased insulin requirements, and improved HbA_{1c} levels (336).

However, important clinical questions remain regarding which individuals with type 1 diabetes and severe obesity are most likely to benefit from surgical intervention, and which type of procedure should be recommended. Hypoglycemia is the most frequently reported complication following bariatric surgery in this population, with an incidence exceeding 50% (337,338). Furthermore, there have been several cases of iatrogenic DKA when insulin is stopped after surgery, in line with guidance for type 2 diabetes.

SECTION 14: OLDER PEOPLE

Key Points

- Patient safety, particularly avoidance of hypoglycemia, is a key priority for older individuals.
- Glycemic treatment approaches and targets should be based on functional and cognitive status, available care partner support, history of hypoglycemia and impaired awareness, fear of hypoglycemia and hyperglycemia, and life expectancy, rather than chronological age.
- The use of advanced technologies in older individuals living independently, including those with mild cognitive impairment, is effective and safe and

should not be discontinued or a priori excluded solely based on age.

- Simplification of insulin management may be needed to maintain patient safety and prevent diabetes symptoms. In some cases, this may require insulin to be administered by a caregiver or the older person to be supervised while self-dosing.

The population of older adults with type 1 diabetes is growing because of the increasing incidence of type 1 diabetes, more individuals being diagnosed in adulthood, and rising life expectancy (339,340). This group of older adults is highly heterogeneous. Many older adults remain healthy and should be managed in a similar way to younger individuals. However, as comorbidities and frailty develop, reassessment of diabetes management, including potential modifications to insulin therapy, is essential.

Glycemic treatment strategies and targets should be personalized based on functional and cognitive status, availability of care partner support, history of hypoglycemia and IAH, fear of both hypoglycemia and hyperglycemia, and life expectancy, rather than being based on chronological age. Patient safety is paramount. Given the heightened vulnerability to hypoglycemia, older adults with type 1 diabetes may require adjusted glucose targets to reduce the risk of hypoglycemia (18). If insulin administration becomes challenging, simplification of insulin regimens may be appropriate, especially in those with frailty or significant functional or cognitive decline (18).

Advanced diabetes technologies can be highly beneficial in older adults and should not be discontinued or excluded solely based on age (170,341,342). CGM reduces hypoglycemia without increasing hyperglycemia in both healthy older adults and those with mild cognitive impairment (170,341). However, older individuals may be less familiar with and more apprehensive about new technologies. Clear explanations and training, tailored to both the individual and their caregivers, are essential. Training typically takes longer in older adults than in younger individuals (343), but, with adequate education and support, older adults can use these tools effectively, improve glycemic levels, and feel safer.

Alarm settings should be personalized to meet the needs and preferences of the individual, and data sharing with a supportive caregiver can be especially helpful as health declines (344). When selecting new devices, factors such as usability (affected by dexterity, vision, hearing, and cognition) and cost should be considered.

Similar to CGM, AID systems reduce hypoglycemia and improve TIR in older adults who are healthy or who have mild cognitive impairment (345–348). A personalized approach is often needed during both initial training and ongoing use. The potential role of a caregiver should be evaluated, as they may also require training, although the autonomy of the older adult should be preserved as much as possible.

Other technologies, such as mobile apps for reminders and activity tracking, or digital home assistants, have not been extensively studied in this population but may support independence in those with specific impairments. Given the diversity and evolving health status of older adults with type 1 diabetes, and the growing availability of advanced technologies, treatment strategies including the use of newer devices should be regularly reevaluated as health, living situations, support systems, and device usability change.

When someone moves into a residential care home, insulin should not be withheld but, where possible and safe, the insulin regimen should be simplified and glucose targets relaxed to avoid the risk of hypoglycemia (18). Appropriate staff training and clear protocols that include written insulin administration plans and hypoglycemia management protocols are essential.

SECTION 15: PREGNANCY INCLUDING PRECONCEPTION AND POSTNATAL CARE

Key Points

- Health care professionals should discuss pregnancy prior to conception, including the effects of pregnancy on diabetes and vice versa, glycemic targets, and a review of medications.
- All pregnant individuals with type 1 diabetes should be referred to a clinic with expertise in the management of diabetes in pregnancy if available.

- Glycemic targets are more stringent during pregnancy to avoid adverse pregnancy outcomes. These goals should be discussed and reviewed with the pregnant individual.
- Insulin requirements fluctuate significantly during pregnancy and postpartum and with breastfeeding. Frequent glucose monitoring is required to guide dose adjustments.
- Automated insulin delivery (AID) systems should be made available prior to, during, and after pregnancy, but health care professionals should be aware that not all AID systems are approved for use in pregnancy.

Preconception Planning

Effective pregnancy management in type 1 diabetes begins before conception, as planned pregnancies are associated with better outcomes for both the mother and the child. Glycemic levels should be optimized before conception, as this reduces the risk of congenital anomalies, miscarriage, and other adverse pregnancy outcomes (349,350). Ideally, women should maintain a near-normal HbA_{1c}, preferably <48 mmol/mol (6.5%), at least 3 months prior to conception, as this lowers the risk of adverse pregnancy outcomes related to hyperglycemia during embryogenesis.

All medications should be reviewed and any associated with fetal harm should be removed. RAAS inhibitors are contraindicated in pregnancy because of well-established fetal risks and should be discontinued before conception and replaced with pregnancy-safe agents such as nifedipine or labetalol (19). Statins, which are commonly taken by people with type 1 diabetes, should not be used during pregnancy.

Education should be provided on the need for more intensive blood glucose monitoring and insulin dose adjustments and the importance of maintaining a balanced diet. Folic acid supplementation (typically 5 mg daily) is strongly recommended before conception and during early pregnancy to reduce the risk of neural tube defects, which is elevated in women with diabetes.

Managing diabetes-related complications prior to pregnancy, such as hypertension, nephropathy, and retinopathy, is essential. Optimizing blood pressure

and glucose levels, along with comprehensive eye examinations, can help prevent the progression of these conditions during pregnancy. Until optimal glucose levels are reached and comorbidities are well-managed, effective contraception should be used. The choice of contraceptive method should be both safe and reliable, tailored to the individual's medical history and personal preferences (19).

During Pregnancy

When possible, individuals should be referred early to a clinic with expertise in the management of diabetes in pregnancy. Pregnant individuals with type 1 diabetes require more intensive and frequent monitoring than is typically available in standard diabetes care. Diabetes in pregnancy is best managed by a multidisciplinary team, including a diabetologist or endocrinologist, obstetrician, dietitian, diabetes nurse or educator, and diabetes midwife (19). A detailed discussion of pregnancy management in individuals with type 1 diabetes is beyond the scope of this report and so only a brief description is given here (19).

Hyperglycemia during pregnancy increases the risk of complications for both the pregnant individual and the developing fetus and may also impact long-term child development (349,350). Therefore, individuals with type 1 diabetes should be supported in achieving optimal glycemic levels.

CGM use has been approved for use in pregnancy in both the U.S. and Europe, depending on the device. The CONCEPTT trial demonstrated that CGM use during pregnancy was associated with improved outcomes (351). As such, CGM should be encouraged during pregnancy, with the understanding that BGM remains necessary in situations where CGM accuracy may be compromised. Importantly, CGM target glucose ranges during pregnancy differ from standard recommendations, with TIR defined as 3.5–7.8 mmol/L (63–140 mg/dL) (Fig. 4) (352,353). Consequently, CGM reports should be interpreted using these adjusted thresholds, and alarm settings may need to be modified to alert users earlier to hyperglycemia while avoiding unnecessary alarms for appropriately lower glucose levels. Where CGM is unavailable, capillary BGM targets are shown in Table 4.

Hypoglycemia remains the primary barrier to optimizing glycemic levels during pregnancy. It is particularly common in the first half of pregnancy, partly due to IAH and pregnancy-related nausea and vomiting (354). Conversely, pregnant individuals with type 1 diabetes are at risk for DKA at lower blood glucose levels than in the nonpregnant state and should receive education on DKA prevention and detection (355).

AID systems are emerging as standard of care for managing type 1 diabetes during pregnancy. Randomized clinical trials have shown that AID systems can improve glycemic levels and quality of life during pregnancy (356–362). However, these studies have not demonstrated significant improvements in fetal outcomes, such as neonatal birth weight, hypoglycemia rates, or perinatal mortality, probably because they were not sufficiently powered to detect differences in these outcomes.

Not all AID systems are approved for use in pregnancy, and regulatory approval varies between countries. ADA recognizes that AID systems may help improve glycemic management during pregnancy in selected individuals with type 1 diabetes. Accordingly AID systems that do not incorporate pregnancy-specific glucose targets or algorithms may be considered when used with assistive techniques and support from experienced multidisciplinary teams (19).

The Endocrine Society and European Society of Endocrinology recommend the use of AID systems where available, provided there is local expertise to support their implementation and management (363). In contrast, NICE explicitly recommends pregnancy-specific AID systems for people with type 1 diabetes who are pregnant or planning pregnancy, with training and support delivered by a specialist multidisciplinary team (364).

The UK National Pregnancy in Diabetes Audit has recently reported that 53% of individuals using pregnancy-specific AID systems had an HbA_{1c} below 43 mmol/mol (6.1%), compared with 36% using an AID system without a pregnancy-specific algorithm and 37% using MDI (365). Use of pregnancy-specific AID systems was also associated with lower rates of preterm birth, large-for-gestational-age infants, and serious adverse outcomes, with comparable rates of neonatal unit admission. Rates of major congenital anomalies, stillbirth, and neonatal death

among those using pregnancy-specific AID systems were the lowest recorded since national data collection began.

Further research is needed to evaluate the efficacy and safety of the various AID systems currently on the market for use during pregnancy.

Postpartum Care

Postpartum care for individuals with type 1 diabetes is a critical aspect of long-term health management and preparation for future pregnancies. Following delivery, hormonal shifts and physiological changes often lead to enhanced insulin sensitivity, resulting in a marked reduction in insulin requirements. During this period, factors such as breastfeeding, irregular sleep, and inconsistent eating patterns can further elevate the risk of hypoglycemia, making careful insulin dose adjustments essential (366,367). Close monitoring of glucose levels is vital to prevent hypoglycemic episodes and maintain stable glycemic levels. AID systems may improve glycemic levels during the peripartum and postpartum periods (359,360).

Breastfeeding is strongly encouraged due to its benefits for both mother and child. However, it can influence insulin needs, often leading to reduced requirements (366). Individuals should receive guidance on balancing insulin therapy with breastfeeding and the importance of maintaining optimal glycemic levels to support effective lactation. Continued education on nutrition and medication taking is essential to support stable glucose levels during the postpartum period.

Finally, this phase offers an opportunity to revisit long-term diabetes management goals and prepare for future pregnancies if desired. Counseling on contraceptive options and timing of subsequent pregnancies should be provided, emphasizing the importance of preconception care to achieve optimal glycemic levels prior to conception.

SECTION 16: IN-HOSPITAL MANAGEMENT

Key Points

- People with type 1 diabetes are at increased risk of developing diabetic ketoacidosis in hospital.
- A target glucose range of 7.8–10.0 mmol/L (140–180 mg/dL) should be

used for most noncritically and critically ill patients. A glucose goal of 5.6–10.0 mmol/L (100–180 mg/dL) may be used in noncritically ill individuals if this can be attained without hypoglycemia.

- Adopting diabetes technology in hospitals may benefit individuals with diabetes by helping overcome the unmet need for better inpatient glycemic management and alleviating the workload of clinical staff. Where feasible, noncritically ill adults with type 1 diabetes should be allowed to use diabetes technology with appropriate support.

It is important that the hospital teams recognize the key differences between type 1 diabetes and type 2 diabetes because of the high risk of DKA if insulin is withheld in people with type 1 diabetes (22). Therefore, inpatients with type 1 diabetes should be clearly identified to avoid common errors, such as omission of mealtime insulin or withholding of basal insulin for procedures or surgery. People with type 1 diabetes, particularly those with concomitant chronic kidney disease, are at higher risk of hypoglycemia, which should be avoided by careful basal insulin dosing, mealtime insulin bolusing based on carbohydrate matching, and correction dosing for hyperglycemia (20).

There have been no large RCTs specifically assessing glycemic targets for inpatients with type 1 diabetes. Therefore, we recommend following type 2 diabetes guidelines, which recommend target glucose ranges of 7.8–10.0 mmol/L (140–180 mg/dL) for the majority of noncritically and critically ill patients (20). A glucose goal of 5.6–10.0 mmol/L (100–180 mg/dL) may be used in noncritically ill individuals if it can be attained without hypoglycemia, for example in individuals continuing to use their home AID system (20).

Data on use of diabetes technologies, including CGM and AID, are still limited in the inpatient setting (368). One of the reasons is lack of formal approval of those devices for inpatient use by the U.S. FDA (369). Adopting diabetes technology in hospitals may potentially benefit individuals with diabetes by helping overcome the unmet need for better inpatient glycemic management while alleviating the workload of clinical staff (368). Evidence supporting CGM effectiveness

and safety in noncritically ill inpatients is growing, including in those on hemodialysis, and after abdominal surgery and solid organ transplantation (370–376). Hospital-wide CGM policies with electronic health record integration have been used successfully, with good satisfaction reported by nurses and adults with diabetes (377). The evidence concerning effective usage of AID systems in hospitalized individuals, however, is quite limited (378). Although randomized trials are lacking, the benefits of AID use in hospitalized individuals with type 1 diabetes may include improved glycemic outcomes, reduced staff workload, and increased patient satisfaction (369). However, the use of AID should be withheld in people hospitalized for intercurrent illness and/or receiving drugs (e.g., steroids) that significantly impair glucose homeostasis, as well as in people who are no longer capable of managing AID, unless appropriately trained staff are available for support.

