



Cost-Effectiveness Analysis of Tenueligliptin V/S Glimepiride as an Add-on to Metformin in Type 2 Diabetes Mellitus



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Highlights

- The metformin-glimepiride combination demonstrated superior cost-effectiveness compared to metformin-tenueligliptin for managing type 2 diabetes.
- Both therapies provided satisfactory blood glucose reduction, but the glimepiride-based regimen achieved this at a significantly lower drug acquisition cost.
- For cost-conscious patients without contraindications, metformin and glimepiride is recommended as the more economically efficient initial combination therapy.

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ABSTRACT

Background: Diabetes is a major lifestyle disorder. Proper glycemic control is needed to prevent the development of patients with Type 2 diabetes. Glimepiride is a second-generation sulphonyl urea with better safety and efficacy, and is commonly prescribed compared to other drugs of that class. tenueligliptin is a relatively newer dipeptidyl peptidase inhibitor with proven clinical efficacy with metformin. A more cost-effective treatment option is important in the case of people with a poor economic background for better adherence and thereby preventing complications and economic burden.

Methods: A prospective observation study was conducted in a tertiary care hospital to assess and compare the cost-effectiveness of glimepiride 1mg and tenueligliptin 20 mg when added to metformin 500 mg. a total of 112 patients were selected as per the inclusion criteria, 56 of them allocated to group A are taking glimepiride 1 mg and metformin 500 mg once daily. Group B patients are those who are on tenueligliptin 20 mg and metformin 500 mg once daily. Fasting blood glucose and post-prandial blood glucose of each patient in both groups were recorded as baseline.

Results: The average cost-effectiveness ratio is calculated using the average cost of therapy for 4 months and the average reduction in effectiveness for 4 months. In this study, cost data included direct costs of purchasing the drugs. cost-effectiveness analysis revealed that glimepiride metformin combination showed better effectiveness in terms of both FBG and PPBG reduction. Tenueligliptin and metformin therapy have shown satisfactory reduction in blood glucose levels, but it is much costlier Compared to tenueligliptin, glimepiride is a better choice as an add-on drug in the absence of any contraindication in patients who are highly concerned about cost, to reduce economic burden, and to improve adherence, when used as an initial combination in patients with Type 2 diabetes mellitus. .

Keywords: Diabetes mellitus, Cost effectiveness, Metformin hydrochloride, Glimepiride, Tenueligliptin



Introduction

Diabetes mellitus is a lifestyle disorder and a cause of premature mortality due to cardiovascular conditions and renal failure¹. The development and progression of diabetes related complications can be minimized by good glycemic control². The purpose of pharmacotherapy in the management of type 2 diabetes mellitus is to control hyperglycemia and prevent hyperglycemia-related complications³. Evidence suggests that combined therapy with oral hypoglycemics is more effective than single drug therapy⁴. Metformin is the first drug of choice for diabetes control because of cardiovascular safety, low chance of hypoglycemia, durability, no incidence of weight gain, advantages in weight reduction, blood pressure reduction, and low cost⁵. Along with metformin, other classes of oral antidiabetics are added to attain better glycemic control⁶. The second-generation sulphonyl urea glimepiride is commonly added with metformin since it has having relatively better safety and efficacy profile among its group⁷. Teneiglipitin is a dipeptidyl peptidase inhibitor that has got clinical benefits along with metformin⁸.

Being a chronic condition, diabetes needs lifelong treatment. Economic burden is also a factor of consideration while prescribing medications⁹. chronicity of diabetes affects the financial status of the individual and family. The prevalence of diabetes mellitus is increasing day by day, so a great deal of expenditure for the country is also increasing¹⁰. For long-term therapy of diseased, along with the safety of medications, cost also becomes a major concern¹¹.

This study planned to compare cost cost-effectiveness of the combination therapy of teneiglipitin 20 mg and metformin 500 mg, with the glimepiride 1 mg and metformin 500 mg combination. Both glimepiride and teneiglipitin are used as add-on therapy with metformin, but the costs are different. For long-term therapy, the cost-effectiveness is a factor for economic considerations like resource allocation, insurance, and making health policy decisions. In resource-constrained settings like India, assessing the cost-effectiveness of antidiabetic therapies is crucial for rational prescribing. Scientific evaluation beyond just drug price helps

physicians choose optimal regimens for socioeconomically disadvantaged patients. Although DPP-4 inhibitors are often preferred in Western studies, Indian data comparing them with sulfonylureas is limited. A recent study found that while both metformin-glimepiride and metformin-teneiglipitin combinations showed similar glycaemic control, the former was significantly more cost-effective. Short-term effects on BMI were comparable. These findings stress the importance of local pharmacoeconomic evaluations and the need for larger, long-term studies in Indian populations¹². A recent study comparing the efficacy and safety of Teneiglipitin versus Glimepiride as add-on therapy to Metformin in patients with type 2 diabetes found that Teneiglipitin was well tolerated and more effective in improving both glycaemic and lipid profiles. Patients receiving the Metformin-Teneiglipitin combination demonstrated better overall metabolic control compared to those on the Metformin-Glimepiride regimen, indicating Teneiglipitin as a superior second-line option. As a modern DPP-4 inhibitor, Teneiglipitin offers notable advantages in terms of safety and efficacy, making it a promising choice in the evolving landscape of anti-diabetic therapies. However, given that the study included only 30 patients per group, larger-scale studies with extended follow-up are warranted to validate these findings¹³.

