Research: Epidemiology

Continuing rise of Type 2 diabetes incidence in children and young people in the UK

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Abstract

Aims To estimate the incidence of Type 2 diabetes in children aged <17 years, compare this with similar data 10 years ago, and characterize clinical features at diagnosis in the UK and Republic of Ireland.

Methods Using the British Paediatric Surveillance Unit reporting framework, cases of Type 2 diabetes diagnosed in children aged <17 years between 1 April 2015 and 30 April 2016 were reported each month.

Results A total of 106 cases were reported, giving a UK incidence of 0.72/100 000 (95% CI 0.58–0.88). Children from ethnic minorities had significantly higher incidence compared with white children (0.44/100 000) with rates of 2.92/100 000 and 1.67/100 000, in Asian and BACBB (black/African/Caribbean/black British) children respectively. Sixty-seven percent were girls and 81% had a family history of Type 2 diabetes. The mean BMI SD score at diagnosis was 2.89 (2.88, girls; 2.92, boys); 81% were obese. Children of Asian ethnicity had a significantly lower BMI SD score compared with white children (P < 0.001). There was a trend in increased incidence from 2005 to 2015, with a rate ratio of 1.35 (95% CI 0.99–1.84), although this was not statistically significant (P = 0.062). There was statistical evidence of increased incidence among girls (P = 0.03) and children of South-Asian ethnicity (P = 0.01) when comparing the 2005 and 2015 surveys.

Conclusions Type 2 diabetes remains far less common than Type 1 diabetes in childhood in the UK, but the number of cases continues to rise, with significantly increased incidence among girls and South-Asian children over a decade. Female gender, family history, non-white ethnicity and obesity were found to be strongly associated with the condition.


Introduction

Type 2 diabetes accounts for 90% of diabetes globally [1] and prevalence of Type 2 diabetes in adults is increasing in the UK [2]. The highest rates of Type 2 diabetes in children and adolescents have been reported in studies from the USA [3], with the incidence rising from nine to 12.5 cases per 100 000 between 2002 and 2012 [4]. European countries report a lower incidence, with an Austrian study reporting 0.29 per 100 000 per year between 1999 and 2007 [3]. Recent results from a multinational European database of >27 000 children found that Type 2 diabetes accounted for 1.3% of all cases of diabetes. In Europe, the UK has the highest reported prevalence of childhood Type 2 diabetes [5]. In 2005, the UK annual incidence of Type 2 diabetes in children aged <17 years was 0.53 per 100 000 per year [6]. Among children in the UK, Type 2 diabetes is less common than Type 1 diabetes. The National Paediatric Diabetes Audit 2015–2016 reported that Type 2 diabetes accounted for 2.2% of the total [7].

Risk factors for developing childhood Type 2 diabetes are similar to those for adulthood, and include obesity, family history and ethnicity [8]. A previous UK study reported 95% of children were overweight or obese at diagnosis [6], with similar findings from other countries [9]. In the UK, incidence was substantially higher in ethnic minority groups: 3.9 per 100 000 in BACBB children, and 1.25 per 100 000 in children of South-Asian ethnicity, compared with 0.35 per 100 000 in white children [6], with similar findings among non-white ethnicities in other studies [3]. Family history is an
What’s new?

- The 2015/2016 UK incidence of Type 2 diabetes in children aged <17 years was 0.72 per 100 000 per year.
- The incidence of Type 2 diabetes amongst girls and South-Asian children has risen significantly over the last decade.
- Female gender, family history, non-white ethnicity and obesity were strongly associated with Type 2 diabetes in childhood.
- Comorbidities are commonly identified at diagnosis, including 37% with non-alcoholic fatty liver disease and 21% with hypertension.
- The most common presenting complaint at diagnosis after osmotic symptoms (polyuria, polydipsia and weight loss) was recurrent, mainly genital, infections, although over a third of cases were asymptomatic and detected on obesity screening investigations.