Noncritically ill adults with type 1 diabetes using diabetes devices (CGM, insulin pumps, AID systems) should be allowed to use them during outpatient procedures or in inpatient settings when proper supervision is available and the individuals/caregivers have clear mentation, previous training, and education and are capable of managing the device(s) (379–381). Institutions should have clear guidelines and protocols in place to manage inpatient type 1 diabetes safely, including enabling selected adults who can monitor their glucose and self-administer insulin safely to continue managing their diabetes during hospitalization.

If a dedicated inpatient diabetes service is available, it should be consulted for glycemic management, DSMES and discharge planning (20).

CONCLUSION

The 2021 consensus report significantly influenced the clinical management of type 1 diabetes in adults, becoming a landmark reference for health care professionals and informing national and international diabetes policy (4,5). It raised awareness of adult-onset type 1 diabetes, endorsed personalized glycemic targets and promoted the adoption of CGM and TIR metrics. Importantly, it underscored the emotional burden of living with diabetes and the need to embed psychosocial care into routine clinical

practice. This updated report reflects the continued evolution of type 1 diabetes management, incorporating advances in diagnosis, therapy, and technology. Yet, the writing group acknowledges persistent and substantial evidence gaps across prevention, diagnosis, and treatment. People with type 1 diabetes deserve high-quality research to guide their care. We also recognize the ongoing disparities in access to treatment and advocate strongly for equitable, person-centered services to ensure that all individuals with type 1 diabetes receive the care they need and deserve.

Acknowledgments. We acknowledge R. Bannuru, K. Balapattabi, N. El-Sayed, S. Challa SivaKanaka and R. Kalyani (ADA) and P. Niemann, M. Schöнке, L. Rother, N. Buckley-Muehge, M. Meireles and M. Osadolor (EASD), the Committee for Clinical Affairs and Board of the EASD, and the Professional Practice Committee of the ADA for their coordination, guidance, and support throughout the development process. We also acknowledge those who responded to the public consultation and thank the peer reviewers for their thoughtful input and valuable feedback, which helped strengthen the report. Finally, we acknowledge A. Jones (University of Exeter, Exeter, U.K.) for his invaluable and tireless help with section 2.

M.S.K. and J.P. are editors of *Diabetes Care* but were not involved in any of the decisions regarding review of the manuscript or its acceptance.

Funding. The report was jointly commissioned and funded by the ADA and EASD. The authors did not receive any payment for their involvement in the writing group.

Duality of Interest. R.I.G.H. has received honoraria for speaker engagements from Boehringer Ingelheim, Eli Lilly, Encore, Liberum, Novo Nordisk, and ROVI Pharma and consultancy fees from Spotlight-AQ. J.H.D. has served on an advisory board for Liom and was on a speaker's bureau for Novo Nordisk. He is a consultant for Gan & Lee. A.H.-F. is a member of the ADA Board of Directors, editor of ADCEs in Practice, and an auditor for the ADA's Education Recognition Program. She is a participant in a speaker's bureau for Abbott Diabetes Care and Xeris. She is also a member of Xeris's advisory board. I.B.H. receives industry research funding from COUR Pharma and Tandem. He is a consultant for Roche and Abbott Diabetes Care. T.K. has served on advisory boards for Abbott, Ascensia, Bioton, Boehringer Ingelheim, Dexcom, Eli Lilly, Medtronic, Roche, Sanofi, and Ypsomed. He has received research funding from Medtronic and is a participant in speakers' bureaus for Abbott, Ascensia, Bioton, Boehringer Ingelheim, Eli Lilly, Medtronic, Novo Nordisk, Roche, Sanofi, and Servier. K.N. receives research funding from, is a member of the advisory board for, and is a stockholder in Novo Nordisk. She is an advisory board member for Medtronic, Tandem, Convatec, and Abbott Diabetes Care and receives

research funding from Dexcom, Medtronic, and Zealand Pharma. J.P. is a consultant to Sanofi, Novo Nordisk, and Kriya therapeutics. E.R. serves on the advisory board for Abbott, Air Liquide SI, Dexcom, Insulet, Sanofi, Roche, Novo Nordisk, and Eli Lilly and has received research support from Dexcom and Tandem. J.S.S. is a member of the board of directors for Elvinix and SAB-BIO. He has been an advisor and consultant to 4Immune, AbbVie, Abvance, ActoBiotics, Adocia, AiTA, AstraZeneca, Avotres, Bayer, Biomea Fusion, COUR Pharmaceuticals, Dexcom, Eli Lilly, ICON, Immuthera, Inhale Biologics, Integrated Nanotherapeutics, i2o Inc, Kriya Therapeutics, Novo Nordisk, Oramed, Regor, Remedy Plan, Respond Health, Sanofi, Signos, Structure Therapeutics, Vertex, vTv, and WiNK. He is a shareholder or option holder in 4Immune, Abvance, AiTA, Avotres, Dexcom, Elvinix, Oramed SAB-BIO, Signos, vTv, and WiNK. F.J.S. is a consultant for Abbott, Eli Lilly, Sanofi, and Novo Nordisk and serves on the speakers' bureaus for Abbott and Sanofi. He has received research funding from Sanofi and Novo Nordisk. R.S.W. has received research funding from Eli Lilly, Tandem, Insulet, Diasome, DEKA Research and Development Corp/Sequel Med Tech, MannKind, and Amgen. A.L.P. serves on the advisory board for Vertex Pharmaceuticals and Medscape. She has received research support from Abbott Diabetes Care, Insulet, and Zucara. The authors declare that there are no other relationships or activities that might bias, or be perceived to bias, their work.

Author Contributions. R.I.G.H. and A.L.P. were co-chairs for the consensus statement writing group. A.H.-F., I.B.H., M.S.K., J.P., J.S.S., and R.S.W. were the writing group members for the ADA. J.H.D., T.K., B.L., K.N., E.R., and F.J.S. were the writing group members for the EASD. All authors were responsible for drafting the article and revising it critically for important intellectual content. All authors approved the version to be published.

Data and Resource Availability. No new data were created or analyzed in this article. All data supporting the findings of this consensus report are available within the article and its electronic supplementary material, or from the references cited within the article.

References

- Ogle GD, Wang F, Haynes A, et al. Global type 1 diabetes prevalence, incidence, and mortality estimates 2025: results from the International Diabetes Federation Atlas, 11th Edition, and the T1D Index Version 3.0. *Diabetes Res Clin Pract* 2025;225:112277
- Fang M, Wang D, Echouffo-Tcheugui JB, Selvin E. Age at Diagnosis in U.S. Adults With Type 1 Diabetes. *Ann Intern Med* 2023;176:1567–1568
- Miller RG, Secrest AM, Sharma RK, Songer TJ, Orchard TJ. Improvements in the life expectancy of type 1 diabetes: the Pittsburgh Epidemiology of Diabetes Complications study cohort. *Diabetes* 2012;61:2987–2992
- Holt RIG, DeVries JH, Hess-Fischl A, et al. The management of type 1 diabetes in adults. A consensus report by the American Diabetes Association (ADA) and the European Association for the Study of Diabetes (EASD). *Diabetologia* 2021;64:2609–2652
- Holt RIG, DeVries JH, Hess-Fischl A, et al. The management of type 1 diabetes in adults. A Consensus Report by the American Diabetes Association (ADA) and the European Association for the Study of Diabetes (EASD). *Diabetes Care* 2021;44:2589–2625
- American Diabetes Association Professional Practice Committee for Diabetes. 9. Pharmacologic approaches to glycemic treatment: Standards of Care in Diabetes—2026. *Diabetes Care* 2026;49:S183–S215
- International Diabetes Federation. Type 1 diabetes, 2026. Accessed 2 June 2026. Available from <https://idf.org/about-diabetes/type-1-diabetes/>
- National Institute for Health and Care Excellence. Type 1 diabetes in adults: diagnosis and management, 2015. NICE guideline [NG17]. Accessed 17 July 2026. Available from <https://www.nice.org.uk/guidance/ng17>
- Gattrell WT, Logullo P, van Zuuren EJ, et al. ACCORD (ACcurate CONsensus Reporting Document): a reporting guideline for consensus methods in biomedicine developed via a modified Delphi. *PLoS Med* 2024;21:e1004326
- American Diabetes Association Professional Practice Committee for Diabetes. Introduction and methodology: Standards of Care in Diabetes—2026. *Diabetes Care* 2026;49:S1–S5
- American Diabetes Association Professional Practice Committee for Diabetes. 2. Diagnosis and classification of diabetes: Standards of Care in Diabetes—2026. *Diabetes Care* 2026;49:S27–S49
- American Diabetes Association Professional Practice Committee for Diabetes. 5. Facilitating positive health behaviors and well-being to improve health outcomes: Standards of Care in Diabetes—2026. *Diabetes Care* 2026;49:S89–S131
- American Diabetes Association Professional Practice Committee for Diabetes. 6. Glycemic goals, hypoglycemia, and hyperglycemic crises: Standards of Care in Diabetes—2026. *Diabetes Care* 2026;49:S132–S149
- American Diabetes Association Professional Practice Committee for Diabetes. 7. Diabetes technology: Standards of Care in Diabetes—2026. *Diabetes Care* 2026;49:S150–S165
- American Diabetes Association Professional Practice Committee for Diabetes. 10. Cardiovascular disease and risk management: Standards of Care in Diabetes—2026. *Diabetes Care* 2026;49:S216–S245
- American Diabetes Association Professional Practice Committee for Diabetes. 11. Chronic kidney disease and risk management: Standards of Care in Diabetes—2026. *Diabetes Care* 2026;49:S246–S260
- American Diabetes Association Professional Practice Committee for Diabetes. 12. Retinopathy, neuropathy, and foot care: Standards of Care in Diabetes—2026. *Diabetes Care* 2026;49:S261–S276
- American Diabetes Association Professional Practice Committee for Diabetes. 13. Older adults: Standards of Care in Diabetes—2026. *Diabetes Care* 2026;49:S277–S296
- American Diabetes Association Professional Practice Committee for Diabetes. 15. Management of diabetes in pregnancy: Standards of Care in Diabetes—2026. *Diabetes Care* 2026;49:S321–S338
- American Diabetes Association Professional Practice Committee for Diabetes. 16. Diabetes care in the hospital: Standards of Care in Diabetes—2026. *Diabetes Care* 2026;49:S339–S355
- Phillip M, Achenbach P, Addala A, et al. Consensus guidance for monitoring individuals with islet autoantibody-positive pre-stage 3 type 1 diabetes. *Diabetes Care* 2024;47:1276–1298
- Umpierrez GE, Davis GM, ElSayed NA, et al. Hyperglycaemic crises in adults with diabetes: a consensus report. *Diabetologia* 2024;67:1455–1479
- Sherr JL, Heinemann L, Fleming GA, et al. Automated insulin delivery: benefits, challenges, and recommendations. A consensus report of the Joint Diabetes Technology Working Group of the European Association for the Study of Diabetes and the American Diabetes Association. *Diabetes Care* 2022;45:3058–3074
- Moser O, Zaharieva DP, Adolfsson P, et al. The use of automated insulin delivery around physical activity and exercise in type 1 diabetes: a position statement of the European Association for the Study of Diabetes (EASD) and the International Society for Pediatric and Adolescent Diabetes (ISPAD). *Diabetologia* 2025;68:255–280
- Fleming GA, Petrie JR, Bergenstal RM, Holl RW, Peters AL, Heinemann L. Diabetes digital app technology: benefits, challenges, and recommendations. A consensus report by the European Association for the Study of Diabetes (EASD) and the American Diabetes Association (ADA) Diabetes Technology Working Group. *Diabetologia* 2020;63:229–241
- Kovacs A, Bunduc S, Veres DS, et al. One third of cases of new-onset diabetic ketosis in adults are associated with ketosis-prone type 2 diabetes—a systematic review and meta-analysis. *Diabetes Metab Res Rev* 2024;40:e3743
- Michels AW, Brusko TM, Evans-Molina C, Homann D, Richardson SJ, Powers AC. Challenges and opportunities for understanding the pathogenesis of type 1 diabetes: an Endocrine Society scientific statement. *J Clin Endocrinol Metab* 2025;110:2496–2508
- Davis AK, DuBose SN, Haller MJ, et al.; T1D Exchange Clinic Network. Prevalence of detectable C-peptide according to age at diagnosis and duration of type 1 diabetes. *Diabetes Care* 2015;38:476–481
- Umpierrez GE, Smiley D, Kitabchi AE. Narrative review: ketosis-prone type 2 diabetes mellitus. *Ann Intern Med* 2006;144:350–357
- Thomas NJ, Lynam AL, Hill AV, et al. Type 1 diabetes defined by severe insulin deficiency occurs after 30 years of age and is commonly treated as type 2 diabetes. *Diabetologia* 2019;62:1167–1172
- Muñoz C, Floreen A, Garey C, et al. Misdiagnosis and diabetic ketoacidosis at diagnosis of type 1 diabetes: patient and caregiver perspectives. *Clin Diabetes* 2019;37:276–281
- Hope SV, Wienand-Barnett S, Shepherd M, et al. Practical classification guidelines for diabetes in patients treated with insulin: a cross-sectional study of the accuracy of diabetes diagnosis. *Br J Gen Pract* 2016;66:e315–322
- Shields BM, Peters JL, Cooper C, et al. Can clinical features be used to differentiate type 1 from type 2 diabetes? A systematic review of the literature. *BMJ Open* 2015;5:e009088
- Thomas NJ, Jones SE, Weedon MN, Shields BM, Oram RA, Hattersley AT. Frequency and phenotype of type 1 diabetes in the first six decades of life: a cross-sectional, genetically stratified survival analysis from UK Biobank. *Lancet Diabetes Endocrinol* 2018;6:122–129

