

1 Insulin Resistance and Pharmacotherapy Effectiveness in 2 Patients with Long -Term Diabetes Mellitus Type 2

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6 Abstract

7 Introduction: With the development of the availability of genetic research, a multiplex
8 approach to determining the pharmacotherapy tactics has become possible, taking into
9 account the individual characteristics of the patient. Nevertheless, the modern tactics of
10 pharmacotherapy "to failure" has been adopted, which over time leads to the intensification of
11 therapy. A personalized approach to the pharmacotherapy of type 2 diabetes mellitus by
12 determining the genotype of endothelial synthase of nitric oxide will predict the effectiveness
13 of metformin monotherapy in the debut of the disease and reduce the risk of decompensation
14 and complications with long-term type 2 diabetes mellitus. The aim of the study was to
15 monitor the effectiveness of metformin pharmacotherapy for long-term type 2 DM, depending
16 on the patient's genotype.

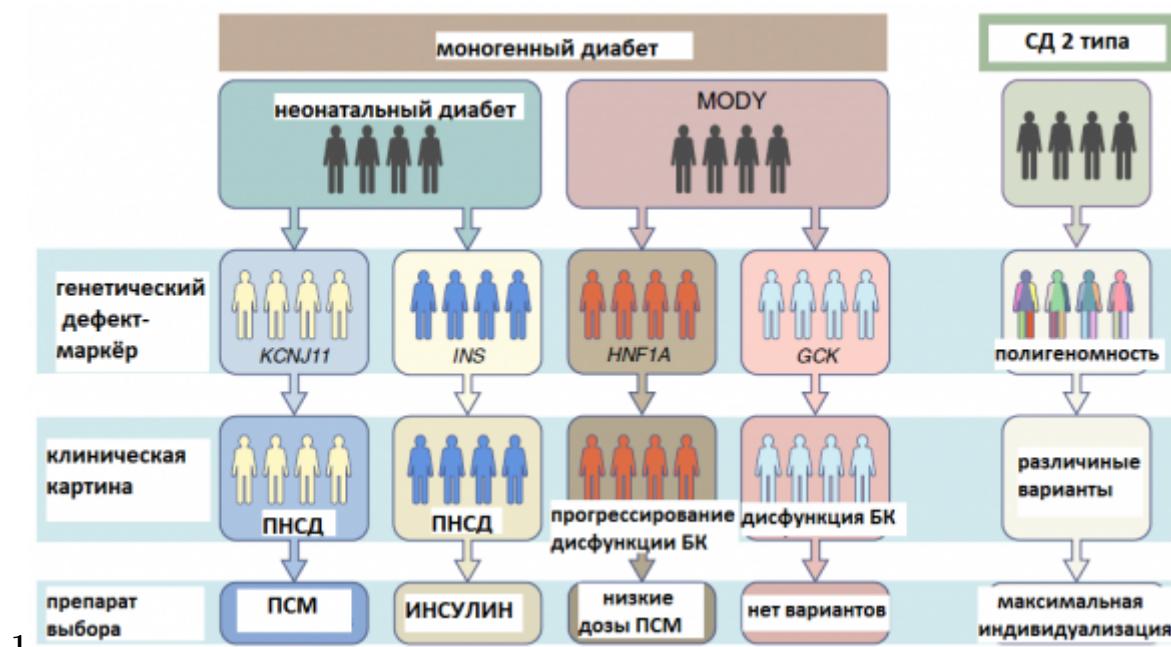
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18 **Index terms**— type 2 diabetes mellitus, monotherapy, metformin, eNOS3 gene polymorphism, insulin
19 resistance.
20 of genetic research, a multiplex approach to determining the pharmacotherapy tactics has become possible,
21 taking into account the individual characteristics of the patient. Nevertheless, the modern tactics of pharma-
22 cotherapy "to failure" has been adopted, which over time leads to the intensification of therapy. A personalized
23 approach to the pharmacotherapy of type 2 diabetes mellitus by determining the genotype of endothelial synthase
24 of nitric oxide will predict the effectiveness of metformin monotherapy in the debut of the disease and reduce the
25 risk of decompensation and complications with long-term type 2 diabetes mellitus.

26 The aim of the study was to monitor the effectiveness of metformin pharmacotherapy for long-term type 2 DM,
27 depending on the patient's genotype. Materials and methods: A single-center, randomized, prospective study
28 of 200 patients referred for planned hospitalization. The single nucleotide polymorphism of the gene eNOS3,
29 the level of glycated hemoglobin. Homeostatic model 2 (HOMA 2) was applied to evaluate insulin resistance
30 in patients. In consistency with the results, patients were divided into three groups: with the ??, TC and TT
31 genotype.

32 Results and discussion: Patients of the CC genotype group achieved and kept glycated hemoglobin below the
33 target level in 80% of cases, when metformin was used at a dose of 1700 mg per day. No one of the patients, who
34 representatives of the TC and TT genotypes reached the target values of glycated hemoglobin. Thus, the use of
35 metformin as monotherapy to compensate for carbohydrate metabolism is not enough for TC and TT genotypes,
36 and combination hypoglycemic pharmacotherapy, including insulin therapy, is required.

37 Conclusion: Identification of the presence of a particular eNOS3 gene allele provides the possibility of earlier
38 prescribing appropriate drugs in an individual dose to reduce insulin resistance and likely limit the progression
39 of diabetes mellitus, as well as increase the degree of compensation for carbohydrate metabolism.



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Figure 1: ??????? 1 :

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Figure 2: ????????. ??? ?? ?????, ??????? ?????????????? ??????? ?????????????? «?? ??????»,
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Figure 3: ? ??? ????. ??????????????????. ??????????: ?????????? ??????? ??? ???. ????. ???? ?????? ????
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Abstract-Introduction: With the development of the availability

Figure 4: ?????????? ?????????? ?????????????????? ?????????? ???????, ? ????. ??????????
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40 **1 Keywords**
41 ^{1 2}

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16 : type 2 diabetes mellitus, monotherapy, methformin, eNOS3 gene polymor- 17
 Yeaphism, insulin resistance. I. ?????????? ???????????, ??? ?????????? ?????? (?)D?”) Year
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