Diabetes mellitus was defined according to American Diabetes Association [12] definition with fasting glucose level >7 mmol/l, random glucose level >11.1 mmol/l or stimulated glucose level >11.1 mmol/l after a standard oral glucose tolerance test or HbA1c >48 mmol/mol (6.5%). Type 2 diabetes was distinguished from other types of diabetes using the following criteria: (1) presence of raised insulin level (>132 pmol/l) or raised C peptide level (>0.6 mmol/l) [13,14] or (2) the child was managed off insulin therapy for >9 months in the absence of typical Type 1 diabetes auto-antibodies. The latter definition is likely to be based on a clinical re-evaluation after diagnosis when a clinical course suggests a diagnosis other than Type 1 diabetes. Exclusion criteria included: Type 1 diabetes (positive auto-antibodies and/or persisting insulin requirement from diagnosis); maturity-onset diabetes of the young; diabetes developing in a person with a known diabetes-associated syndrome, such as Prader–Willi or Bardet–Biedl syndromes; diagnosis of diabetes while on medical therapy with a known diabetogenic medication and pancreatic failure.

The initial study proforma collected information about the child, clinical presentation, diagnostic details and management. Clinicians were asked to provide evidence of comorbidities including hypertension (defined as systolic blood pressure above the 95th centile for sex, age and height centile [15]), renal disease (presence of macro/microalbuminuria), polycystic ovarian syndrome (evidenced by oligo or secondary amennorhoea, ultrasonography or a biochemical picture of luteinizing hormone:follicle-stimulating hormone ratio >3, low sex hormone binding globulin) and non-alcoholic fatty liver disease (alanine aminotransferase >50 mg/dl for boys and >44 mg/dl for girls in obese children or ultrasonography indicating hepatic steatosis with other aetiologies excluded [16]). A section allowed clinicians to report other relevant clinical features. A follow-up proforma was sent 1 year after diagnosis. Case eligibility was determined on review of the questionnaire by the investigators (T.C., J.H.S and T.B.).

The methodology was the same used with the BPSU study in 2004–2005 [6] to allow direct comparison.

Methods

A prospective monthly surveillance of >3400 consultant paediatricians in the UK and Republic of Ireland using the British Paediatric Surveillance Unit (BPSU), based at the Royal College of Paediatrics and Child Health, UK was undertaken to identify new cases of Type 2 diabetes in children aged <17 years. The BPSU is an active reporting system in which an orange report card containing a list of conditions is sent monthly, by post or electronically, to all consultant paediatricians. Respondents report cases they have seen in the previous month for conditions named on the card or tick ‘Nothing to report’. The BPSU forward the reporting clinician’s details to the research team, who send a proforma requesting the child’s clinical details. The study ran for 13 months (as per previous BPSU studies allowing a lead-in month) from April 2015 to April 2016.

-Focused on early-onset Type 2 diabetes in young people.
-Fewer new cases of Type 2 diabetes were detected compared with Type 1 diabetes.
-Female gender, non-white ethnicity and obesity were strongly associated with Type 2 diabetes.
-Comorbidities are commonly identified at diagnosis, including 37% with non-alcoholic fatty liver disease and 21% with hypertension.
-The most common presenting complaint at diagnosis after osmotic symptoms was recurrent, mainly genital, infections, although over a third of cases were asymptomatic and detected on obesity screening investigations.
UK, Mid-2015’ table, published by the Office for National Statistics (www.ons.gov.uk). Ethnicity-specific incidence rates were calculated for English and Welsh children only using the ‘Ethnic Group by Age in England and Wales 2011’ data (www.ons.gov.uk). Data for ethnicity by age were only available for these two countries. The 95% CIs for incidence rates were calculated using the exact Poisson method [17]. Poisson regression models were used to compare the incidences rates in the different ethnic groups. Each ethnic group was compared with the white ethnicity, baseline group, and the P values from the Wald test are reported in Table 1. BMI z-scores (SD scores) at diagnosis were calculated from weight and height, age and gender using the 1990 UK growth standard curves [18]. For each ethnic group, we reported the mean difference in BMI SD score from the white ethnic group, as the baseline group, 95% CI, and the Wald test P value (Table 2). Overweight and obesity were defined as described in Cole et al. [19] as follows: SD scores ≥1.30 and ≥2.37, for boys and ≥1.19 and ≥2.25 for girls, respectively. Simple linear regression models were used to compare the BMI SD scores according to ethnic group. Because the BMI SD scores are adjusted, by definition, for age and gender, the resulting estimates consider any potential confounding effects of these variables when comparing the BMI across the ethnic groups.