35. Hillier TA, Pedula KL. Characteristics of an adult population with newly diagnosed type 2 diabetes: the relation of obesity and age of onset. *Diabetes Care* 2001;24:1522–1527
36. Westphal SA. The occurrence of diabetic ketoacidosis in non-insulin-dependent diabetes and newly diagnosed diabetic adults. *Am J Med* 1996;101:19–24
37. Nakagami T, Qiao Q, Carstensen B, et al.; DECODE-DECODA Study Group. Age, body mass index and type 2 diabetes-associations modified by ethnicity. *Diabetologia* 2003;46:1063–1070
38. Prior MJ, Prout T, Miller D, Ewart R, Kumar D. C-peptide and the classification of diabetes mellitus patients in the Early Treatment Diabetic Retinopathy Study. Report number 6. The ETDRS Research Group. *Ann Epidemiol* 1993;3:9–17
39. Chung WK, Erion K, Florez JC, et al. Precision medicine in diabetes: a consensus report from the American Diabetes Association (ADA) and the European Association for the Study of Diabetes (EASD). *Diabetologia* 2020;63:1671–1693
40. Buzzetti R, Tuomi T, Mauricio D, et al. Management of latent autoimmune diabetes in adults: a consensus statement from an international expert panel. *Diabetes* 2020;69:2037–2047
41. Jones AG, McDonald TJ, Shields BM, Hagopian W, Hattersley AT. Latent autoimmune diabetes of adults (LADA) is likely to represent a mixed population of autoimmune (type 1) and nonautoimmune (type 2) diabetes. *Diabetes Care* 2021;44:1243–1251
42. Sabbah E, Savola K, Ebeling T, et al. Genetic, autoimmune, and clinical characteristics of childhood- and adult-onset type 1 diabetes. *Diabetes Care* 2000;23:1326–1332
43. Bingley PJ. Clinical applications of diabetes antibody testing. *J Clin Endocrinol Metab* 2010;95:25–33
44. Littorin B, Sundkvist G, Hagopian W, et al. Islet cell and glutamic acid decarboxylase antibodies present at diagnosis of diabetes predict the need for insulin treatment. A cohort study in young adults whose disease was initially labeled as type 2 or unclassifiable diabetes. *Diabetes Care* 1999;22:409–412
45. Lynam A, McDonald T, Hill A, et al. Development and validation of multivariable clinical diagnostic models to identify type 1 diabetes requiring rapid insulin therapy in adults aged 18–50 years. *BMJ Open* 2019;9:e031586
46. Tridgell DM, Spiekerman C, Wang RS, Greenbaum CJ. Interaction of onset and duration of diabetes on the percent of GAD and IA-2 antibody-positive subjects in the type 1 diabetes genetics consortium database. *Diabetes Care* 2011;34:988–993
47. Thomas NJ, Walkey HC, Kaur A, et al. The relationship between islet autoantibody status and the genetic risk of type 1 diabetes in adult-onset type 1 diabetes. *Diabetologia* 2023;66:310–320
48. Thomas NJ, Jones AG. The challenges of identifying and studying type 1 diabetes in adults. *Diabetologia* 2023;66:2200–2212
49. Eason RJ, Thomas NJ, Hill AV, et al.; StartRight Study Group. Routine islet autoantibody testing in clinically diagnosed adult-onset type 1 diabetes can help identify misclassification and the possibility of successful insulin cessation. *Diabetes Care* 2022;45:2844–2851
50. Rohlfing C, Petroski G, Connolly SM, Hanson S, Little RR, Kabytaev K. A comparative analysis of current C-peptide assays compared to a reference method: can we overcome inertia to standardization? *Clin Chem Lab Med* 2025;63:1124–1131
51. Schleicher E, Kabytaev K, Nauck M, et al. Call for standardization of C-peptide measurement. *J Diabetes Sci Technol* 2025;19:322968251362848
52. Little RR, Wielgosz RI, Josephs R, et al. Implementing a reference measurement system for C-peptide: successes and lessons learned. *Clin Chem* 2017;63:1447–1456
53. Foteinopoulou E, Clarke CAL, Pattenden RJ, et al. Impact of routine clinic measurement of serum C-peptide in people with a clinician-diagnosis of type 1 diabetes. *Diabet Med* 2021;38:e14449
54. Balasubramanyam A, Garza G, Rodriguez L, et al. Accuracy and predictive value of classification schemes for ketosis-prone diabetes. *Diabetes Care* 2006;29:2575–2579
55. Hohberg C, Pfützner A, Forst T, et al. Successful switch from insulin therapy to treatment with pioglitazone in type 2 diabetes patients with residual beta-cell function: results from the PioSwitch study. *Diabetes Obes Metab* 2009;11:464–471
56. Lee A, Morley J. Classification of type 2 diabetes by clinical response to metformin-troglitazone combination and C-peptide criteria. *Endocr Pract* 1999;5:305–313
57. Bell DS, Mayo MS. Improved glycemic control with use of oral hypoglycemic therapy with or without insulin. *Endocr Pract* 1998;4:82–85
58. Hendriks AEJ, Marcovecchio ML, Evans ML, et al.; INNODIA Consortium. Early detection of β -cell decline using home dried blood spot c-peptide levels in new-onset type 1 diabetes. *Diabetes Care* 2025;48:1484–1492
59. Shields BM, Shepherd M, Hudson M, et al.; UNITED study team. Population-based assessment of a biomarker-based screening pathway to aid diagnosis of monogenic diabetes in young-onset patients. *Diabetes Care* 2017;40:1017–1025
60. Pearson ER, Starkey BJ, Powell RJ, Gribble FM, Clark PM, Hattersley AT. Genetic cause of hyperglycaemia and response to treatment in diabetes. *Lancet* 2003;362:1275–1281
61. Greeley SAW, Polak M, Njølstad PR, et al. ISPAD Clinical Practice Consensus Guidelines 2022: the diagnosis and management of monogenic diabetes in children and adolescents. *Pediatr Diabetes* 2022;23:1188–1211
62. De Franco E, Flanagan SE, Houghton JAL, et al. The effect of early, comprehensive genomic testing on clinical care in neonatal diabetes: an international cohort study. *Lancet* 2015;386:957–963
63. Pearson ER, Flechtner I, Njølstad PR, et al.; Neonatal Diabetes International Collaborative Group. Switching from insulin to oral sulfonylureas in patients with diabetes due to Kir6.2 mutations. *N Engl J Med* 2006;355:467–477
64. Carlsson A, Shepherd M, Ellard S, et al. Absence of islet autoantibodies and modestly raised glucose values at diabetes diagnosis should lead to testing for MODY: lessons from a 5-year pediatric Swedish national cohort study. *Diabetes Care* 2020;43:82–89
65. Shields BM, McDonald TJ, Ellard S, Campbell MJ, Hyde C, Hattersley AT. The development and validation of a clinical prediction model to determine the probability of MODY in patients with young-onset diabetes. *Diabetologia* 2012;55:1265–1272
66. Misra S, Shields B, Colclough K, et al. South Asian individuals with diabetes who are referred for MODY testing in the UK have a lower mutation pick-up rate than white European people. *Diabetologia* 2016;59:2262–2265
67. Besser REJ, Shepherd MH, McDonald TJ, et al. Urinary C-peptide creatinine ratio is a practical outpatient tool for identifying hepatocyte nuclear factor 1- α /hepatocyte nuclear factor 4- α maturity-onset diabetes of the young from long-duration type 1 diabetes. *Diabetes Care* 2011;34:286–291
68. McDonald TJ, Colclough K, Brown R, et al. Islet autoantibodies can discriminate maturity-onset diabetes of the young (MODY) from type 1 diabetes. *Diabet Med* 2011;28:1028–1033
69. Thanabalasingham G, Pal A, Selwood MP, et al. Systematic assessment of etiology in adults with a clinical diagnosis of young-onset type 2 diabetes is a successful strategy for identifying maturity-onset diabetes of the young. *Diabetes Care* 2012;35:1206–1212
70. Insel RA, Dunne JL, Atkinson MA, et al. Staging presymptomatic type 1 diabetes: a scientific statement of JDRF, the Endocrine Society, and the American Diabetes Association. *Diabetes Care* 2015;38:1964–1974
71. Besser REJ, Ng SM, Gregory JW, Dayan CM, Randell T, Barrett T. General population screening for childhood type 1 diabetes: is it time for a UK strategy? *Arch Dis Child* 2022;107:790–795
72. Mathieu C, Cherubini V, Kapellen T, et al. Advancing early-stage type 1 diabetes care: future focused pathways for identification, screening, diagnosis, and monitoring. *Diabetes Obes Metab* 2026;28:6836–6849
73. Haller MJ, Bell KJ, Besser REJ, et al. ISPAD Clinical Practice Consensus Guidelines 2024: screening, staging, and strategies to preserve beta-cell function in children and adolescents with type 1 diabetes. *Horm Res Paediatr* 2024;97:529–545
74. The Eurodiab Ace Study Group and The Eurodiab Ace Substudy 2 Study Group. (1998) Familial risk of type 1 diabetes in European children. *Diabetologia* 1998;41:1151–1156
75. Karges B, Prinz N, Placzek K, et al. A comparison of familial and sporadic type 1 diabetes among young patients. *Diabetes Care* 2021;44:1116–1124
76. Hemminki K, Li X, Sundquist J, Sundquist K. Familial association between type 1 diabetes and other autoimmune and related diseases. *Diabetologia* 2009;52:1820–1828
77. Turtinen M, Härkönen T, Parkkola A, Ilonen J, Knip M, Finnish Pediatric Diabetes Register. Characteristics of familial type 1 diabetes: effects of the relationship to the affected family member on phenotype and genotype at diagnosis. *Diabetologia* 2019;62:2025–2039
78. Misra S, Shukla AK. Teplizumab: type 1 diabetes mellitus preventable? *Eur J Clin Pharmacol* 2023;79:609–616
79. Nathan DM, Genuth S, Lachin J, et al.; Diabetes Control and Complications Trial Research Group. The effect of intensive treatment of diabetes on the development and progression of long-term complications in insulin-dependent diabetes mellitus. *N Engl J Med* 1993;329:977–986

