

SUPPLEMENTARY DATA

Supplementary Table 1. Case listing of bladder cancer related to pioglitazone therapy

CASE	Event year	From clinical study	Age	Gender	Therapy duration (days)	Concomitant Drugs	Co-morbidity
4172635	2004	No	88	M	n.a.	gliclazide acarbose ezetimibe metformin ramipril atorvastatin	
4125536	2004	Yes	73	M	337	lansoprazole	
5653392	2004	Yes	53	M	n.a.	isepamicin bezafibrate famotidine ketoprofen ofloxacin sildenafil sulpiride valsartan ethyl icosapentate	
5731398	2004	Yes	75	M	n.a.	dextromethorphan tranexamic acid telmisartan salicylamide promethazine glimepiride metformin cloperastine amoxicillin cefdinir carbocisteine paracetamol caffeine	
6462605	2005	No	67	M	n.a.	methotrexate tamsulosin sitagliptin simvastatin	psoriatic arthropathy

SUPPLEMENTARY DATA

CASE	Event year	From clinical study	Age	Gender	Therapy duration (days)	Concomitant Drugs	Co-morbidity
						ramipril metformin infliximab folic acid escitalopram ranitidine	
5960713	2005	No	54	M	63	lisinopril simvastatin gemfibrozil clopidogrel	
5996136	2005	Yes	69	M	418	atenolol acetylsalicylic acid	
6280536	2006	No	56	F	n.a.	oxybutynin interferon beta-1a	multiple sclerosis
6209285	2006	No	81	F	n.a.	atorvastatin celecoxib fluticasone gabapentin levothyroxine sodium lisinopril metformin multivitamins risedronic acid salbutamol salmeterol	
6144746	2006	No	62	F	n.a.	salmeterol ipratropium bromide lisinopril potassium insulin glargine vitamins duloxetine	

SUPPLEMENTARY DATA

CASE	Event year	From clinical study	Age	Gender	Therapy duration (days)	Concomitant Drugs	Co-morbidity
						salbutamol glimepiride fluticasone esomeprazole cetirizine calcium gluconate calcium gluceptate acarbose exenatide spironolactone valsartan	
6093650	2006	Yes	66	M	796	gemfibrozil metoprolol simvastatin vitamins nos	
6136429	2006	No	76	M	153	acetylsalicylic acid glimepiride	
6833097	2006	No	73	M	n.a.	acetylsalicylic acid atenolol enalapril	
6061282	2006	Yes	84	M	172	atorvastatin brinzolamide latanoprost levofloxacin timolol	
6070139	2006	Yes	74	M	n.a.	triazolam acetylsalicylic acid nateglinide telmisartan	
6139667	2006	No	79	M	69	simvastatin tolterodine tamsulosin paracetamol	renal failure chronic

SUPPLEMENTARY DATA

CASE	Event year	From clinical study	Age	Gender	Therapy duration (days)	Concomitant Drugs	Co-morbidity
						nicotinic acid mometasone hyoscyamine furosemide esomeprazole darbepoetin alfa clopidogrel amlodipine thiamine metoprolol	
6367838	2007	No	73	M	n.a.		
6288824	2007	No	71	M	n.a.	paracetamol bupropion cyanocobalamin darbepoetin alfa dexamethasone enalapril folic acid loratadine tamsulosin prochlorperazine atorvastatin	non-small cell lung cancer
6654052	2007	No	68	M	1215	omega-3 glipizide furosemide fluticasone clopidogrel tiotropium bromide	
6354279	2007	Yes	71	M	126	clopidogrel pantoprazole	
6517839	2007	No	69	F	318	atenolol atorvastatin acarbose glimepiride	

SUPPLEMENTARY DATA

CASE	Event year	From clinical study	Age	Gender	Therapy duration (days)	Concomitant Drugs	Co-morbidity
6579398	2008	No	70	M	2361	verapamil lisinopril hydrochlorothiazide doxazosin atorvastatin	
6892641	2008	No	74	F	762		
6913281	2008	Yes	71	M	n.a.		
6950249	2008	No	77	M	92	furosemide atenolol glimepiride simvastatin acetylsalicylic acid spironolactone	
7131298	2008	No	53	F	277	escitalopram glimepiride lisinopril metformin verapamil alprazolam	
6796353	2008	No	80	F	n.a.	glibenclamide risedronic acid	
7128421	2009	Yes	73	M	n.a.		
7015483	2009	No	66	M	414	lisinopril metoprolol amlodipine acetylsalicylic acid	
7017054	2009	No	75	M	n.a.	olmesartan medoxomil carvedilol	
7025092	2009	No	60	F	n.a.	exenatide	

SUPPLEMENTARY DATA

CASE	Event year	From clinical study	Age	Gender	Therapy duration (days)	Concomitant Drugs	Co-morbidity
						metformin	

SUPPLEMENTARY DATA

Supplementary Table 2. Stratified analysis for age, gender and event year of ROR for pioglitazone-related bladder cancer

	pioglitazone				
	Cases*	All ADR	ROR	95%CI	p _{MH}
Gender					
M	23	16,867	3.86	2.37-6.26	<0.001
F	8	20,974	5.19	2.15-12.11	<0.001
Age					
65 yr	6	22,738	2.42	0.92-5.98	0.038
> 65 yr	25	15,103	5.10	3.14-8.23	<0.001
Event year					
2004	4	4,221	4.77	1.30-15.88	0.003
2005	3	7,581	3.73	0.83-14.35	0.030
2006	9	10,415	5.24	2.29-11.64	<0.001
2007	5	8,165	3.85	1.26-10.90	0.004
2008	6	4,813	5.23	1.85-14.07	<0.001
2009	4	2646	3.10	0.90-9.53	0.029

*Cases of bladder cancer