Overall UK incidence, gender-specific and ethnic-specific incidence rates for 2015 were compared with the corresponding rates for 2005 using incidence rate ratio, defined as the ratio of incidence in 2015 to 2005 (with null value equal to 1). P values computed from the z-test for log-ratio of incidence rates are reported in Table 3. For comparisons of ethnic group-specific rates, findings were based on the England figures only, to be comparable with 2005 results.

R Statistical software (version 3.3.2) were used for the analysis. The descriptive statistics included all 106 cases (UK and Republic of Ireland).

**Results**

A total of 155 case notifications were received during the 13-month study period. The mean monthly BPSU card return compliance rate was 92%. For the 155 notifications, 148 questionnaires (95%) were received (Fig. 1), 29 cases were excluded (eight were diagnosed outside the study period, in seven cases another type of diabetes was diagnosed, in five cases the individual was aged >17 years at diagnosis, and nine cases did not meet the criteria for diabetes or the diabetes was associated with known associated syndrome/medical treatment, or was made outside the UK/Republic of Ireland). After case review, 119 cases met the diagnostic criteria for Type 2 diabetes. Thirteen of these were excluded, as duplicate cases (two clinicians reporting cases with the same, unique National Health Service identification number). A total of 106 cases were identified as newly confirmed (104 from the UK, two from the Republic of Ireland).

In all, 67% of cases were in girls (71/106). The median (range) age at diagnosis was 14.3 (7.9–16.9) years [girls: 14.2 (7.9–16.9) years; boys 14.5 (9.9–16.7) years], 44% of the cases the children were white, in 4% of cases the ethnic group was uncertain and 52% of the cases were in ethnic minority groups, predominantly Asian/Asian-British (65%) and BACBB (25%). Of the total cohort, 34% were Asian or Asian-British (36 cases) and 13% were BACBB (14 cases).

**Table 1** Incidence rates for Type 2 diabetes in children aged 0–16 years in England and Wales, according to ethnic group