80. Diabetes Control Complications Trial Epidemiology of Diabetes Interventions Complications Study Research Group. Intensive diabetes treatment and cardiovascular outcomes in type 1 diabetes: the DCCT/EDIC study 30-year follow-up. *Diabetes Care* 2016;39:686–693
81. Malik FS, Sauder KA, Isom S, et al.; SEARCH for Diabetes in Youth Study. Trends in glycemic control among youth and young adults with diabetes: the SEARCH for Diabetes in Youth Study. *Diabetes Care* 2022;45:285–294
82. Foster NC, Beck RW, Miller KM, et al. State of type 1 diabetes management and outcomes from the T1D Exchange in 2016–2018. *Diabetes Technol Ther* 2019;21:66–72
83. Udsen FW, Hangaard S, Bender C, et al. The effectiveness of telemedicine solutions in type 1 diabetes management: a systematic review and meta-analysis. *J Diabetes Sci Technol* 2023;17:782–793
84. Hood KK, Wong JJ. Telehealth for people with diabetes: poised for a new approach. *Lancet Diabetes Endocrinol* 2022;10:8–10
85. Duke DC, Barry S, Wagner DV, Speight J, Choudhary P, Harris MA. Distal technologies and type 1 diabetes management. *Lancet Diabetes Endocrinol* 2018;6:143–156
86. Verhoeven F, Tanja-Dijkstra K, Nijland N, Eysenbach G, van Gemert-Pijnen L. Asynchronous and synchronous teleconsultation for diabetes care: a systematic literature review. *J Diabetes Sci Technol* 2010;4:666–684
87. Davis J, Fischl AH, Beck J, et al. 2022 National standards for diabetes self-management education and support. *Diabetes Care* 2022;45:484–494
88. Chatterjee S, Davies MJ, Heller S, Speight J, Snoek FJ, Khunti K. Diabetes structured self-management education programmes: a narrative review and current innovations. *Lancet Diabetes Endocrinol* 2018;6:130–142
89. Cooke D, Bond R, Lawton J, et al.; U.K. NIHR DAFNE Study Group. Structured type 1 diabetes education delivered within routine care: impact on glycemic control and diabetes-specific quality of life. *Diabetes Care* 2013;36:270–272
90. Michie S, Wood CE, Johnston M, Abraham C, Francis JJ, Hardeman W. Behaviour change techniques: the development and evaluation of a taxonomic method for reporting and describing behaviour change interventions (a suite of five studies involving consensus methods, randomised controlled trials and analysis of qualitative data). *Health Technol Assess* 2015;19:1–188
91. Joubert M, Benhamou P-Y, Schaepelynck P, et al. Remote monitoring of diabetes: a cloud-connected digital system for individuals with diabetes and their health care providers. *J Diabetes Sci Technol* 2019;13:1161–1168
92. Sharma V, Feldman M, Sharma R. Telehealth technologies in diabetes self-management and education. *J Diabetes Sci Technol* 2024;18:148–158
93. Powers MA, Bardsley JK, Cypress M, et al. Diabetes self-management education and support in adults with type 2 diabetes: a consensus report of the American Diabetes Association, the Association of Diabetes Care & Education Specialists, the Academy of Nutrition and Dietetics, the American Academy of Family Physicians, the American Academy of PAs, the American Association of Nurse Practitioners, and the American Pharmacists Association. *Diabetes Care* 2020;43:1636–1649
94. Ahn DT, Stahl R. Is there an app for that? The pros and cons of diabetes smartphone apps and how to integrate them into clinical practice. *Diabetes Spectr* 2019;32:231–236
95. Mozzillo E, Zito E, Maffei C, et al. Unhealthy lifestyle habits and diabetes-specific health-related quality of life in youths with type 1 diabetes. *Acta Diabetol* 2017;54:1073–1080
96. Hassanein M, Afandi B, Yakoob Ahmedani M, et al. Diabetes and Ramadan: practical guidelines 2021. *Diabetes Res Clin Pract* 2022;185:109185
97. Builes-Montañón CE, Ortiz-Cano NA, Ramirez-Rincón A, Rojas-Henao NA. Efficacy and safety of carbohydrate counting versus other forms of dietary advice in patients with type 1 diabetes mellitus: a systematic review and meta-analysis of randomised clinical trials. *J Hum Nutr Diet* 2022;35:1030–1042
98. Tikkanen-Dolenc H, Wadén J, Forsblom C, et al.; FinnDiane Study Group. Frequent and intensive physical activity reduces risk of cardiovascular events in type 1 diabetes. *Diabetologia* 2017;60:574–580
99. Tikkanen-Dolenc H, Wadén J, Forsblom C, et al.; FinnDiane Study Group. Physical activity reduces risk of premature mortality in patients with type 1 diabetes with and without kidney disease. *Diabetes Care* 2017;40:1727–1732
100. Chimen M, Kennedy A, Nirantharakumar K, Pang TT, Andrews R, Narendran P. What are the health benefits of physical activity in type 1 diabetes mellitus? A literature review. *Diabetologia* 2012;55:542–551
101. Tatovic D, Narendran P, Dayan CM. A perspective on treating type 1 diabetes mellitus before insulin is needed. *Nat Rev Endocrinol* 2023;19:361–370
102. De Cock D, Schreurs L, Steenackers N, et al. The effect of physical activity on glycaemic control in people with type 1 diabetes mellitus: a systematic literature review and meta-analysis. *Diabet Med* 2024;41:e15415
103. Tikkanen-Dolenc H, Wadén J, Forsblom C, et al.; FinnDiane Study Group. Frequent physical activity is associated with reduced risk of severe diabetic retinopathy in type 1 diabetes. *Acta Diabetol* 2020;57:527–534
104. Lemaster JW, Reiber GE, Smith DG, Heagerty PJ, Wallace C. Daily weight-bearing activity does not increase the risk of diabetic foot ulcers. *Med Sci Sports Exerc* 2003;35:1093–1099
105. Riddell MC, Gallen IW, Smart CE, et al. Exercise management in type 1 diabetes: a consensus statement. *Lancet Diabetes Endocrinol* 2017;5:377–390
106. Adolfsson P, Taplin CE, Zaharieva DP, et al. ISPAD Clinical Practice Consensus Guidelines 2022: exercise in children and adolescents with diabetes. *Pediatr Diabetes* 2022;23:1341–1372
107. Zimmer RT, Auth A, Schierbauer J, et al. (Hybrid) closed-loop systems: from announced to unannounced exercise. *Diabetes Technol Ther* 22 December 2023 [Epub ahead of print]. DOI: 10.1089/dia.2023.0293
108. Reutrakul S, Thakkinstian A, Anothaisintawee T, et al. Sleep characteristics in type 1 diabetes and associations with glycemic control: systematic review and meta-analysis. *Sleep Med* 2016;23:26–45
109. Perfect MM. Sleep-related disorders in patients with type 1 diabetes mellitus: current insights. *Nat Sci Sleep* 2020;12:101–123
110. van Dijk M, Donga E, van Dijk JG, et al. Disturbed subjective sleep characteristics in adult patients with long-standing type 1 diabetes mellitus. *Diabetologia* 2011;54:1967–1976
111. Denic-Roberts H, Costacou T, Orchard TJ. Subjective sleep disturbances and glycemic control in adults with long-standing type 1 diabetes: the Pittsburgh's Epidemiology of Diabetes Complications study. *Diabetes Res Clin Pract* 2016;119:1–12
112. Charlton J, Gill J, Elliott L, Whittaker A, Farquharson B, Strachan M. A review of the challenges, glycaemic risks and self-care for people with type 1 diabetes when consuming alcoholic beverages. *Practical Diabetes* 2020;37:7–12c
113. Kinney GL, Akturk HK, Taylor DD, Foster NC, Shah VN. Cannabis use is associated with increased risk for diabetic ketoacidosis in adults with type 1 diabetes: findings from the T1D Exchange Clinic Registry. *Diabetes Care* 2020;43:247–249
114. Pastor A, Conn J, MacIsaac RJ, Bonomo Y. Alcohol and illicit drug use in people with diabetes. *Lancet Diabetes Endocrinol* 2020;8:239–248
115. Karakus KE, Akturk HK. Cannabis hyperemesis syndrome can mimic diabetic gastroparesis in cannabis users with type 1 diabetes. *Can J Diabetes* 2025;49:218–219
116. Barnard KD, Dyson P, Sinclair JMA, et al. Alcohol health literacy in young adults with type 1 diabetes and its impact on diabetes management. *Diabet Med* 2014;31:1625–1630
117. Engler PA, Ramsey SE, Smith RJ. Alcohol use of diabetes patients: the need for assessment and intervention. *Acta Diabetol* 2013;50:93–99
118. Uruska A, Araszkiwicz A, Uruski P, Zozulinska-Ziolkiewicz D. Higher risk of microvascular complications in smokers with type 1 diabetes despite intensive insulin therapy. *Microvasc Res* 2014;92:79–84
119. Helmink MAG, Hageman SHJ, Eliasson B, et al. Lifetime and 10-year cardiovascular risk prediction in individuals with type 1 diabetes: the LIFE-T1D model. *Diabetes Obes Metab* 2024;26:2229–2238
120. Pan A, Wang Y, Talaei M, Hu FB. Relation of smoking with total mortality and cardiovascular events among patients with diabetes mellitus: a meta-analysis and systematic review. *Circulation* 2015;132:1795–1804
121. Jensen MH, Cichosz SL, Hirsch IB, Vestergaard P, Hejlesen O, Seto E. Smoking is associated with increased risk of not achieving glycemic target, increased glycemic variability, and increased risk of hypoglycemia for people with type 1 diabetes. *J Diabetes Sci Technol* 2021;15:827–832
122. Tarlton C, James S, Dixon B, Craft J. Travel health practices, behaviours and experiences of people living with type 1 diabetes. *J Travel Med* 2024;31:taae023
123. Cox DJ, Frier BM, Bruggeman B, et al. Diabetes and driving: a statement of the American Diabetes Association. *Diabetes Care* 2024;47:1889–1896
124. Anderson JE, Greene MA, Griffin JW, et al.; American Diabetes Association. Diabetes and

- employment. *Diabetes Care* 2014;37 Suppl 1: S112–S117
125. Durrani T, East B, Fischer A, et al. Employment and diabetes: a statement of the American Diabetes Association. *Diabetes, Obesity, and CardioMetabolic Care* 2026;1:572–581
 126. Gonzalez JS, Tanenbaum ML, Commissariat PV. Psychosocial factors in medication adherence and diabetes self-management: implications for research and practice. *Am Psychol* 2016; 71:539–551
 127. de Groot M, Golden SH, Wagner J. Psychological conditions in adults with diabetes. *Am Psychol* 2016;71:552–562
 128. Hill-Briggs F, Adler NE, Berkowitz SA, et al. Social determinants of health and diabetes: a scientific review. *Diabetes Care* 2020;44:258–279
 129. Franc S, Bensaid S, Schaepelynck P, Orlando L, Lopes P, Charpentier G. Impact of chronic emotions and psychosocial stress on glycemic control in patients with type 1 diabetes. Heterogeneity of glycemic responses, biological mechanisms, and personalized medical treatment. *Diabetes Metab* 2023;49:101486
 130. Nicolucci A, Kovacs Burns K, Holt RIG, et al.; DAWN2 Study Group. Diabetes Attitudes, Wishes and Needs second study (DAWN2): cross-national benchmarking of diabetes-related psychosocial outcomes for people with diabetes. *Diabet Med* 2013;30:767–777
 131. van Duinkerken E, Snoek FJ, de Wit M. The cognitive and psychological effects of living with type 1 diabetes: a narrative review. *Diabet Med* 2020;37:555–563
 132. Fisher L, Polonsky WH, Hessler DM, et al. Understanding the sources of diabetes distress in adults with type 1 diabetes. *J Diabetes Complications* 2015;29:572–577
 133. Housni A, Katz A, Kichler JC, Nakhla M, Secours L, Brazeau A-S. Predictors of stigma perception by people with type 1 diabetes: a cross-sectional analysis of the BETTER registry. *Diabetes Metab Syndr* 2024;18:103112
 134. Hessler DM, Fisher L, Polonsky WH, et al. Diabetes distress is linked with worsening diabetes management over time in adults with Type 1 diabetes. *Diabet Med* 2017;34:1228–1234
 135. Holt RIG. Diabetes and depression. In: Feingold KR, Ahmed SF, Anawalt B, et al. (Eds) *Endotext*. MDText.com, South Dartmouth, MA
 136. Nefs G, Hendrieckx C, Reddy P, et al. Comorbid elevated symptoms of anxiety and depression in adults with type 1 or type 2 diabetes: results from the International Diabetes MILES Study. *J Diabetes Complications* 2019;33: 523–529
 137. Snoek FJ, Bremmer MA, Hermanns N. Constructs of depression and distress in diabetes: time for an appraisal. *Lancet Diabetes Endocrinol* 2015;3:450–460
 138. Pouwer F, Schram MT, Iversen MM, Nouwen A, Holt RIG. How 25 years of psychosocial research has contributed to a better understanding of the links between depression and diabetes. *Diabet Med* 2020;37: 383–392
 139. Sultan S, Epel E, Sachon C, Vaillant G, Hartemann-Hurtier A. A longitudinal study of coping, anxiety and glycemic control in adults with type 1 diabetes. *Psychol Health* 2008;23: 73–89
 140. Wild D, von Maltzahn R, Brohan E, Christensen T, Clauson P, Gonder-Frederick L. A critical review of the literature on fear of hypoglycemia in diabetes: implications for diabetes management and patient education. *Patient Educ Couns* 2007;68:10–15
 141. McKechnie V, Oliver N, Amiel SA, Fox JRE. Hyperglycaemia aversion in type 1 diabetes: a grounded theory study. *Br J Health Psychol* 2024;29:254–271
 142. Jacob P, Potts L, Maclean RH, et al.; HARPdoc study group. Characteristics of adults with type 1 diabetes and treatment-resistant problematic hypoglycaemia: a baseline analysis from the HARPdoc RCT. *Diabetologia* 2022;65: 936–948
 143. Dean YE, Motawea KR, Aslam M, et al. Association between type 1 diabetes mellitus and eating disorders: a systematic review and meta-analysis. *Endocrinol Diabetes Metab* 2024;7: e473
 144. Hirvelä L, Haukka J, Keski-Rahkonen A, Sipilä PN. Eating disorders among people with and without type 1 diabetes: incidence and treatment in a nationwide population-based cohort. *Diabetologia* 2025;68:766–777
 145. Goddard G, Oxlad M. Caring for individuals with type 1 diabetes mellitus who restrict and omit insulin for weight control: evidence-based guidance for healthcare professionals. *Diabetes Res Clin Pract* 2022;185:109783
 146. Xie X-N, Lei X, Xiao C-Y, Li Y-M, Lei X-Y. Association between type 1 diabetes and neurodevelopmental disorders in children and adolescents: a systematic review and meta-analysis. *Front Psychiatry* 2022;13:982696
 147. Liu S, Kuja-Halkola R, Larsson H, et al. Neurodevelopmental disorders, glycemic control, and diabetic complications in type 1 diabetes: a nationwide cohort study. *J Clin Endocrinol Metab* 2021;106:e4459–e4470
 148. Jacobson AM, Ryan CM, Braffett BH, et al.; DCCT/EDIC Research Group. Cognitive performance declines in older adults with type 1 diabetes: results from 32 years of follow-up in the DCCT and EDIC Study. *Lancet Diabetes Endocrinol* 2021;9:436–445
 149. Hermanns N, Kulzer B, Ehrmann D. Person-reported outcomes in diabetes care: what are they and why are they so important? *Diabetes Obes Metab* 2024;26 Suppl 1:30–45
 150. Barnard-Kelly K, Marrero D, de Wit M, et al. Towards the standardisation of adult person-reported outcome domains in diabetes research: a consensus statement development panel. *Diabet Med* 2024;41:e15332
 151. Snoek FJ, Kersch NYA, Eldrup E, et al. Monitoring of Individual Needs in Diabetes (MIND): baseline data from the cross-national Diabetes Attitudes, Wishes, and Needs (DAWN) MIND study. *Diabetes Care* 2011;34:601–603
 152. Pouwer F, Snoek FJ, van der Ploeg HM, Adèr HJ, Heine RJ. Monitoring of psychological well-being in outpatients with diabetes: effects on mood, HbA(1c), and the patient's evaluation of the quality of diabetes care: a randomized controlled trial. *Diabetes Care* 2001;24:1929–1935
 153. Beran M, Muzambi R, Geraets A, et al.; European Depression in Diabetes (EDID) Research Consortium. The bidirectional longitudinal association between depressive symptoms and HbA1c: a systematic review and meta-analysis. *Diabet Med* 2022;39:e14671
 154. d'Emden H, McDermott B, D'Silva N, et al. Psychosocial screening and management of young people aged 18–25 years with diabetes. *Intern Med J* 2017;47:415–423
 155. Hamilton K, Forde R, Due-Christensen M, et al. Which diabetes specific patient reported outcomes should be measured in routine care? A systematic review to inform a core outcome set for adults with Type 1 and 2 diabetes mellitus: the European Health Outcomes Observatory (H2O) programme. *Patient Educ Couns* 2023;116: 107933
 156. Nano J, Carinci F, Okunade O, et al.; Diabetes Working Group of the International Consortium for Health Outcomes Measurement (ICHOM). A standard set of person-centred outcomes for diabetes mellitus: results of an international and unified approach. *Diabet Med* 2020;37:2009–2018
 157. Pursey KM, Hart M, Jenkins L, McEvoy M, Smart CE. Screening and identification of disordered eating in people with type 1 diabetes: a systematic review. *J Diabetes Complications* 2020;34:107522
 158. Fisher L, Guzman S, Polonsky W, Hessler D. Bringing the assessment and treatment of diabetes distress into the real world of clinical care: time for a shift in perspective. *Diabet Med* 2024;41:e15446
 159. Young-Hyman D, de Groot M, Hill-Briggs F, Gonzalez JS, Hood K, Peyrot M. Psychosocial care for people with diabetes: a position statement of the American Diabetes Association. *Diabetes Care* 2016;39:2126–2140
 160. Snoek FJ, Anarte-Ortiz MT, Anderbro T, et al. Roles and competencies of the clinical psychologist in adult diabetes care-A consensus report. *Diabet Med* 2024;41:e15312
 161. Embaye J, de Wit M, Snoek F. A self-guided web-based app (MyDiaMate) for enhancing mental health in adults with type 1 diabetes: insights from a real-world study in the Netherlands. *JMIR Diabetes* 2024;9:e52923
 162. Yakubu TI, Pauer S, West NC, Tang TS, Gorges M. Impact of digitally enabled peer support interventions on diabetes distress and depressive symptoms in people living with type 1 diabetes: a systematic review. *Curr Diab Rep* 2024;25:1
 163. Li Y, Storch EA, Ferguson S, Li L, Buys N, Sun J. The efficacy of cognitive behavioral therapy-based intervention on patients with diabetes: a meta-analysis. *Diabetes Res Clin Pract* 2022;189: 109965
 164. Diribe O, Palmer K, Kennedy A, et al. A systematic literature review of psychological interventions for adults with type 1 diabetes. *Diabetes Ther* 2024;15:367–380
 165. Winkley K, Upsher R, Stahl D, et al. Psychological interventions to improve self-management of type 1 and type 2 diabetes: a systematic review. *Health Technol Assess* 2020; 24:1–232
 166. van der Feltz-Cornelis C, Allen SF, Holt RIG, Roberts R, Nouwen A, Sartorius N. Treatment for comorbid depressive disorder or subthreshold depression in diabetes mellitus: systematic review and meta-analysis. *Brain Behav* 2021;11:e01981
 167. van Steenberghe-Weijnenburg KM, de Vroeghe L, Ploeger RR, et al. Validation of the

