

S4 Table. Weighted mean outcomes and regression results. Mean outcomes for fixed-dose combination and weighted mean outcomes for loose-dose combination cohort in baseline (one year) and follow-up period (three years) with the respective difference-in-difference estimator, its standard error, and the p-value.

S5 Table. Weighted mean outcomes and regression results.

	Year 1					
	FDC (756)		LDC (234)		DiD estimation	
	Baseline	Follow-up	Baseline	Follow-up	ATT	P-value
<i>Adherence and Persistence</i>						
Proportion of days covered	0.69	0.88	0.69	0.66	0.22	<0.001
Days until discontinuation ^a	276.69	N/A	157.79	N/A	119.04	<0.001
<i>Indicators for health care utilization</i>						
Outpatient cases due to diabetes	14.07 5.98	14.06 6.15	14.80 5.78	14.96 6.01	-0.16 -0.07	0.63 0.28
Pharmaceutical prescriptions due to diabetes	22.51 4.23	21.05 4.14	24.12 4.83	26.68 9.32	-4.02 -4.59	<0.001 <0.001
Proportion with emergency visits due to T2D comorbidities	0.12 0.03	0.07 0.01	0.16 0.03	0.09 0.01	0.02 0.01	0.04 0.02
<i>Therapeutic safety</i>						
Proportion with adverse drug events	0.08	0.08	0.11	0.07	0.04	0.03
<i>Comorbidities</i>						
Proportion with microangiopathic complications						
Eye complication	0.17	0.17	0.16	0.17	0.00	0.04
Renal failure	0.13	0.12	0.14	0.14	0.00	0.03
Diabetic foot syndrome/ Peripher neuropathy	0.21	0.21	0.21	0.19	0.02	0.04
Proportion with macroangiopathic complications						
Myocardial infarction	0.04	0.03	0.04	0.03	0.00	0.02
Ischemic heart disease	0.18	0.18	0.20	0.19	0.00	0.03
Angina pectoris	0.03	0.03	0.03	0.02	0.00	0.02
Heart failure	0.05	0.04	0.06	0.05	0.01	0.01
Cerebrovascular disease	0.08	0.09	0.09	0.09	0.00	0.02
<i>Treatment modification^b</i>						
Proportion with alternative						
Oral antidiabetic prescription	N/A	0.25	N/A	0.29	0.00	0.03
Insulins	N/A	0.06	N/A	0.03	0.03	0.01

FDC: fixed-dose combination, LDC: loose-dose combination, ATT: average treatment effect on the treated represents excess outcomes attributable to FDC, SE: standard error

^arefers to discontinuation of treatment in year one of post-index period, estimated using weighted least squares

^bbased on individuals who modified their dual therapy (N: FDC=732, LDC=229)

S5 Table. (Continued) Weighted mean outcomes and regression results.

	Year 2						DiD estimation ATT	P-value	SE			
	FDC (756)		LDC (234)									
	Baseline	Follow-up	Baseline	Follow-up								
<i>Adherence and Persistence</i>												
Proportion of days covered	0.69	0.83	0.69	0.58	0.25	<0.001	0.02					
Days until discontinuation ^a	N/A	N/A	N/A	N/A	N/A			N/A				
<i>Indicators for health care utilization</i>												
Outpatient cases due to diabetes	14.07	13.95	14.80	14.14	0.54		0.63					
due to diabetes	5.98	6.05	5.78	5.66	0.17		0.29					
Pharmaceutical prescriptions due to diabetes	22.51	20.20	24.12	25.29	-3.48	<0.001	1.14					
due to diabetes	4.23	3.29	4.83	7.55	-3.66	<0.001	0.27					
Proportion with emergency visits due to T2D comorbidities	0.12	0.08	0.16	0.08	0.03		0.04					
0.03	0.02	0.03	0.03	-0.01			0.02					
<i>Therapeutic safety</i>												
Proportion with adverse drug events	0.08	0.08	0.11	0.08	0.03		0.03					
<i>Comorbidities</i>												
Proportion with microangiopathic complications												
Eye complication	0.17	0.18	0.16	0.21	-0.03		0.04					
Renal failure	0.13	0.18	0.14	0.19	0.01		0.04					
Diabetic foot syndrome/ Peripherie neuropathy	0.21	0.30	0.21	0.26	0.03		0.04					
Proportion with macroangiopathic complications												
Myocardial infarction	0.04	0.03	0.04	0.02	0.02		0.02					
Ischemic heart disease	0.18	0.19	0.20	0.23	-0.02		0.03					
Angina pectoris	0.03	0.02	0.03	0.02	0.01		0.02					
Heart failure	0.05	0.06	0.06	0.07	0.01		0.02					
Cerebrovascular disease	0.08	0.10	0.09	0.11	0.00		0.02					
<i>Treatment modification^b</i>												
Proportion with alternative												
Oral antidiabetic prescription	N/A	0.58	N/A	0.59	0.02		0.04					
Insulins	N/A	0.17	N/A	0.14	0.03		0.03					