<table>
<thead>
<tr>
<th>Group</th>
<th>Ethnicity</th>
<th>Population</th>
<th>No. cases</th>
<th>Incidence rate*†</th>
<th>95% CI</th>
<th>P value (Wald test)</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>All Ethnicities</td>
<td>11 274 750</td>
<td>88</td>
<td>0.78</td>
<td>0.63–0.96</td>
<td>Baseline group</td>
</tr>
<tr>
<td></td>
<td>White: total</td>
<td>8 921 552</td>
<td>39</td>
<td>0.44</td>
<td>0.31–0.60</td>
<td></td>
</tr>
<tr>
<td>2</td>
<td>Mixed: total</td>
<td>577 065</td>
<td>&lt;5</td>
<td>0.52</td>
<td>0.11–1.32</td>
<td>0.772</td>
</tr>
<tr>
<td></td>
<td>White and black</td>
<td>84 471</td>
<td>&lt;5</td>
<td>2.37</td>
<td>0.29–8.55</td>
<td>0.020</td>
</tr>
<tr>
<td></td>
<td>Other mixed</td>
<td>124 917</td>
<td>&lt;5</td>
<td>0.80</td>
<td>0.02–4.46</td>
<td>0.550</td>
</tr>
<tr>
<td>3</td>
<td>Asian: total</td>
<td>1 096 304</td>
<td>32</td>
<td>2.92</td>
<td>2.00–4.12</td>
<td>2×10⁻¹³</td>
</tr>
<tr>
<td></td>
<td>Indian</td>
<td>287 027</td>
<td>&lt;5</td>
<td>1.05</td>
<td>0.22–3.05</td>
<td>0.146</td>
</tr>
<tr>
<td></td>
<td>Pakistani</td>
<td>388 889</td>
<td>15</td>
<td>3.86</td>
<td>2.16–6.36</td>
<td>8×10⁻¹¹</td>
</tr>
<tr>
<td></td>
<td>Bangladeshi</td>
<td>163 308</td>
<td>8</td>
<td>4.90</td>
<td>2.11–9.65</td>
<td>5×10⁻⁸</td>
</tr>
<tr>
<td></td>
<td>Other Asian</td>
<td>201 002</td>
<td>6</td>
<td>2.99</td>
<td>1.10–6.50</td>
<td>1×10⁻⁵</td>
</tr>
<tr>
<td>4</td>
<td>BACBB: total</td>
<td>537 938</td>
<td>9</td>
<td>1.67</td>
<td>0.77–3.18</td>
<td>3×10⁻⁴</td>
</tr>
<tr>
<td></td>
<td>Caribbean</td>
<td>111 569</td>
<td>&lt;5</td>
<td>1.79</td>
<td>0.22–6.48</td>
<td>0.052</td>
</tr>
<tr>
<td></td>
<td>African</td>
<td>314 374</td>
<td>5</td>
<td>1.59</td>
<td>0.52–3.71</td>
<td>0.007</td>
</tr>
<tr>
<td></td>
<td>Other Black</td>
<td>111 996</td>
<td>&lt;5</td>
<td>1.79</td>
<td>0.22–6.45</td>
<td>0.052</td>
</tr>
<tr>
<td>5</td>
<td>Other: total</td>
<td>141 891</td>
<td>&lt;5</td>
<td>0.70</td>
<td>0.02–3.93</td>
<td>0.637</td>
</tr>
</tbody>
</table>

* Rates are presented per 100 000/year.
† Poisson regression model was used to compare the incidence rates of the ethnicity groups (labeled 1–5 in the first column). Each ethnic group and subgroup was compared with group 1, as the baseline group. There was strong evidence that groups 3 (and all its subgroups except those of Indian origin) and 4 had a higher rate than group 1. For black/African/Caribbean/black British there was evidence of a higher rate than group 1. Whereas there was no evidence that groups 2 and 3 were differed from 1.
The mean BMI SD score at diagnosis was 2.89 (girls, 2.88; boys, 2.92). Of the 106 cases, 102 (96%) were at least overweight and 81% (86/106) were obese. A total of 2% had a normal BMI and 2% had no BMI measured. Comparisons of BMI at diagnosis for each ethnic group are shown in Fig. 2, and Table 2 shows comparisons of the mean BMI by ethnicity, compared with white children. Asian children had a lower BMI at presentation than white children (ethnicity, compared with white children. Asian children had a lower BMI at presentation than white children.

A family history of Type 2 diabetes was present in 81% of cases, with a first-degree relative affected in 70% of cases (17% both parents affected, 50% mother alone affected, 23% father alone affected, 10% sibling alone affected) and a second-degree relative in 11%.

In 35% of cases (37/106) the child was asymptomatic at diagnosis, with diagnosis being made after an incidental glucose finding or during a clinical assessment for obesity-related comorbidities. In 53% of cases (56/106), osmotic symptoms suggestive of diabetes were present (either polyuria, polydipsia, nocturia or weight loss). Recurrent infections were reported in 18% of cases (19/106), with genital/perineal infections being the most common (47%). Other infections included cellulitis/skin abscesses (26%) and urinary tract infections (11%). Lethargy was reported in 15% of the children (16/106). Half of all cases had evidence of ketonuria at diagnosis. Fewer than five cases presented with diabetic ketoacidosis or a hyperosmolar hyperglycaemic state.