- PHQ-9 as a screening instrument for depression in diabetes patients in specialized outpatient clinics. *BMC Health Serv Res* 2010;10:235
168. De Hert M, Dekker JM, Wood D, Kahl KG, Holt RIG, Möller H-J. Cardiovascular disease and diabetes in people with severe mental illness position statement from the European Psychiatric Association (EPA), supported by the European Association for the Study of Diabetes (EASD) and the European Society of Cardiology (ESC). *Eur Psychiatry* 2009;24:412–424
169. Rizos EC, Markozannes G, Charitakis N, et al. Continuous glucose monitoring in type 1 diabetes, type 2 diabetes, and diabetes during pregnancy: a systematic review with meta-analysis of randomized controlled trials. *Diabetes Technol Ther* 2025;27:537–552
170. Pratley RE, Kanapka LG, Rickels MR, et al.; Wireless Innovation for Seniors With Diabetes Mellitus (WISDM) Study Group. Effect of continuous glucose monitoring on hypoglycemia in older adults with type 1 diabetes: a randomized clinical trial. *JAMA* 2020;323:2397–2406
171. van Beers CAJ, DeVries JH, Kleijer SJ, et al. Continuous glucose monitoring for patients with type 1 diabetes and impaired awareness of hypoglycaemia (IN CONTROL): a randomised, open-label, crossover trial. *Lancet Diabetes Endocrinol* 2016;4:893–902
172. Klonoff DC, Gabbay M, Moon SJ, Wilmot EG. Importance of FDA-integrated continuous glucose monitors to ensure accuracy of continuous glucose monitoring. *J Diabetes Sci Technol* 2025;19:1392–1399
173. Mathieu C, Irace C, Wilmot EG, et al. Minimum expectations for market authorization of continuous glucose monitoring devices in Europe—eCGM compliance status. *Diabetes Obes Metab* 2025;27:1025–1031
174. Divilly P, Martine-Edith G, Zaremba N, et al.; Hypo-RESOLVE Consortium. Relationship between sensor-detected hypoglycemia and patient-reported hypoglycemia in people with type 1 and insulin-treated type 2 diabetes: the Hypo-METRICS Study. *Diabetes Care* 2024;47:1769–1777
175. Sacks DB, Arnold M, Bakris GL, et al. Executive summary: guidelines and recommendations for laboratory analysis in the diagnosis and management of diabetes mellitus. *Diabetes Care* 2023;46:1740–1746
176. Kong YW, Morrison D, Lu JC, Lee MH, Jenkins AJ, O'Neal DN. Continuous ketone monitoring: exciting implications for clinical practice. *Diabetes Obes Metab* 2024;26(Suppl 7):47–58
177. Dhatariya K, Bergenstal RM, Al-Sofiani M, et al. Continuous ketone monitoring for people with diabetes: international expert recommendations on the application of a new technology. *Lancet Diabetes Endocrinol* 2026;14:82–92
178. Tozzo V, Genco M, Omololu SO, et al. Estimating glycemia from HbA1c and CGM: analysis of accuracy and sources of discrepancy. *Diabetes Care* 2024;47:460–466
179. Beck RW, Connor CG, Mullen DM, Wesley DM, Bergenstal RM. The fallacy of average: how using HbA1c alone to assess glycemic control can be misleading. *Diabetes Care* 2017;40:994–999
180. Beck RW, Bergenstal RM, Cheng P, et al. The relationships between time in range, hyperglycemia metrics, and HbA1c. *J Diabetes Sci Technol* 2019;13:614–626
181. Yapanis M, James S, Craig ME, O'Neal D, Ekinli EI. Complications of diabetes and metrics of glycemic management derived from continuous glucose monitoring. *J Clin Endocrinol Metab* 2022;107:e2221–e2236
182. Gorst C, Kwok CS, Aslam S, et al. Long-term glycemic variability and risk of adverse outcomes: a systematic review and meta-analysis. *Diabetes Care* 2015;38:2354–2369
183. Hirsch IB, Juneja R, Beals JM, Antalis CJ, Wright EE. The evolution of insulin and how it informs therapy and treatment choices. *Endocr Rev* 2020;41:733–755
184. Russell-Jones D, Babazono T, Cailleateau R, et al. Once-weekly insulin icodec versus once-daily insulin degludec as part of a basal-bolus regimen in individuals with type 1 diabetes (ONWARDS 6): a phase 3a, randomised, open-label, treat-to-target trial. *Lancet* 2023;402:1636–1647
185. Bergenstal RM, Weinstock RS, Mathieu C, et al. Once-weekly insulin efsitora alfa versus once-daily insulin degludec in adults with type 1 diabetes (QWINT-5): a phase 3 randomised non-inferiority trial. *Lancet* 2024;404:1132–1142
186. Masierek M, Nabrdalik K, Janota O, Kwiendacz H, Macherski M, Gumprecht J. The review of insulin pens-past, present, and look to the future. *Front Endocrinol (Lausanne)* 2022;13:827484
187. Gibney MA, Arce CH, Byron KJ, Hirsch LJ. Skin and subcutaneous adipose layer thickness in adults with diabetes at sites used for insulin injections: implications for needle length recommendations. *Curr Med Res Opin* 2010;26:1519–1530
188. Danne TPA, Joubert M, Hartvig NV, Kaas A, Knudsen NN, Mader JK. Association between treatment adherence and continuous glucose monitoring outcomes in people with diabetes using smart insulin pens in a real-world setting. *Diabetes Care* 2024;47:995–1003
189. Hellman J, Hartvig NV, Kaas A, Møller JB, Sørensen MR, Jendle J. Associations of bolus insulin injection frequency and smart pen engagement with glycaemic control in people living with type 1 diabetes. *Diabetes Obes Metab* 2024;26:301–310
190. MacLeod J, Im GH, Smith M, Vigersky RA. Shining the spotlight on multiple daily insulin therapy: real-world evidence of the InPen smart insulin pen. *Diabetes Technol Ther* 2024;26:33–39
191. Adolfsson P, Hartvig NV, Kaas A, Møller JB, Hellman J. Increased time in range and fewer missed bolus injections after introduction of a smart connected insulin pen. *Diabetes Technol Ther* 2020;22:709–718
192. Harbison R, Hecht M, MacLeod J. Building a data-driven multiple daily insulin therapy model using smart insulin pens. *J Diabetes Sci Technol* 2022;16:610–616
193. Boughton CK, Hovorka R. The role of automated insulin delivery technology in diabetes. *Diabetologia* 2024;67:2034–2044
194. Kesavadev J, Srinivasan S, Saboo B, Krishna B M, Krishnan G. The do-it-yourself artificial pancreas: a comprehensive review. *Diabetes Ther* 2020;11:1217–1235
195. Boughton CK, Hovorka R. The artificial pancreas. *Curr Opin Organ Transplant* 2020;25:336–342
196. Heise T, Stender-Petersen K, Hövelmann U, et al. Pharmacokinetic and pharmacodynamic properties of faster-acting insulin aspart versus insulin aspart across a clinically relevant dose range in subjects with type 1 diabetes mellitus. *Clin Pharmacokinet* 2017;56:649–660
197. Kapitzka C, Nowotny I, Lehmann A, et al. Similar pharmacokinetics and pharmacodynamics of rapid-acting insulin lispro products SAR342434 and US- and EU-approved humalog in subjects with type 1 diabetes. *Diabetes Obes Metab* 2017;19:622–627
198. Hirsch IB, Beck RW, Marak MC, et al.; INHALE-3 Study Group. A randomized trial comparing inhaled insulin plus basal insulin versus usual care in adults with type 1 diabetes. *Diabetes Care* 2025;48:353–360
199. Beck RW, Bailey RJ, Klein KR, et al.; INHALE-3 Study Group. Inhaled technosphere insulin plus insulin degludec for adults with type 1 diabetes: the INHALE-3 extension study. *Diabetes Technol Ther* 2025;27:170–178
200. McGill JB, Peters A, Buse JB, et al. Comprehensive pulmonary safety review of inhaled technosphere insulin in patients with diabetes mellitus. *Clin Drug Investig* 2020;40:973–983
201. Spaan NA, Teplova AE, Renard E, Spaan JAE. Implantable insulin pumps: an effective option with restricted dissemination. *Lancet Diabetes Endocrinol* 2014;2:358–360
202. van Dijk PR, Logtenberg SJJ, Groenier KH, Gans ROB, Kleefstra N, Bilo HJ. Continuous intraperitoneal insulin infusion in type 1 diabetes: a 6-year post-trial follow-up. *BMC Endocr Disord* 2014;14:30
203. Dalla Libera A, Toffanin C, Drecogna M, Galderisi A, Pillonetto G, Cobelli C. In silico design and validation of a time-varying PID controller for an artificial pancreas with intraperitoneal insulin delivery and glucose sensing. *APL Bioeng* 2023;7:026105
204. Tian T, Aaron RE, Huang J, et al. Lipohypertrophy and insulin: an update from the Diabetes Technology Society. *J Diabetes Sci Technol* 2023;17:1711–1721
205. Mader JK, Fornengo R, Hassoun A, et al. Relationship between lipohypertrophy, glycemic control, and insulin dosing: a systematic meta-analysis. *Diabetes Technol Ther* 2024;26:351–362
206. Bochanen N, Decochez K, Heleu E, et al. Lipohypertrophy Monitoring Study (LIMO): effect of single use of 4 mm pen needles combined with education on injection site rotation on glycaemic control: confirmation of an unpleasant truth. *Diabet Med* 2022;39:e14672
207. Sola-Gazagnes A, Pecquet C, Berré S, et al. Insulin allergy: a diagnostic and therapeutic strategy based on a retrospective cohort and a case-control study. *Diabetologia* 2022;65:1278–1290
208. Masouri MM, Ebrahimi R, Noori S. An updated systematic review and meta-analysis on the efficacy and safety of metformin as add-on therapy to insulin in patients with type 1 diabetes. *Endocrinol Diabetes Metab* 2025;8:e70060
209. Petrie JR, Chaturvedi N, Ford I, et al.; REMOVAL Study Group. Cardiovascular and metabolic effects of metformin in patients with type 1 diabetes (REMOVAL): a double-blind, randomised, placebo-controlled trial. *Lancet Diabetes Endocrinol* 2017;5:597–609