Sitaglipitin, a selective DPP-4 inhibitor, offers a novel approach in the management of type 2 diabetes mellitus. In a comparative study, the combination of metformin and sitagliptin was evaluated against metformin and glimepiride in 60 patients over 24 weeks. Both groups showed significant reductions in fasting blood glucose (FBG), postprandial blood glucose (PPBG), and HbA1c. However, the metformin-sitaglipitin group showed a greater reduction in PPBg at the end of the study. The incidence of adverse effects such as hypoglycaemia and gastrointestinal discomfort was noted in both groups. These findings suggest that sitagliptin may be a safer and effective second-line agent alongside metformin¹⁴.

A study by John et al. (2021) highlighted that the majority of patients were managed with a combination of Metformin and Glimepiride,

which demonstrated effective control of HbA1c and random blood sugar (RBS) levels. This combination was found to be beneficial in achieving glycaemic targets in type 2 diabetes patients. Additionally, it showed a higher clinical effectiveness ratio (CER) compared to other combinations. The findings suggest that Metformin + Glimepiride therapy not only improves glycaemic control but is also a cost-effective option in diabetes management¹⁵.

A recent study evaluated the effects of Canagliflozin and Tenofovir as third-line agents in patients with uncontrolled type 2 diabetes already on Metformin and Glimepiride. Both drugs demonstrated significant improvements in glycaemic parameters, including HbA1c, fasting, and postprandial glucose levels. Additionally, Canagliflozin showed greater benefits in reducing blood pressure and improving lipid profiles, particularly HDL-C levels. While both agents retained their glycaemic efficacy in this treatment position, Canagliflozin appeared more effective for patients with comorbid hypertension and diabetic dyslipidaemia. These findings support their potential pleiotropic roles beyond glucose control¹⁶. To assess and compare the cost-effectiveness of glimepiride and tenofovir 20 mg when added to metformin 500 mg.

Methods

The study was an observational comparative one conducted over a period of 6 months in a tertiary care hospital. Newly diagnosed patients of type 2 diabetes mellitus of either sex with fasting blood glucose more than 126mg/dL were included. Pregnant and lactating patients were excluded.

Procedure

Patients were selected as per the inclusion criteria. Patients enrolled in group A are taking

glimepiride 1 mg and metformin 500 mg once daily. Group B patients are those who are on tenofovir 20 mg and metformin 500 mg once daily. Fasting blood glucose and post-prandial blood glucose of each patient in both groups were recorded as baseline. First review values of FBG and PPBG recorded at the end of the second month, after the first, final review values recorded at the end of the fourth month. The efficacy of two drugs with metformin was assessed by a reduction in terms of FBG, PPBG at the first review from baseline and at the second review from the first review. The average cost-effectiveness ratio is calculated using the average cost of therapy for 4 months and the average reduction in effectiveness for 4 months. In this study, cost data included direct costs of purchasing the drugs.

Result and Discussion

A total of 112 patients, 56 each in group A and B, enrolled in the study. Other comorbidities observed were hypertension and hyperlipidemia.

In group A, patients are categorized into 5 ranges based on the values of their FBG level. In each category individual patient's reduction in FBG from review 1 and baseline, from review 2 and review 1 calculated, and the average of each group was taken. Then the total average value was calculated, and the same procedure was applied to group B, for PPBG in groups A and B. In group A average reduction in FBG level from baseline to review 1 was 5.14mg/dL, and from review 1 to review 2, it was 7.32mg/dL. The total average reduction in FBG level is found to be 6.23mg/dL. In group B reduction in FBG level from baseline to review 1 is 4.55mg/dL, and from review 1 to review 2 is 5.67mg/dL, and the total average reduction in FBG is 5.08mg/dL.

Table 1. Drugs used in treatment of varicose veins [22 – 31]

FBS in mg/dL	Group A (metformin500 mg+ glimepiride 1 mg) no of patients			Group B (metformin500mg+ tenofovir 20 mg) number of Patients		
	Baseline	Review 1	Review 2	Baseline	Review 1	Review 2
120-139	6	8	13	3	4	7
140-159	12	15	16	20	23	28
160-179	16	16	14	19	16	12
180-199	16	13	11	9	11	7
>200	6	4	2	5	2	2

Table 2. Drugs used in treatment of varicose veins [22 – 31]

PPBG in mg/dL	Group A (metformin 500mg+ glimepiride 1 mg)			Group B (metformin 500 mg+ teneligliptin 20 mg)		
	Baseline	Review 1	Review 2	baseline	Review 1	Review 2
<or= 180		2	4		2	8
181-209	22	27	33	26	28	28
210-239	17	11	16	18	14	12
240-269	9	8	6	9	9	6
>270	8	8	7	3	3	2

Cost-effectiveness analysis was conducted using costs and effectiveness for both drug combinations. In this study, cost data included direct costs of purchasing the drugs.