At diagnosis, non-alcoholic fatty liver disease was reported in 37%, hypertension in 21%, dyslipidaemia in 9%, renal disease in 3% and psychological comorbidities in 2% of cases.

A total of 104 newly confirmed cases were from the UK (excluding two from Republic of Ireland), of which 94 were diagnosed in the last consecutive 12 months of the study (May 2015 to April 2016) and were used for calculations of incidence. The estimated UK population aged 0–16 years at mid-2015 was 13 008 432 (www.ons.gov.uk), resulting in a national UK incidence of Type 2 diabetes in children aged 0–16 years of 0.72 per 100 000 (95% CI 0.58–0.88). For the ethnic breakdown of incidence, data from England and Wales were available. When compared with white children...
**FIGURE 1** Flowchart of cases’ classification. D, Duplicate report; DP, Duplicate positive (i.e. duplicate of confirmed case); IRE, Republic of Ireland; NP, No response received

**FIGURE 2** Box-plots for the BMI z-scores (SD scores) of children aged 0–16 years in UK, grouped by their ethnicity

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DIABETIC rate ratio 1.35, 95% CI 0.99–1.84), although this was not statistically significant (P=0.062; Table 3). There was statistical evidence that the incidence had increased among girls (P=0.03) and South-Asian children (P=0.01) over the decade (Table 3).

Discussion

In the UK, the incidence of Type 2 diabetes among girls and South-Asian children is increasing. The first study to quantify accurately the incidence in the UK was in 2005 [6], using a similar methodology to that used in the present study and thereby allowing comparison a decade later. Sixty-seven cases were reported in 2004–2005, while the present study identified 106 cases, with incidence rising from 0.53 per 100 000 (95% CI 0.41–0.68) in 2004–2005 to 0.72 per 100 000 (95% CI 0.58–0.88). The incidence in white children in 2004–2005 was 0.35 per 100 000 (95% CI 0.2–0.5) and in 2015–2016 it was 0.43 per 100 000 (95% CI 0.30–0.58), demonstrating a modest increase. Generally, the higher national incidence is a result of significantly higher rates among ethnic minorities, despite Asian and BACBB children accounting for 9.7% and 4.8% of the population in England and Wales respectively. Compared with white children, Asian children had an incidence nearly seven times and BACBB children nearly four times greater. The rise in cases among Asian children means that they represent a higher proportion of cases compared with BACBB children, while previously the reverse was described [6].

Previous estimates of the prevalence of Type 2 diabetes among young people of Asian ethnicity may be underestimates because a proportion were initially misdiagnosed as having Type 1 diabetes [20]. The UK incidence is still lower than in the USA [4], Japan and Taiwan [3] but higher than in previous studies from Europe [3]. The higher incidence in ethnic minority groups has been described in many previous studies [4]. It is notable that only one case in 2005 and two cases from 2015 were reported from the Republic of Ireland. It could be speculated that this relates to the less diverse ethnic background of the citizens in that country.

A major driver of the development of Type 2 diabetes is obesity. Most children in our cohort were overweight or obese. With 53% of the cases presenting with osmotic symptoms including weight loss, the BMI recorded at diagnosis may be lower than earlier in the disease process. Asian children present with Type 2 diabetes at a significantly lower BMI than do white children, which has been described previously [21] and may be related to ethnic differences in visceral fat distribution which corresponds to increased metabolic risk. Obesity prevalence in the UK has increased dramatically over the last 20 years. Childhood obesity increased between 1994 and 2003, although the overall trend stabilized between 2004 and 2013. In the 11–14-year age group, however, there was a continued trend in increased obesity albeit at a slower rate than the previous decade [22]. The increased incidence of Type 2 diabetes may not be solely explained by rising obesity rates but may also be attributable to greater awareness of the disease in the paediatric population, changing demographics in the UK and better screening for comorbidities related to obesity.

