

Supplementary Information

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**Trends in Pregabalin Use and Prescribing Patterns in the Adult Population: A 10-Year
Pharmacoepidemiologic Study**

Nofar Benassayag Kaduri, Reuven Dressler, Wiessam Abu Ahmad, Victoria Rotshild

Correspondence to:

Nofar Benassayag Kaduri, Pharmacoepidemiology Research Unit, Institute for Drug Research, School
of Pharmacy, The Hebrew University of Jerusalem, Israel

Nofar.benassayag@mail.huji.ac.il

Table S1 The International Classification of Diseases Version 10 (ICD-10) diagnosis codes of indications [1]

Indications of pregabalin	
Indication	ICD-10
Peripheral neuropathy	
Postherpetic neuralgia	B02.2x, G53.0
Diabetic neuropathy	E08.x, E09.x, E10.x, E11.x, E13.x
Other neuropathic pain	G50.0, G50.1, M60.x, G54.4, G54.6, G55.0, G55.1, G56.0, G56.2, G56.4, G56.9, G57.0, G57.1, G57.8, G57.9, G58.0, G58.7, G58.8, G62.9, G95.0, G95.2, G95.8, M48.0, M50.1, M53.0, M53.1, M54.1, M54.3, M54.4, M79.2
Fibromyalgia (Central neuropathy)	M60.8x, M60.9, M79.0, M79.1, M79.7, M79.9
Generalized anxiety disorders	F41.1,
Social anxiety disorder	F40.x

ICD-10 International Classification of Diseases Version 10

Table S2 The Anatomical Therapeutic Chemical classification system with Defined Daily Doses (ATC/DDD) [2].

Group	Name	ATC5	DDD (mg)	Administration route
	pregabalin	N03AX16*	300	Oral
Benzodiazepines	diazepam	N05BA01	10	Oral, parenteral, rectal
	oxazepam	N05BA04	50	Oral, rectal
	lorazepam	N05BA06	2.5	Oral, parenteral, sublingual
	alprazolam	N05BA12	1	Oral
	clonazepam	N03AE01	8	Oral, parenteral
Z-Drugs	zopiclone	N05CF01	7.5	Oral
	zolpidem	N05CF02	10	Oral, sublingual
Opioids	buprenorphine	N07BC51	8	Sublingual
		N02AE01	1.2	Parenteral, sublingual, transdermal
	fentanyl	N01AH01	0.7	Parenteral
		N02AB03	1.2	Transdermal
	methadone	N07BC02	25	Oral, parenteral
	morphine	N02AA01	100	Oral
		N02AA01	30	Parenteral, rectal
	oxycodone	N02AA05	75	Oral
		N02AA05	30	parenteral
	pethidine	N02AB02	400	Oral, parenteral, rectal

ATC Anatomical Therapeutic Chemical, *CHS* Clalit Health Services, *DDD* Defined Daily Dose

* In January 2023, pregabalin's ATC code was changed from N03AX16 to N02BF02. As the study data was gathered from the CHS database prior to the change in code in 2022, the old code is indicated.

Table S3 Annual pregabalin prescription rate (ages 18-34)

Year	Rate (per 100)	95% CI	Odds ratio	Chi-square	p-value [‡]
2010	0.21	0.20-0.22	Ref.		
2011	0.29	0.28-0.30	1.50	0.20	1.000
2012	0.38	0.37-0.39	1.99	0.67	0.992
2013	0.38	0.37-0.40	1.98	0.65	0.992
2014	0.46	0.45-0.48	1.97	0.64	0.993
2015	0.51	0.50-0.52	2.43	1.24	0.937
2016	0.55	0.54-0.57	2.46	1.28	0.932
2017	0.70	0.68-0.72	3.45	2.85	0.578
2018	0.85	0.83-0.86	3.93	3.74	0.389
2019	0.80	0.79-0.82	3.93	3.73	0.390

CHS Clalit Health Services, **CI** confidence interval

Rate (per 100) - Pregabalin prescriptions per 100 CHS members.

[‡] Adjusted by Sidak method [3].

Table S4 Annual pregabalin prescription rate (ages 35-54)

Year	Rate (per 100)	95% CI	Odds ratio	Chi-square	p-value [‡]
2010	1.58	1.55-1.61	Ref.		
2011	2.03	2.00-2.06	1.36	0.72	0.990
2012	2.50	2.46-2.53	1.63	1.99	0.788
2013	2.53	2.05-2.56	1.66	2.18	0.741
2014	2.82	2.79-2.85	1.84	3.32	0.472
2015	3.05	3.02-3.09	2.02	4.64	0.248
2016	3.24	3.21-3.28	2.17	5.95	0.125
2017	3.60	3.56-3.64	2.39	7.89	0.044
2018	3.85	3.82-3.89	2.60	9.95	0.014
2019	3.84	3.80-3.88	2.53	9.41	0.019

CHS Clalit Health Services, **CI** confidence interval

Rate (per 100) - Pregabalin prescriptions per 100 CHS members.

[‡] Adjusted by Sidak method [3].

Table S5 Annual pregabalin prescription rate (ages 55-74)

Year	Rate (per 100)	95% CI	Odds ratio	Chi-square	p-value [‡]
2010	6.09	6.04-6.15	Ref.		
2011	8.19	8.13-8.26	1.38	2.43	0.680
2012	10.19	10.12-10.26	1.75	8.08	0.040
2013	10.58	10.51-10.65	1.82	9.57	0.018
2014	11.53	11.45-11.60	1.20	13.28	0.002
2015	12.49	12.42-12.57	2.20	17.96	<0.001
2016	13.04	12.96-13.11	2.30	20.55	<0.001
2017	13.73	13.65-13.81	2.44	24.34	<0.001
2018	13.89	13.82-13.97	2.48	25.60	<0.001
2019	13.18	13.11-13.26	2.35	22.10	<0.001

CHS Clalit Health Services, **CI** confidence interval

Rate (per 100) - Pregabalin prescriptions per 100 CHS members.

[‡] Adjusted by Sidak method [3].

Table S6 Annual pregabalin prescription rate (ages 75+)

Year	Rate (per 100)	95% CI	Odds ratio	Chi-square	p-value [‡]
2010	10.24	10.13-10.36	Ref.		
2011	14.20	14.07-14.33	1.46	1.99	0.788
2012	17.77	17.63-17.91	1.93	6.76	0.081
2013	19.04	18.90-19.19	2.09	8.76	0.027
2014	21.03	20.88-21.18	2.35	12.41	0.004
2015	23.15	22.99-23.30	2.68	17.25	<0.001
2016	24.44	24.28-24.60	2.84	19.69	<0.001
2017	25.71	25.55-25.88	3.05	22.94	<0.001
2018	25.91	25.75-26.07	3.09	23.63	<0.001
2019	24.11	23.96-24.27	2.83	19.66	<0.001

CHS Clalit Health Services, **CI** confidence interval

Rate (per 100) - Pregabalin prescriptions per 100 CHS members.

[‡] Adjusted by Sidak method [3].

Supplements references

- [1] Classification of Diseases, 10th Revision, Clinical Modification (ICD-10-CM) n.d. <https://icd10cmtool.cdc.gov/?fy=FY2023> (accessed July 10, 2021).
- [2] ATC/DDD Index 2021 n.d. https://www.whocc.no/atc_ddd_index/ (accessed July 10, 2021).
- [3] Sidak Z. Rectangular Confidence Regions for the Means of Multivariate Normal Distributions. vol. 62. 1967.