210. Kong MF, King P, Macdonald IA, et al. Infusion of pramlintide, a human amylin analogue, delays gastric emptying in men with IDDM. *Diabetologia* 1997;40:82–88
211. Kong MF, Stubbs TA, King P, et al. The effect of single doses of pramlintide on gastric emptying of two meals in men with IDDM. *Diabetologia* 1998;41:577–583
212. Fineman MS, Koda JE, Shen LZ, et al. The human amylin analog, pramlintide, corrects postprandial hyperglucagonemia in patients with type 1 diabetes. *Metabolism* 2002;51:636–641
213. Chapman I, Parker B, Doran S, et al. Effect of pramlintide on satiety and food intake in obese subjects and subjects with type 2 diabetes. *Diabetologia* 2005;48:838–848
214. Whitehouse F, Kruger DF, Fineman M, et al. A randomized study and open-label extension evaluating the long-term efficacy of pramlintide as an adjunct to insulin therapy in type 1 diabetes. *Diabetes Care* 2002;25:724–730
215. Ratner RE, Want LL, Fineman MS, et al. Adjunctive therapy with the amylin analogue pramlintide leads to a combined improvement in glycemic and weight control in insulin-treated subjects with type 2 diabetes. *Diabetes Technol Ther* 2002;4:51–61
216. Hollander PA, Levy P, Fineman MS, et al. Pramlinide as an adjunct to insulin therapy improves long-term glycemic and weight control in patients with type 2 diabetes: a 1-year randomized controlled trial. *Diabetes Care* 2003;26:784–790
217. Ratner RE, Dickey R, Fineman M, et al. Amylin replacement with pramlintide as an adjunct to insulin therapy improves long-term glycaemic and weight control in type 1 diabetes mellitus: a 1-year, randomized controlled trial. *Diabet Med* 2004;21:1204–1212
218. von Herrath M, Bain SC, Bode B, et al.; Anti-IL-21-liraglutide Study Group investigators and contributors. Anti-interleukin-21 antibody and liraglutide for the preservation of β -cell function in adults with recent-onset type 1 diabetes: a randomised, double-blind, placebo-controlled, phase 2 trial. *Lancet Diabetes Endocrinol* 2021;9:212–224
219. Nauck MA, Meier JJ. GLP-1 receptor agonists in type 1 diabetes: a MAG1C bullet? *Lancet Diabetes Endocrinol* 2020;8:262–264
220. Mathieu C, Zinman B, Hemmingsson JU, et al.; ADJUNCT ONE Investigators. Efficacy and safety of liraglutide added to insulin treatment in type 1 diabetes: the ADJUNCT ONE treat-to-target randomized trial. *Diabetes Care* 2016;39:1702–1710
221. Ahrén B, Hirsch IB, Pieber TR, et al.; ADJUNCT TWO Investigators. Efficacy and safety of liraglutide added to capped insulin treatment in subjects with type 1 diabetes: the ADJUNCT TWO randomized trial. *Diabetes Care* 2016;39:1693–1701
222. Shah VN, Akturk HK, Kruger D, et al. Semaglutide in adults with type 1 diabetes and obesity. *NEJM Evid* 2025;4:EVIDo2500173
223. Akturk HK, Dong F, Snell-Bergeon JK, Karakus KE, Shah VN. Efficacy and safety of tirzepatide in adults with type 1 diabetes: a proof of concept observational study. *J Diabetes Sci Technol* 2025;19:292–296
224. Snaith JR, Frampton R, Samocha-Bonet D, Greenfield JR. Tirzepatide in adults with type 1 diabetes: a phase 2 randomized placebo-controlled clinical trial. *Diabetes Care* 2026;49:161–170
225. Snaith JR, Holmes-Walker DJ, Greenfield JR. Reducing type 1 diabetes mortality: role for adjunctive therapies? *Trends Endocrinol Metab* 2020;31:150–164
226. Liu Y, Yang S, Jiang A, Zou D, Chen Z, Su N. Risk of diabetic ketoacidosis caused by sodium glucose cotransporter-2 inhibitors in patients with type 1 diabetes: a systematic review and network meta-analysis of randomized controlled trials. *Front Endocrinol (Lausanne)* 2024;15:1453067
227. Taylor SI, Blau JE, Rother KI, Beitelshes AL. SGLT2 inhibitors as adjunctive therapy for type 1 diabetes: balancing benefits and risks. *Lancet Diabetes Endocrinol* 2019;7:949–958
228. Danne T, Garg S, Peters AL, et al. International consensus on risk management of diabetic ketoacidosis in patients with type 1 diabetes treated with sodium-glucose cotransporter (SGLT) inhibitors. *Diabetes Care* 2019;42:1147–1154
229. McCoy RG, Herrin J, Galindo RJ, et al. Rates of hypoglycemic and hyperglycemic emergencies among U.S. adults with diabetes, 2011–2020. *Diabetes Care* 2023;46:e69–e71
230. Sherr JL, Laffel LM, Liu J, et al. Severe hypoglycemia and impaired awareness of hypoglycemia persist in people with type 1 diabetes despite use of diabetes technology: results from a cross-sectional survey. *Diabetes Care* 2024;47:941–947
231. Lin YK, Fisher SJ, Pop-Busui R. Hypoglycemia unawareness and autonomic dysfunction in diabetes: lessons learned and roles of diabetes technologies. *J Diabetes Investig* 2020;11:1388–1402
232. Geddes J, Schopman JE, Zammitt NN, Frier BM. Prevalence of impaired awareness of hypoglycaemia in adults with type 1 diabetes. *Diabet Med* 2008;25:501–504
233. Cryer PE. Mechanisms of hypoglycemia-associated autonomic failure in diabetes. *N Engl J Med* 2013;369:362–372
234. Clarke WL, Cox DJ, Gonder-Frederick LA, Julian D, Schlundt D, Polonsky W. Reduced awareness of hypoglycemia in adults with IDDM. A prospective study of hypoglycemic frequency and associated symptoms. *Diabetes Care* 1995;18:517–522
235. Pedersen-Bjergaard U, Pramming S, Thorsteinsson B. Recall of severe hypoglycaemia and self-estimated state of awareness in type 1 diabetes. *Diabetes Metab Res Rev* 2003;19:232–240
236. Gold AE, MacLeod KM, Frier BM. Frequency of severe hypoglycemia in patients with type I diabetes with impaired awareness of hypoglycemia. *Diabetes Care* 1994;17:697–703
237. Speight J, Barendse SM, Singh H, et al. Characterizing problematic hypoglycaemia: iterative design and preliminary psychometric validation of the Hypoglycaemia Awareness Questionnaire (HypoA-Q). *Diabet Med* 2016;33:376–385
238. Söholm U, Broadley M, Zaremba N, et al.; Hypo-RESOLVE consortium. The impact of hypoglycaemia on daily functioning among adults with diabetes: a prospective observational study using the Hypo-METRICS app. *Diabetologia* 2024;67:2160–2174
239. International Hypoglycaemia Study Group. Hypoglycaemia, cardiovascular disease, and mortality in diabetes: epidemiology, pathogenesis, and management. *Lancet Diabetes Endocrinol* 2019;7:385–396
240. Diabetes Control and Complications Trial/Epidemiology of Diabetes Interventions Complications Study Research Group. Long-term effect of diabetes and its treatment on cognitive function. *N Engl J Med* 2007;356:1842–1852
241. Yeoh E, Choudhary P, Nwokolo M, Ayis S, Amiel SA. Interventions that restore awareness of hypoglycemia in adults with type 1 diabetes: a systematic review and meta-analysis. *Diabetes Care* 2015;38:1592–1609
242. Renard E, Joubert M, Villard O, et al.; iDCL Trial Research Group. Safety and efficacy of sustained automated insulin delivery compared with sensor and pump therapy in adults with type 1 diabetes at high risk for hypoglycemia: a randomized controlled trial. *Diabetes Care* 2023;46:2180–2187
243. De Meulemeester J, Keymeulen B, De Block C, et al. One-year real-world benefits of Tandem Control-IQ technology on glucose management and person-reported outcomes in adults with type 1 diabetes: a prospective observational cohort study. *Diabetologia* 2025;68:948–960
244. Phillip M, Nimri R, Bergenstal RM, et al. Consensus recommendations for the use of automated insulin delivery technologies in clinical practice. *Endocr Rev* 2023;44:254–280
245. Tidemand KG, Laugesen C, Ranjan AG, Skovhus LB, Nørgaard K. Frequency of rebound hyperglycemia in adults with type 1 diabetes treated with different insulin delivery modalities. *Diabetes Technol Ther* 2025;27:60–65
246. Benoit SR, Hora I, Pasquel FJ, et al. Trends in emergency department visits and inpatient admissions for hyperglycemic crises in adults with diabetes in the U.S., 2006–2015. *Diabetes Care* 2020;43:1057–1064
247. Stougaard EB, Amadi H, Søndergaard E, et al. Time trends in the incidence of diabetic ketoacidosis leading to hospital admission among adults with type 1 diabetes: a nationwide Danish register study. *Diabetes Care* 2023;46:1897–1902
248. Kalscheuer H, Seufert J, Lanzinger S, et al.; DPV Initiative. Event rates and risk factors for the development of diabetic ketoacidosis in adult patients with type 1 diabetes: analysis from the DPV Registry based on 46,966 patients. *Diabetes Care* 2019;42:e34–e36
249. Everett E, Mathioudakis NN. Association of socioeconomic status and DKA readmission in adults with type 1 diabetes: analysis of the US National Readmission Database. *BMJ Open Diabetes Res Care* 2019;7:e000621
250. Garrett CJ, Moulton CD, Choudhary P, Amiel SA, Fonagy P, Ismail K. The psychopathology of recurrent diabetic ketoacidosis: a case-control study. *Diabet Med* 2021;38:e14505
251. Bonora BM, Avogaro A, Fadini GP. Sodium-glucose co-transporter-2 inhibitors and diabetic ketoacidosis: an updated review of the literature. *Diabetes Obes Metab* 2018;20:25–33
252. Jacobsen LM, Schatz D. Immunotherapy-based strategies for treatment of type 1 diabetes. *Horm Res Paediatr* 2025;98:425–434
253. Ramos EL, Dayan CM, Chatenoud L, et al.; PROTECT Study Investigators. Teplizumab and β -cell function in newly diagnosed type 1 diabetes. *N Engl J Med* 2023;389:2151–2161
254. Haller MJ, Long SA, Blanchfield JL, et al.; Type 1 Diabetes TrialNet ATG-GCSF Study Group.