Family history of Type 2 diabetes clearly plays a role in developing the condition in childhood. The interplay between genetic predisposition and obesogenic environment (which families often share) can be difficult to characterize. Impaired glucose handling is seen more frequently in non-obese children with a family history of Type 2 diabetes than in those with no family history [23], suggesting the importance of genetic predisposition. Two-thirds of the cases were female, significantly higher than in 2005 [6]. This female preponderance has been described previously [10] and the UK National Paediatric Diabetes Audit reported a female: male ratio of ~2:1 [7]. Girls tend to reach puberty earlier and puberty is associated with fat accumulation. This may contribute to greater insulin resistance in girls than in boys of a similar age. Physical activity levels also tend to be lower among adolescent girls than boys [24] and this may be detrimental to metabolic health.

The present study suggests the importance of screening for Type 2 diabetes in obese individuals, as 35% of our cases were asymptomatic at diagnosis. UK national guidelines recommend assessment (including consideration of measurement of fasting glucose and insulin, HbA1c, and an oral glucose tolerance test) of comorbidities related to obesity, including diabetes in those above the 98th centile for BMI [25]. Recurrent infections, especially those affecting the genitalia, were the most common presentation after osmotic symptoms. We advocate screening for Type 2 diabetes in overweight children presenting with recurrent perineal or cutaneous infections. Ketoacidosis is a rare presentation in Type 2 diabetes but does occur, as evidenced by cases in our cohort.

Non-alcoholic fatty liver disease was the most commonly reported comorbidity (37%) and has previously been reported in approximately half of adolescents with Type 2 diabetes [26]. A recent study in children and adolescents with non-alcoholic fatty liver disease reported a prevalence of impaired glucose tolerance of 23.4% and of Type 2 diabetes of 6.5% [27]. Our finding of hypertension in 21% of children at diagnosis is consistent with other studies, which have reported rates of hypertension in adolescents of 10–55% at diagnosis [28,29], although it was lower than the 34% rate reported in the 2005 cohort [30]. In 11 cases no blood pressure data were available, therefore, cases of
hypothesis may have been missed. Our observation of renal disease in 3% of cases is low compared with previous studies [29], which have reported rates of 7–22% of microalbuminuria at diagnosis in adolescents. This relatively small number of cases with renal disease is similar to the number observed in the 2005 cohort, in which early nephropathy was rare (4%) [30]. This may reflect either a lack of screening for renal disease by clinicians, as 32% of cases did not report data on albuminuria, earlier diagnosis of diabetes, or a lower incidence of microalbuminuria. Certainly, as the rate of progression of microalbuminuria and renal disease is rapid in adolescents with Type 2 diabetes, with 28% prevalence of microalbuminuria reported at 1.3 years after diagnosis [29], careful screening and monitoring is crucial. Follow-up data on this cohort will give valuable insights into any progression and change in prevalence of nephropathy.

Surveys relying on clinicians reporting data have the potential to underestimate the true incidence of a disease. Clinical information was available from 95% of notifications, thus there was only a small proportion (5%) of missing data resulting from incomplete reporting. Over the study period, there was a mean monthly orange card return of 92%, leaving potential for cases not being reported. There was a mean monthly case notification card return of 93% in the study in 2005 [6]; therefore, the potential for error was similar in the two studies. It is recommended that children with diabetes are managed in secondary care, however, it is possible that some young people, especially those aged 16–17 years, with Type 2 diabetes are managed in primary care. These cases would not be detected by the BPSU reporting system.

Our data suggest the number of cases of childhood Type 2 diabetes is rising in the UK, although Type 2 diabetes in children is still unusual compared Type 1 diabetes [7]. Female gender, non-white ethnicity and a family history of Type 2 diabetes were strongly associated with the disease. Follow-up data from the present cohort will characterize the management, clinical course and development of comorbidities in this population. Further work is needed to prevent the development of Type 2 diabetes and to understand the processes leading to Type 2 diabetes developing amongst children with obesity. The long-term health and economic implications of the rising incidence in the UK will become evident in the coming decades, although this should be of great concern to clinicians and policy makers.

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Competing interests

None declared.

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