Type 2 diabetes in adults: choosing medicines

Factors to take into account when choosing, reviewing and changing medicines

Prescribing guidance

Choosing treatments
Base the choice of medicine on:
• the person’s individual clinical circumstances, for example comorbidities, contraindications, weight, and risks from polypharmacy
• the person’s individual preferences and needs
• the effectiveness of the drug treatments in terms of metabolic response and cardiovascular and renal protection
• safety (see MHRA guidance, the BNF and individual SPCs) and tolerability of the drug treatment
• monitoring requirements
• the licensed indications or combinations available
• cost (if 2 drugs in the same class are appropriate, choose the option with the lowest acquisition cost)

Reviewing and changing treatments
At each point, think about and discuss the following with the person:
• stopping medicines that are not tolerated
• stopping medicines that have had no impact on glycaemic control or weight, unless there is an additional clinical benefit, such as cardiovascular or renal protection, from continued treatment
• how to optimise their current treatment regimen before thinking about changing treatments, taking into account factors such as:
  – adverse effects
  – adherence to existing medicines
  – the need to revisit advice about diet and lifestyle
  – prescribed doses and formulations
• whether switching rather than adding drugs could be effective

High risk of cardiovascular disease
Adults with type 2 diabetes who have:
• QRISK2 more than 10% in adults aged 40 and over or
• an elevated lifetime risk of cardiovascular disease (defined as the presence of 1 or more cardiovascular risk factors in someone under 40).

Cardiovascular disease risk factors: hypertension, dyslipidaemia, smoking, obesity, and family history (in a first-degree relative) of premature cardiovascular disease.

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This is a summary of the advice in the NICE guideline on type 2 diabetes in adults: management.
How to choose first-line medicines

Rescue therapy
For symptomatic hyperglycaemia, consider insulin or a sulfonylurea and review when blood glucose control has been achieved.

First-line treatment

Assess HbA1c, cardiovascular risk and kidney function

Not at high CVD risk

Offer
- Metformin
- Or if GI disturbance
- Metformin MR

Chronic heart failure or established atherosclerotic CVD

Offer
- Metformin
- Or if GI disturbance
- Metformin MR

and as soon as metformin tolerability is confirmed, offer
- SGLT2 inhibitor ('flozin')
  - with proven cardiovascular benefit

High risk of CVD

QRISK2 of 10% or higher elevated lifetime risk

Offer
- Metformin
- Or if GI disturbance
- Metformin MR

and as soon as metformin tolerability is confirmed, consider
- SGLT2 inhibitor ('flozin')
  - with proven cardiovascular benefit

Person's HbA1c not controlled below individually agreed threshold, or the person develops CVD or a high risk of CVD

See treatment options if further interventions are needed

Established atherosclerotic CVD includes coronary heart disease, acute coronary syndrome, previous myocardial infarction, stable angina, prior coronary or other revascularisation, cerebrovascular disease (ischaemic stroke and transient ischaemic attack) and peripheral arterial disease.

NICE technology appraisals recommend SGLT2 inhibitors as monotherapy options in people:
- who cannot have metformin
- for whom diet and exercise alone do not provide adequate glycaemic control.

The SGLT2 inhibitors are recommended only if a dipeptidyl peptidase-4 (DPP-4) inhibitor would otherwise be prescribed and a sulfonylurea or pioglitazone is not appropriate.

In February 2022, using ertugliflozin to reduce cardiovascular risk when blood glucose is well controlled was off label. See NICE's information on prescribing medicines.
When exercising their judgement, professionals and practitioners are expected to take this guideline fully into account, alongside the individual needs, preferences and values of their patients or the people using their service. It is not mandatory to apply the recommendations, and the guideline does not override the responsibility to make decisions appropriate to the circumstances of the individual, in consultation with them and their families and carers or guardian.

This information is a summary of the recommendations, please consult the guideline for the full recommendations. All supplementary information is taken from the BNF or the SPCs.

In February 2022, using ertugliflozin to reduce cardiovascular risk when blood glucose is well controlled was off label. See NICE's information on prescribing medicines.

See summaries of product characteristics (SPCs), British national formulary (BNF) or the Medicines and Healthcare products Regulatory Agency (MHRA) for up-to-date information.

### Summary of first-line medicines

<table>
<thead>
<tr>
<th>Medicine</th>
<th>Options and BNF link</th>
<th>Contraindications or special warnings (see SPCs)</th>
<th>Effect on weight</th>
<th>Hypoglycaemia risk</th>
<th>Renal impairment</th>
<th>Hepatic impairment</th>
</tr>
</thead>
<tbody>
<tr>
<td>DPP-4 inhibitor ('gliptins')</td>
<td>Alogliptin Linagliptin Saxagliptin Sitagliptin Vildagliptin</td>
<td>Ketoacidosis</td>
<td>None</td>
<td>Low</td>
<td>Dose reduction or caution (not for linagliptin)</td>
<td>Dose reduction or caution or avoid (not for linagliptin and sitagliptin)</td>
</tr>
<tr>
<td>Metformin</td>
<td>Metformin</td>
<td>Acute metabolic acidosis</td>
<td>None</td>
<td>Low</td>
<td>Dose reduction or avoid. Check the BNF monograph for eGFR thresholds</td>
<td>Withdraw if tissue hypoxia likely</td>
</tr>
<tr>
<td>Pioglitazone</td>
<td>Pioglitazone</td>
<td>Ketoacidosis, history of heart failure, previous or active bladder cancer, uninvestigated macroscopic haematuria</td>
<td>Gain</td>
<td>Low</td>
<td>No warnings</td>
<td>Avoid</td>
</tr>
<tr>
<td>SGLT2 inhibitor ('flozins')</td>
<td>Canagliflozin Dapagliflozin Empagliflozin Ertugliflozin</td>
<td>Ketoacidosis</td>
<td>Loss</td>
<td>Low</td>
<td>Dose reduction or caution or avoid. Check the BNF monographs for eGFR thresholds</td>
<td>Caution or avoid. Check the BNF monographs for severity</td>
</tr>
<tr>
<td>Sulfonylurea</td>
<td>Gliclazide Glimepiride Glipizide Tolbutamide</td>
<td>All sulfonylureas: ketoacidosis Gliclazide and tolbutamide: avoid where possible in acute porphyrias</td>
<td>Gain</td>
<td>Moderate High in older people</td>
<td>Dose reduction or caution or avoid. Check the BNF monographs for eGFR thresholds</td>
<td>Caution or avoid. Check the BNF monographs for severity</td>
</tr>
</tbody>
</table>

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