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CLINICAL INQUIRIES

Does metformin prevent diabetes in at-risk adults?

J Fam Pract. 2013 August;62(8):436-437

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Author and Disclosure Information





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EVIDENCE-BASED ANSWER

Yes. Metformin therapy reduces the risk of developing diabetes for adults with one or more risk factors for at least 5 years (strength of recommendation[SOR]: A, consistent meta-analyses) and perhaps as long as 10 years (SOR: B, randomized clinical trial[RCT]).

Lifestyle modification with diet and exercise is approximately twice as effective as metformin for preventing diabetes, especially in older patients (SOR:B,RCTs).

Evidence summary

Three meta-analyses of studies lasting from 2 months to 5 years found that metformin reduced the risk of developing overt diabetes in at-risk adults when compared with placebo (TABLE). 1-3

Metformin is likely effective for as long as 10 years, based on long-term follow-up of patients in the Diabetes Prevention Program (DPP). In this trial, investigators randomized 3234 at-risk patients to 3 groups: metformin 850 mg twice daily; lifestyle modification (7% weight loss, 150 minutes of physical activity per week, and a oneto-one 16-lesson curriculum covering diet, exercise, and behavior modification); or placebo. At a mean 2.8-year follow-up, the incidence of diabetes was 31% lower in the metformin group (95% confidence interval [CI], 17%-43%) and 58% lower in the lifestyle modification group than in the placebo group (95% CI, 48%-66%; P<.001 for both comparisons).

At the close of the DPP trial, investigators offered lifestyle intervention to all 3 groups. Patients in the original metformin group continued to take metformin (with participants unblinded to assignment); patients in the original lifestyle intervention group were offered additional lifestyle support. 5 At a median follow-up of 10 years after initial enrollment in the DPP trial, metformin reduced the incidence of overt diabetes by 18% compared with placebo (95% CI, 7%-28%), and lifestyle intervention reduced it by 34% (95% CI, 24%-42%; no statistic of comparison supplied).

Lifestyle modification works better than metformin in older adults

Another analysis found that metformin was equally effective in preventing diabetes in older and younger

1 of 2 3/6/19, 6:00 PM patients, whereas lifestyle modification was more effective in older patients. Investigators followed the patients in the DPP trial for an additional 5 months and stratified the effect of metformin and lifestyle modification by age.⁶

Metformin's effectiveness didn't change significantly in older adults compared with younger adults (hazard ratio [HR] for developing diabetes at age 60-85 years vs 25-44 years=1.45; 95% CI, 0.98-2.16; P=.06). In contrast, lifestyle modification worked better in older adults than younger adults (HR at age 60-85 years vs 25-44 years=0.47; 95% CI, 0.28- 0.78; P<.01).

Recommendations

The American Diabetes Association says that physicians may consider using metformin to prevent type 2 diabetes in patients at the highest risk, such as patients with multiple risk factors, especially if they show progression of hyperglycemia (HbA1c \geq 6%, for example) despite lifestyle intervention.⁷

Evidence-based answers from the Family Physicians Inquiries Network

▼ References

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