- Low-dose anti-thymocyte globulin preserves C-peptide, reduces HbA1c, and increases regulatory to conventional T-cell ratios in new-onset type 1 diabetes: two-year clinical trial data. *Diabetes* 2019;68:1267–1276
255. Quattrin T, Haller MJ, Steck AK, et al.; T1GER Study Investigators. Golumumab and beta-cell function in youth with new-onset type 1 diabetes. *N Engl J Med* 2020;383:2007–2017
256. Waibel M, Wentworth JM, So M, et al.; BANDIT Study Group. Baricitinib and β -cell function in patients with new-onset type 1 diabetes. *N Engl J Med* 2023;389:2140–2150
257. Herold KC, Bundy BN, Long SA, et al.; Type 1 Diabetes TrialNet Study Group. An anti-CD3 antibody, teplizumab, in relatives at risk for type 1 diabetes. *N Engl J Med* 2019;381:603–613
258. Ziegler A-G, Cengiz E, Kay TWH. The future of type 1 diabetes therapy. *Lancet* 2025;406:1520–1534
259. Henriques FL, Buckle I, Forbes JM. Type 1 diabetes mellitus prevention: present and future. *Nat Rev Endocrinol* 2025;21:608–622
260. Hoffmann L, Kohls M, Arnolds S, et al.; EDENT1FI consortium. EDENT1FI Master Protocol for screening of presymptomatic early-stage type 1 diabetes in children and adolescents. *BMJ Open* 2025;15:e088522
261. McQueen RB, Geno Rasmussen C, Waugh K, et al. Cost and cost-effectiveness of large-scale screening for type 1 diabetes in Colorado. *Diabetes Care* 2020;43:1496–1503
262. Ziegler A-G, Kick K, Bonifacio E, et al.; Fr1da Study Group. Yield of a public health screening of children for islet autoantibodies in Bavaria, Germany. *JAMA* 2020;323:339–351
263. Bell KJ, Brodie S, Couper JJ, et al.; Type 1 Diabetes National Screening Pilot Study Group. Protocol for the Australian Type 1 Diabetes National Screening Pilot: assessing the feasibility and acceptability of three general population screening models in children. *Diabet Med* 2024;41:e15419
264. Gruessner AC, Gruessner RWG. The 2022 International Pancreas Transplant Registry Report—a review. *Transplant Proc* 2022;54:1918–1943
265. Rickels MR, Ballou CM, Foster NC, et al. Islet transplantation versus standard of care for type 1 diabetes complicated by severe hypoglycemia from the collaborative islet transplant registry and the T1D Exchange Registry. *Diabetes Care* 2025;48:737–744
266. Niclauss N, Morel P, Berney T. Has the gap between pancreas and islet transplantation closed? *Transplantation* 2014;98:593–599
267. Perrier Q, Jambon-Barbara C, Kessler L, et al.; GRAGIL Network. Impact of islet transplantation on diabetes complications and mortality in patients living with type 1 diabetes. *Diabetes Care* 2025;48:1007–1015
268. Finger EB, Matar AJ, Dunn TB, et al. Evolution of pancreas transplantation at a single institution-50+ years and 2500 transplants. *Ann Surg* 2024;280:604–615
269. Al-Naseem AO, Attia A, Gonnah AR, et al. Pancreas transplantation today: quo vadis? *Eur J Endocrinol* 2023;188:R73–R87
270. Speight J, Woodcock AJ, Reaney MD, et al. Well, I wouldn't be any worse off, would I, than I am now? A qualitative study of decision-making, hopes, and realities of adults with type 1 diabetes undergoing islet cell transplantation. *Transplant Direct* 2016;2:e72
271. Rech Tondin A, Lanzoni G. Islet cell replacement and regeneration for type 1 diabetes: current developments and future prospects. *BioDrugs* 2025;39:261–280
272. Lemos JRN, Skyler JS. Challenges in beta cell replacement for type 1 diabetes. *Horm Res Paediatr* 2025;98:435–449
273. Choksi H, Pleass H, Robertson P, Au E, Rogers N. Long-term metabolic outcomes post-simultaneous pancreas-kidney transplantation in recipients with type 1 diabetes. *Transplantation* 2025;109:1222–1229
274. Sollinger HW, Odorico JS, Becker YT, D'Alessandro AM, Pirsch JD. One thousand simultaneous pancreas-kidney transplants at a single center with 22-year follow-up. *Ann Surg* 2009;250:618–630
275. Kandaswamy R, Sutherland DER. Pancreas versus islet transplantation in diabetes mellitus: how to allocate deceased donor pancreata? *Transplant Proc* 2006;38:365–367
276. Robertson RP, Davis C, Larsen J, Stratta R, Sutherland DER, American Diabetes Association. Pancreas and islet transplantation in type 1 diabetes. *Diabetes Care* 2006;29:935
277. Choudhary P, Rickels MR, Senior PA, et al. Evidence-informed clinical practice recommendations for treatment of type 1 diabetes complicated by problematic hypoglycemia. *Diabetes Care* 2015;38:1016–1029
278. Senior PA, Bellin MD, Alejandro R, et al.; Clinical Islet Transplantation Consortium. Consistency of quantitative scores of hypoglycemia severity and glycemic lability and comparison with continuous glucose monitoring system measures in long-standing type 1 diabetes. *Diabetes Technol Ther* 2015;17:235–242
279. Ryan EA, Shandro T, Green K, et al. Assessment of the severity of hypoglycemia and glycemic lability in type 1 diabetic subjects undergoing islet transplantation. *Diabetes* 2004;53:955–962
280. Barton FB, Rickels MR, Alejandro R, et al. Improvement in outcomes of clinical islet transplantation: 1999–2010. *Diabetes Care* 2012;35:1436–1445
281. Marfil-Garza BA, Imes S, Verhoeff K, et al. Pancreatic islet transplantation in type 1 diabetes: 20-year experience from a single-centre cohort in Canada. *Lancet Diabetes Endocrinol* 2022;10:519–532
282. Catarinella D, Melzi R, Mercalli A, et al. Long-term outcomes of pancreatic islet transplantation alone in type 1 diabetes: a 20-year single-centre study in Italy. *Lancet Diabetes Endocrinol* 2025;13:279–293
283. O'Connell PJ, Holmes-Walker DJ, Goodman D, et al.; Australian Islet Transplant Consortium. Multicenter Australian trial of islet transplantation: improving accessibility and outcomes. *Am J Transplant* 2013;13:1850–1858
284. Lablanche S, Vantuyghem M-C, Kessler L, et al.; TRIMECO trial investigators. Islet transplantation versus insulin therapy in patients with type 1 diabetes with severe hypoglycaemia or poorly controlled glycaemia after kidney transplantation (TRIMECO): a multicentre, randomised controlled trial. *Lancet Diabetes Endocrinol* 2018;6:527–537
285. Bond Z, Malik S, Bashir A, et al. Validation of IgGs criteria for islet transplant functional status using person-reported outcome measures in a cross-sectional study. *Transpl Int* 2023;36:11659
286. Wang Q, Huang Y-X, Liu L, et al. Pancreatic islet transplantation: current advances and challenges. *Front Immunol* 2024;15:1391504
287. Reichman TW, Markmann JF, Odorico J, et al.; VX-880-101 FORWARD Study Group. Stem cell-derived, fully differentiated islets for type 1 diabetes. *N Engl J Med* 2025;393:858–868
288. Nair GG, Tzanakakis ES, Hebrok M. Emerging routes to the generation of functional β -cells for diabetes mellitus cell therapy. *Nat Rev Endocrinol* 2020;16:506–518
289. Montgomery RA, Stern JM, Lonze BE, et al. Results of two cases of pig-to-human kidney xenotransplantation. *N Engl J Med* 2022;386:1889–1898
290. Wang S, Du Y, Zhang B, et al. Transplantation of chemically induced pluripotent stem-cell-derived islets under abdominal anterior rectus sheath in a type 1 diabetes patient. *Cell* 2024;187:6152–6164
291. Hoglebe NJ, Ishahak M, Millman JR. Developments in stem cell-derived islet replacement therapy for treating type 1 diabetes. *Cell Stem Cell* 2023;30:530–548
292. Kioulaphides S, García AJ. Encapsulation and immune protection for type 1 diabetes cell therapy. *Adv Drug Deliv Rev* 2024;207:115205
293. Gerace D, Zhou Q, Kenty JH-R, et al. Engineering human stem cell-derived islets to evade immune rejection and promote localized immune tolerance. *Cell Rep Med* 2023;4:100879
294. Krentz NAJ, Shea LD, Huisling MO, Shaw JAM. Restoring normal islet mass and function in type 1 diabetes through regenerative medicine and tissue engineering. *Lancet Diabetes Endocrinol* 2021;9:708–724
295. Vasavada RC, Dhawan S. Harnessing beta-cell replication: advancing molecular insights to regenerative therapies in diabetes. *Front Endocrinol (Lausanne)* 2025;16:1612576
296. Basile G, Qadir MMF, Mauvais-Jarvis F, et al. Emerging diabetes therapies: bringing back the β -cells. *Mol Metab* 2022;60:101477
297. Weinstock RS, Schütz-Fuhrmann I, Connor CG, et al.; DPV Initiative. Type 1 diabetes in older adults: comparing treatments and chronic complications in the United States T1D Exchange and the German/Austrian DPV registries. *Diabetes Res Clin Pract* 2016;122:28–37
298. Roy MS, Klein R, O'Colmain BJ, Klein BEK, Moss SE, Kempen JH. The prevalence of diabetic retinopathy among adult type 1 diabetic persons in the United States. *Arch Ophthalmol* 2004;122:546–551
299. Silva PS, Horton MB, Clary D, et al. Identification of diabetic retinopathy and ungradable image rate with ultrawide field imaging in a national teleophthalmology program. *Ophthalmology* 2016;123:1360–1367
300. Walton OB, Garoon RB, Weng CY, et al. Evaluation of automated teleretinal screening program for diabetic retinopathy. *JAMA Ophthalmol* 2016;134:204–209
301. Aspelund T, Thörnórisdóttir O, Ólafsdóttir E, et al. Individual risk assessment and information technology to optimise screening frequency for diabetic retinopathy. *Diabetologia* 2011;54:2525–2532
302. Chen K-Y, Chan H-C, Chan C-M. Can fibrin therapy redefine the management of diabetic retinopathy? A comprehensive systematic review