FDC: fixed-dose combination, LDC: loose-dose combination, ATT: average treatment effect on the treated represents excess outcomes attributable to FDC, SE: standard error

^arefers to discontinuation of treatment in year one of post-index period, estimated using weighted least squares

^bbased on individuals who modified their dual therapy (N: FDC=732, LDC=229)

S5 Table. (Continued) Weighted mean outcomes and regression results.

Year 3							
	FDC (756)		LDC (234)		DiD estimation		
	Baseline	Follow-up	Baseline	Follow-up	ATT	P-value	SE
<i>Adherence and Persistence</i>							
Proportion of days covered	0.69	0.81	0.69	0.52	0.29	<0.001	0.03
Days until discontinuation ^a	N/A	N/A	N/A	N/A	N/A		N/A
<i>Indicators for health care utilization</i>							
Outpatient cases	14.07	13.93	14.80	14.14	0.52		0.63
due to diabetes	5.98	6.15	5.78	5.96	-0.02		0.29
Pharmaceutical prescriptions	22.51	20.22	24.12	26.70	-4.87	<0.001	1.77
due to diabetes	4.23	3.16	4.83	7.04	-3.27	<0.001	0.23
Proportion with emergency visits	0.12	0.10	0.16	0.10	0.03		0.04
due to T2D comorbidities	0.03	0.02	0.03	0.01	0.01		0.02
<i>Therapeutic safety</i>							
Proportion with adverse drug events	0.08	0.08	0.11	0.09	0.01		0.03
<i>Comorbidities</i>							
Proportion with microangiopathic complications							
Eye complication	0.17	0.16	0.16	0.23	-0.07		0.05
Renal failure	0.13	0.20	0.14	0.23	-0.02		0.04
Diabetic foot syndrome/ Peripherie neuropathy	0.21	0.31	0.21	0.32	-0.02		0.05
Proportion with macroangiopathic complications							
Myocardial infarction	0.04	0.04	0.04	0.02	0.02		0.02
Ischemic heart disease	0.18	0.21	0.20	0.24	-0.01		0.04
Angina pectoris	0.03	0.03	0.03	0.04	-0.01		0.03
Heart failure	0.05	0.08	0.06	0.08	0.02		0.02
Cerebrovascular disease	0.08	0.10	0.09	0.12	-0.01		0.03
<i>Treatment modification^b</i>							
Proportion with alternative							
Oral antidiabetic prescription	N/A	0.78	N/A	0.79	-0.01		0.03
Insulins	N/A	0.26	N/A	0.29	-0.03		0.04

FDC: fixed-dose combination, LDC: loose-dose combination, ATT: average treatment effect on the treated represents excess outcomes attributable to FDC, SE: standard error

^arefers to discontinuation of treatment in year one of post-index period, estimated using weighted least squares

^bbased on individuals who modified their dual therapy (N: FDC=732, LDC=229)