Unit cost of drugs obtained from the outpatient pharmacy department. Diabetic medications are supplied by the government through the outpatient pharmacy. The cost of treatment in the two groups was calculated as follows.

The cost for each patient for 4 months is calculated by the unit price of each drug/combination per day multiplied by 30 days. Then calculated for four months. Cost of treatment for a patient for 4 months = unit price of metformin 500 mg, glimepiride 1 mg, or unit price of metformin 500 mg, teneligliptin 20 mg x frequency of medication x duration of treatment days. The average cost for the treatment group is equal to the sum of the costs of total patients/number of patients in each treatment group. Let the cost of drugs for each patient for 4 months be C₁, C₂, and C_n for N number of patients. Then the average cost for a group is $C_1 + C_2 + \dots + C_n / N$.

Effectiveness was calculated as the reduction of FBG and PPBG from baseline, review 1, and review 2.

Let the difference between baseline and review 1 be D₁, the difference between review 2 and review 1 be D₂, then the average difference between review 1 and baseline for all patients in a group is $D_{11} + D_{12} + \dots + D_{1n} / N$, similarly for the difference between review 2 and review 1 is $D_{21} + D_{22} + \dots + D_{2n}$, and the average is $D_{21} + D_{22} + \dots + D_{2n} / N$.

Average reduction in glucose level in each group = sum of average reduction/number of patients in each group.

Total Average reduction in glucose level in each patient in terms of FBG and PPBG is (difference between baseline and review 1+ difference between review 1 and review 2)/ 2

Average cost effectiveness ratio ACER = average cost of therapy/average effectiveness.

Incremental Cost Effectiveness Ratio = (total cost Group A – total cost group B)/ (effectiveness of group A – effectiveness of group B)

Calculation of cost for 4 months

Unit price of metformin 500 mg is 0.40 RS and glimepiride 1mg is 0.17RS and teneligliptin 20 mg is 0.70 RS.

Name of drug	Unit price in RS (P)	Cost for 4 months in RS Px30x4
metformin 500 mg	0.40	48
glimepiride 1mg	0.17	20.4
teneligliptin 20 mg	0.70	84

Calculation of cost effectiveness

GROUP	Average Cost for 4months in RS	Average Reduction in FBG mg/dL	Average Reduction in PPBG mg/dL	Average Cost effectiveness ratio in FBG	Average Cost effectiveness ratio in PPBG
A	68.4	6.23	6.9	10.98	9.91
B	132	5.08	6.48	25.98	20.37

Calculation of ICER

group	Cost A in RS	Cost B in RS	Cost A- CostB in RS	Effectiveness A	Effectiveness B	EffectivenessA- EffectivenessB	ICER
FBG	68.4	132	-63.6	6.23	5.08	1.15	-55.30
PPBG	68.4	132	-63.6	6.9	6.48	0.42	-151.42

Conclusion

Group A is having a reduction of 6.23mg/dL FBG level, and group B is showing 5.08 mg/dL. The average cost needed for group A for 4 months of treatment is RS 68.4, and for group B is RS 132. The average cost-effectiveness ratio for group A in FBG level is 10.98, and for group B is 25.98.

In the PPBG group, A has having average reduction of 6.9 mg/dL, and group B has a reduction of 6.48 mg/dL. Average cost effectiveness ratio for group A in PPBG level is 9.91 and for group B is 20.37. In cost effectiveness analysis cost is taken in monetary units and effectiveness in nonmonetary units, such as reduction in FBG and PPBG

Group A is showing the least cost-effectiveness ratio, 10.98 in terms of reduction in FBG level and 9.91 in PPBG level. Hence, group A is more cost-effective. A combination of metformin 500 mg and glimepiride 1 mg shows better effectiveness than the teneligliptin 20 mg and metformin 500 mg combination, in FBG and PPBG reduction, and at least cost too. Teneligliptin 20 mg and metformin 500 mg combination has shown

satisfactory reduction in FBG and PPBG levels, but it is more expensive.

As per the calculation of ICER: The cost is high in the group B and low in group A, and effectiveness in terms of reduction in FBG and PPBG is high in Group A, which means the group A is better. Here group A is dominating.

Limitation of the Study

For long-term, sulphonyl urea remains the second-line add-on therapy with metformin when glycemic control becomes inadequate, but it has side effects like weight gain and hypoglycemia. Dipeptidyl peptidase (DPP-4) inhibitors provide similar efficacy but are free from the above adverse effects and also free from cardiovascular risks. Even though the first combination is cost-effective for patients with a risk of hypoglycemia and cardiovascular problems, the second one is a better choice. Effectiveness of the combinations are different, adverse effects and complications are also different, so considering only the acquisition cost of medications without considering the other medical costs may influence the cost effectiveness ratio.

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