- and meta-analysis of efficacy and safety. *J Diabetes Complications* 2025;39:109178
303. Heerspink HJ, Cherney DZ, Groop P-H, et al. People with type 1 diabetes and chronic kidney disease urgently need new therapies: a call for action. *Lancet Diabetes Endocrinol* 2023;11:536–540
304. Helve J, Sund R, Arffman M, et al. Incidence of end-stage renal disease in patients with type 1 diabetes. *Diabetes Care* 2018;41:434–439
305. Tuttle KR, Reynolds CL, Kornowske LM, et al.; CURE-CKD Consortium. Prevalence and severity of chronic kidney disease in a population with type 1 diabetes from a United States health system: a real-world cohort study. *Lancet Reg Health Am* 2025;47:101130
306. Jansson Sigfrids F, Groop P-H. Progression and regression of kidney disease in type 1 diabetes. *Front Nephrol* 2023;3:1282818
307. Gheith O, Farouk N, Nampoori N, Halim MA, Al-Otaibi T. Diabetic kidney disease: world wide difference of prevalence and risk factors. *J Nephropharmacol* 2016;5:49–56
308. de Boer IH, Khunti K, Sadusky T, et al. Diabetes management in chronic kidney disease: a consensus report by the American Diabetes Association (ADA) and Kidney Disease: Improving Global Outcomes (KDIGO). *Diabetes Care* 2022;45:3075–3090
309. Heerspink HJL, Birkenfeld AL, Cherney DZI, et al.; FINE-ONE Investigators. Finerenone in type 1 diabetes and chronic kidney disease. *N Engl J Med* 2026;394:947–957
310. Kidney Disease: Improving Global Outcomes (KDIGO) CKD Work Group. KDIGO 2024 Clinical Practice Guideline for the Evaluation and Management of Chronic Kidney Disease. *Kidney Int* 105:S117–S314
311. Akerkar A, Rønn PF, Kosjerina V, et al. Cumulative risk of diabetic foot complications in risk groups of type 1 and type 2 diabetes: real-world evidence from a 22-year follow-up study. *Diabetes Obes Metab* 2025;27:2284–2287
312. Schaper NC, van Netten JJ, Apelqvist J, et al.; IWGDF Editorial Board. Practical guidelines on the prevention and management of diabetes-related foot disease (IWGDF 2023 update). *Diabetes Metab Res Rev* 2024;40:e3657
313. Freeman R. Not all neuropathy in diabetes is of diabetic etiology: differential diagnosis of diabetic neuropathy. *Curr Diab Rep* 2009;9:423–431
314. Serhiyenko VA, Serhiyenko AA. Cardiac autonomic neuropathy: risk factors, diagnosis and treatment. *World J Diabetes* 2018;9:1–24
315. Agashe S, Petak S. Cardiac autonomic neuropathy in diabetes mellitus. *Methodist DeBakey Cardiovasc J* 2018;14:251–256
316. Manrique-Acevedo C, Hirsch IB, Eckel RH. Prevention of cardiovascular disease in type 1 diabetes. *N Engl J Med* 2024;390:1207–1217
317. Nathan DM, Cleary PA, Backlund J-YC, et al.; Diabetes Control and Complications Trial/Epidemiology of Diabetes Interventions and Complications (DCCT/EDIC) Study Research Group. Intensive diabetes treatment and cardiovascular disease in patients with type 1 diabetes. *N Engl J Med* 2005;353:2643–2653
318. Lewis CE, Fine LJ, Beddhu S, et al.; Sprint Research Group. Final report of a trial of intensive versus standard blood-pressure control. *N Engl J Med* 2021;384:1921–1930
319. Bi Y, Li M, Liu Y, et al.; BROAD Research Group. Intensive blood-pressure control in patients with type 2 diabetes. *N Engl J Med* 2025;392:1155–1167
320. Hero C, Rawshani A, Svensson A-M, et al. Association between use of lipid-lowering therapy and cardiovascular diseases and death in individuals with type 1 diabetes. *Diabetes Care* 2016;39:996–1003
321. Vistisen D, Andersen GS, Hansen CS, et al. Prediction of first cardiovascular disease event in type 1 diabetes mellitus: the Steno Type 1 Risk Engine. *Circulation* 2016;133:1058–1066
322. Hippisley-Cox J, Coupland C, Brindle P. Development and validation of QRISK3 risk prediction algorithms to estimate future risk of cardiovascular disease: prospective cohort study. *BMJ* 2017;357:j2099
323. Puri R, Nicholls SJ, Shao M, et al. Impact of statins on serial coronary calcification during atheroma progression and regression. *J Am Coll Cardiol* 2015;65:1273–1282
324. Haji M, Erqou S, Fonarow GC, Echouffo-Tcheugui JB. Type 1 diabetes and risk of heart failure: a systematic review and meta-analysis. *Diabetes Res Clin Pract* 2023;202:110805
325. Fang M, Jeon Y, Echouffo-Tcheugui JB, Selvin E. Prevalence and management of obesity in U.S. adults with type 1 diabetes. *Ann Intern Med* 2023;176:427–429
326. Welters A, Tittel SR, Laubner K, et al. Long-term trends of BMI and cardiometabolic risk factors among adults with type 1 diabetes: an observational study from the German/Austrian DPV registry. *Diabetes Res Clin Pract* 2021;178:108973
327. Van der Schueren B, Ellis D, Faradji RN, Al-Ozairi E, Rosen J, Mathieu C. Obesity in people living with type 1 diabetes. *Lancet Diabetes Endocrinol* 2021;9:776–785
328. Igudesman D, Crandell J, Corbin KD, et al.; ACTION Study Group. Weight management in young adults with type 1 diabetes: the advancing care for type 1 diabetes and obesity network sequential multiple assignment randomized trial pilot results. *Diabetes Obes Metab* 2023;25:688–699
329. Turton JL, Raab R, Rooney KB. Low-carbohydrate diets for type 1 diabetes mellitus: a systematic review. *PLoS One* 2018;13:e0194987
330. Ranjan A, Schmidt S, Damm-Frydenberg C, et al. Low-carbohydrate diet impairs the effect of glucagon in the treatment of insulin-induced mild hypoglycemia: a randomized crossover study. *Diabetes Care* 2017;40:132–135
331. Wu N, Bredin SSD, Guan Y, et al. Cardiovascular health benefits of exercise training in persons living with type 1 diabetes: a systematic review and meta-analysis. *J Clin Med* 2019;8:253
332. Gudzone KA, Kushner RF. Medications for obesity: a review. *JAMA* 2024;332:571–584
333. Garg SK, Kaur G, Haider Z, Rodriguez E, Beatson C, Snell-Bergeon J. Efficacy of semaglutide in overweight and obese patients with type 1 diabetes. *Diabetes Technol Ther* 2024;26:184–189
334. Karakus KE, Klein MP, Akturk HK, Shah VN. Changes in basal and bolus insulin requirements with tirzepatide as an adjunctive therapy in adults with type 1 diabetes using Tandem Control-IQ. *Diabetes Ther* 2024;15:1647–1655
335. Brethauer SA, Aminian A, Rosenthal RJ, Kirwan JP, Kashyap SR, Schauer PR. Bariatric surgery improves the metabolic profile of morbidly obese patients with type 1 diabetes. *Diabetes Care* 2014;37:e51–e52
336. Kermansaravi M, Valizadeh R, Jazi AD, et al. Current status of metabolic/bariatric surgery in type 1 diabetes mellitus: an updated systematic review and meta-analysis. *Obes Surg* 2022;32:1726–1733
337. Lannoo M, Dillemans B, Van Nieuwenhove Y, et al. Bariatric surgery induces weight loss but does not improve glycemic control in patients with type 1 diabetes. *Diabetes Care* 2014;37:e173–e174
338. Carette C, Rives-Lange C, Shoung N, et al. Lights and shadows of bariatric surgery: insights from a nationwide administrative database of people living with type 1 diabetes and obesity. *Diabetes Ther* 2025;16:1267–1277
339. Tomic D, Harding JL, Jenkins AJ, Shaw JE, Magliano DJ. The epidemiology of type 1 diabetes mellitus in older adults. *Nat Rev Endocrinol* 2025;21:92–104
340. Yang K, Yang X, Jin C, et al. Global burden of type 1 diabetes in adults aged 65 years and older, 1990–2019: population based study. *BMJ* 2024;385:e078432
341. Miller KM, Kanapka LG, Rickels MR, et al. Benefit of continuous glucose monitoring in reducing hypoglycemia is sustained through 12 months of use among older adults with type 1 diabetes. *Diabetes Technol Ther* 2022;24:424–434
342. Munshi MN, Slynne C, Adam A, et al. Continuous glucose monitoring with geriatric principles in older adults with type 1 diabetes and hypoglycemia: a randomized controlled trial. *Diabetes Care* 2025;48:694–702
343. Weinstock RS, Ragghinaru D, Gal RL, et al. Older adults benefit from virtual support for continuous glucose monitor use but require longer visits. *J Diabetes Sci Technol* 2026;20:994–1002
344. Allen NA, Grigorian EG, Mansfield K, Berg CA, Litchman ML. Continuous glucose monitoring with data sharing in older adults: a qualitative study. *J Clin Nurs* 2023;32:7483–7494
345. Kudva YC, Henderson RJ, Kanapka LG, et al. Automated insulin delivery in older adults with type 1 diabetes. *NEJM Evid* 2025;4:EVID02400200
346. Kudva YC, Henderson RJ, Kanapka LG, et al.; AIDE Study Group. Automated insulin delivery in elderly with type 1 diabetes: a prespecified analysis of the extension phase. *Diabetes Technol Ther* 2025;27:572–575
347. McAuley SA, Trawley S, Vogrin S, et al. Closed-Loop Insulin Delivery Versus Sensor-Augmented Pump Therapy in Older Adults With Type 1 Diabetes (ORACL): a randomized, crossover trial. *Diabetes Care* 2022;45:381–390
348. Boughton CK, Hartnell S, Thabit H, et al. Hybrid closed-loop glucose control compared with sensor augmented pump therapy in older adults with type 1 diabetes: an open-label multicentre, multinational, randomised, crossover study. *Lancet Healthy Longev* 2022;3:e135–e142
349. Jensen DM, Korsholm L, Ovesen P, et al. Peri-conceptual A1C and risk of serious adverse pregnancy outcome in 933 women with type 1 diabetes. *Diabetes Care* 2009;32:1046–1048
350. Abell SK, Boyle JA, de Courten B, et al. Contemporary type 1 diabetes pregnancy outcomes: impact of obesity and glycaemic control. *Med J Aust* 2016;205:162–167

351. Feig DS, Donovan LE, Corcoy R, et al.; CONCEPTT Collaborative Group. Continuous glucose monitoring in pregnant women with type 1 diabetes (CONCEPTT): a multicentre international randomised controlled trial. *Lancet* 2017;390:2347–2359
352. Sanusi AA, Xue Y, McIlwraith C, et al. Association of continuous glucose monitoring metrics with pregnancy outcomes in patients with preexisting diabetes. *Diabetes Care* 2024;47:89–96
353. Battelino T, Danne T, Bergenstal RM, et al. Clinical targets for continuous glucose monitoring data interpretation: recommendations from the International Consensus on Time in Range. *Diabetes Care* 2019;42:1593–1603
354. Nielsen LR, Pedersen-Bjergaard U, Thorsteinsson B, Johansen M, Damm P, Mathiesen ER. Hypoglycemia in pregnant women with type 1 diabetes: predictors and role of metabolic control. *Diabetes Care* 2008;31:9–14
355. Sibai BM, Viteri OA. Diabetic ketoacidosis in pregnancy. *Obstet Gynecol* 2014;123:167–178
356. Teixeira T, Godoi A, Romeiro P, et al. Efficacy of automated insulin delivery in pregnant women with type 1 diabetes: a meta-analysis and trial sequential analysis of randomized controlled trials. *Acta Diabetol* 2024;61:831–840
357. Stewart ZA, Wilinska ME, Hartnell S, et al. Closed-loop insulin delivery during pregnancy in women with type 1 diabetes. *N Engl J Med* 2016;375:644–654
358. Stewart ZA, Wilinska ME, Hartnell S, et al. Day-and-night closed-loop insulin delivery in a broad population of pregnant women with type 1 diabetes: a randomized controlled crossover trial. *Diabetes Care* 2018;41:1391–1399
359. Lee TTM, Collett C, Bergford S, et al.; AiDAPT Collaborative Group. Automated insulin delivery in women with pregnancy complicated by type 1 diabetes. *N Engl J Med* 2023;389:1566–1578
360. Benhalima K, Beunen K, Van Wilder N, et al. Comparing advanced hybrid closed loop therapy and standard insulin therapy in pregnant women with type 1 diabetes (CRISTAL): a parallel-group, open-label, randomised controlled trial. *Lancet Diabetes Endocrinol* 2024;12:390–403
361. Lawton J, Kimbell B, Closs M, et al. Listening to women: experiences of using closed-loop in type 1 diabetes pregnancy. *Diabetes Technol Ther* 2023;25:845–855
362. Benhalima K, Polsky S. Automated insulin delivery in pregnancies complicated by type 1 diabetes. *J Diabetes Sci Technol* 2026;20:38–49
363. Wyckoff JA, Lapolla A, Asias-Dinh BD, et al. Preexisting diabetes and pregnancy: an Endocrine Society and European Society of Endocrinology joint clinical practice guideline. *J Clin Endocrinol Metab* 2025;110:2405–2452
364. National Institute for Health and Care Excellence. (2023) Hybrid closed loop systems for managing blood glucose levels in type 1 diabetes. TA943. Accessed 2 June 2026. Available from <https://www.nice.org.uk/guidance/ta943>
365. NHS Digital. (2026) National Pregnancy in Diabetes Audit - Hybrid closed loop 2025 (01 January 2025 to 31 December 2025). Accessed 14 July 2026. Available from <https://digital.nhs.uk/data-and-information/publications/statistical/national-pregnancy-in-diabetes-audit/2025#highlights>
366. Roeder HA, Moore TR, Ramos GA. Changes in postpartum insulin requirements for patients with well-controlled type 1 diabetes. *Am J Perinatol* 2016;33:683–687
367. Davies HA, Clark JD, Dalton KJ, Edwards OM. Insulin requirements of diabetic women who breast feed. *BMJ* 1989;298:1357–1358
368. Visser MM, Vangoitsenhoven R, Gillard P, Mathieu C. Review article—diabetes technology in the hospital: an update. *Curr Diab Rep* 2024;24:173–182
369. Thabit H, Schofield J. Technology in the management of diabetes in hospitalised adults. *Diabetologia* 2024;67:2114–2128
370. Wright JJ, Williams AJ, Friedman SB, et al. Accuracy of continuous glucose monitors for inpatient diabetes management. *J Diabetes Sci Technol* 2023;17:1252–1255
371. Davis GM, Spanakis EK, Migdal AL, et al. Accuracy of Dexcom G6 continuous glucose monitoring in non-critically ill hospitalized patients with diabetes. *Diabetes Care* 2021;44:1641–1646
372. Villard O, Breton MD, Rao S, et al. Accuracy of a factory-calibrated continuous glucose monitor in individuals with diabetes on hemodialysis. *Diabetes Care* 2022;45:1666–1669
373. Voglová Hagerf B, Protus M, Nemetova L, et al. Accuracy and feasibility of real-time continuous glucose monitoring in critically ill patients after abdominal surgery and solid organ transplantation. *Diabetes Care* 2024;47:956–963
374. Cavalcante Lima Chagas G, Teixeira L, R C Clemente M, et al. Use of continuous glucose monitoring and point-of-care glucose testing in hospitalized patients with diabetes mellitus in non-intensive care unit settings: a systematic review and meta-analysis of randomized controlled trials. *Diabetes Res Clin Pract* 2025;220:111986
375. Olsen MT, Jensen SH, Rasmussen LM, et al. Most hospitalised patients with type 2 diabetes benefit from continuous glucose monitoring compared to point-of-care glucose testing in a non-intensive care unit setting: a heterogeneity of treatment effect analysis. *Diabetes Obes Metab* 2025;27:2857–2863
376. Nielsen CG, Grigonyte-Daraskeviciene M, Olsen MT, et al. Accuracy of continuous glucose monitoring systems in intensive care unit patients: a scoping review. *Intensive Care Med* 2024;50:2005–2018
377. Davis GM, Hughes MS, Brown SA, et al. Automated insulin delivery with remote real-time continuous glucose monitoring for hospitalized patients with diabetes: a multicenter, single-arm, feasibility trial. *Diabetes Technol Ther* 2023;25:677–688
378. Lee MY, Seav SM, Ongwela L, et al. Empowering hospitalized patients with diabetes: implementation of a hospital-wide CGM policy with EHR-integrated validation for dosing insulin. *Diabetes Care* 2024;47:1838–1845
379. Galindo RJ, Umpierrez GE, Rushakoff RJ, et al. Continuous glucose monitors and automated insulin dosing systems in the hospital consensus guideline. *J Diabetes Sci Technol* 2020;14:1035–1064
380. Bailon RM, Partlow BJ, Miller-Cage V, et al. Continuous subcutaneous insulin infusion (insulin pump) therapy can be safely used in the hospital in select patients. *Endocr Pract* 2009;15:24–29
381. Cook CB, Boyle ME, Cisar NS, et al. Use of continuous subcutaneous insulin infusion (insulin pump) therapy in the hospital setting: proposed guidelines and outcome measures. *Diabetes Educ* 2005;31:849–857
382. Milluzzo A, Falorni A, Brozzetti A, et al. Risk for coexistent autoimmune diseases in familial and sporadic type 1 diabetes is related to age at diabetes onset. *Endocr Pract* 2021;27:110–117