

# The association between polypharmacy and its adverse health outcomes in adult patients with type 2 diabetes mellitus: A systematic review and meta-analysis

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## Background

- Type 2 diabetes mellitus (T2DM) remains a significant chronic disease for adults. **Nowadays more adults have been diagnosed with T2DM and classified as early-onset patients.**
- The age-standardized global T2DM prevalence rate was 5,282 per 100,000 population, with a projected increase to 9.5% by 2050, impacting over 1.27 billion individuals.
- The issue of polypharmacy has garnered increasing attention due to the simultaneous development of new drugs and extended life expectancy.
- 72.2% of adult patients with T2DM were prescribed polypharmacy ( $\geq 5$  medicines) according to a database study in China. **Yet the possible detriment of polypharmacy has not been fully explored.**

## Objective

- To present a **summary of the research literature** on polypharmacy and its association with adverse health outcomes in adult T2DM patients.

## Methods

- A **systematic review** was conducted across three databases (PubMed, Web of Science, and ScienceDirect) through October 2023.
- Studies regarding the relationship between polypharmacy and diabetes-related health outcomes, complications as well as multi-morbidity were included.
- For one specific outcome, qualified data from more than two studies were **synthesized in the meta-analysis with a fixed effects model for the odd ratio (OR) and corresponding 95% confidence intervals (CIs)** and  $I^2$  test for the measure of heterogeneity.

## Results

- Among 24 studies that met the inclusion criteria, three were included in the meta-analysis.
- The association between polypharmacy and poor glycemic control was found to be statistically significant (OR=1.84, 95% CI [1.68-2.01],  $P < 0.00001$ ,  $I^2 = 0\%$ ).**

Study or Subgroup	Case Events	Case Total	Control Events	Control Total	Weight	Odds Ratio M-H, Fixed, 95% CI
Wang Jingqing et al. 2021	3280	5591	1245	2852	97.3%	1.83 [1.67, 2.01]
Firehiwot Dawite et al. 2023	34	50	122	262	1.8%	2.44 [1.28, 4.63]
Hasniza Zaman Huri et al. 2015	126	226	8	16	0.9%	1.26 [0.46, 3.48]
<b>Total (95% CI)</b>		<b>5867</b>		<b>3130</b>	<b>100.0%</b>	<b>1.84 [1.68, 2.01]</b>
Total events	3440		1375			
Heterogeneity: $\chi^2 = 1.28$ , $df = 2$ ( $P = 0.53$ ); $I^2 = 0\%$						
Test for overall effect: $Z = 13.27$ ( $P < 0.00001$ )						

Figure 2: Meta-analysis of studies on association between polypharmacy and poor glycemic control

## Results

- Polypharmacy (the daily/concurrent use of  $\geq 5$  medications or the use of  $\geq 1$  oral antidiabetics/oral antidiabetics combined with insulin) was found to be significantly associated with an 84% higher risk of poor glycemic control in T2DM adult patients.
- Included studies also reported the association between polypharmacy and **hospitalization, death, hypoglycemia, complications (diabetic foot ulcers, non-proliferative diabetic retinopathy), and multi-morbidity (fracture, falls, depression relapse, health-related quality of life).**
- Only insignificant association between stroke and polypharmacy (combined  $\geq 2$  types of antidiabetic medications) was reported.

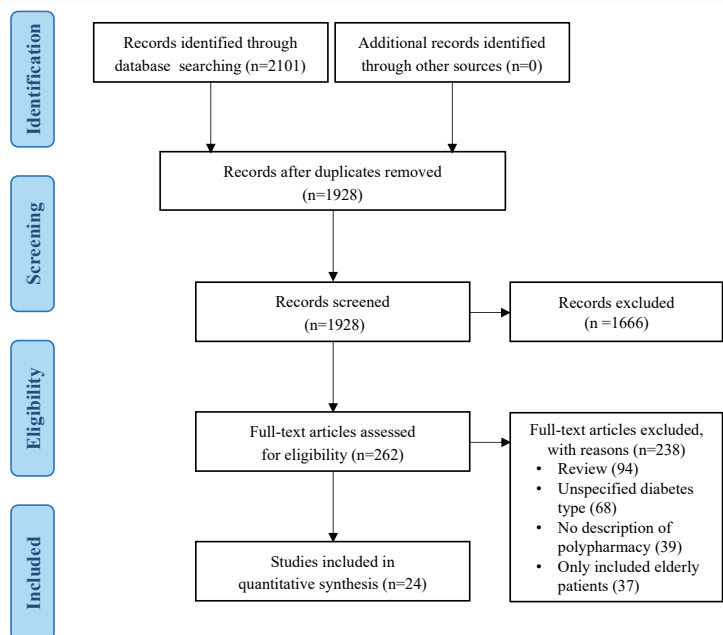


Figure 1: The PRISMA flow diagram for the included studies

## Discussion

- The limitations of this meta-analysis revolve around **the definition of polypharmacy and the reporting of diabetic complications and comorbid conditions.**

## Conclusion

- The current research has revealed a significant association between polypharmacy and several adverse health outcomes in adults with T2DM.
- The evidence underscores the need for **greater caution and improved management in drug therapies** to mitigate these risks.

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