SUPPLEMENTAL FILE 1

Codes used to identify epilepsy in the CPRD

Table S1. Codes used to identify epilepsy in the CPRD

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48134	F255100	Sensory induced epilepsy					
73542	F255300	Visceral reflex epilepsy					
98870	F255311	Partial epilepsy with autonomic symptoms					
55739	F255400	Visual reflex epilepsy					
68946	F255500	Unilateral epilepsy					
40105	F255600	Simple partial epileptic seizure					
		Partial epilepsy without impairment of					
26733	F255y00	consciousness OS					
		Partial epilepsy without impairment of					
27526	F255z00	consciousness NOS					
71719	F257.00	Kojevnikov's epilepsy					
30604	F25B.00	Alcohol-induced epilepsy					
30816	F25C.00	Drug-induced epilepsy					
56359	F25D.00	Menstrual epilepsy					
65673	F25E.00	Stress-induced epilepsy					
30635	F25F.00	Photosensitive epilepsy					
106571	F25H.00	Generalised seizure					
6271	F25X.00	Status epilepticus, unspecified					
38307	F25y.00	Other forms of epilepsy					
55260	F25y000	Cursive (running) epilepsy					
53483	F25y100	Gelastic epilepsy					
		Locl-rlt(foc)(part)idiop epilep&epilptic syn					
9887	F25y200	seiz locl onset					
25330	F25y300	Complex partial status epilepticus					
96641	F25y500	Panayiotopoulos syndrome					
9979	F25yz00	Other forms of epilepsy NOS					
9747	F25z.00	Epilepsy NOS					
3607	F25z.11	Fit (in known epileptic) NOS					
		[X]Other generalized epilepsy and epileptic					
99731	Fyu5000	syndromes					
69831	Fyu5100	[X]Other epilepsy					
59120	Fyu5200	[X]Other status epilepticus					
71801	Fyu5900	[X]Status epilepticus, unspecified					
4109	SC20000	Traumatic epilepsy					
37644	F132100	Progressive myoclonic epilepsy					

Definition of variables

Lists of Medcodes and corresponding Read codes used for each variable can be found in Supplemental file 2. An initial list of Read codes was created for each variable, either based on previous research or by searching specific words in a Read code browser. Medcodes were matched to the Read code list using the CPRD Medical Dictionary. An additional search of the Medcodes was conducted to ensure all appropriate codes were used. Medcodes were used during the analysis in STATA as these are numeric codes. For variables using medication data, lists of prodcodes were generated by searching the CPRD product code list, which lists prodcodes against British National Formulary (BNF) information. • Age: For cases: age = year of death – year of birth. For matched controls: age = year of death of the matched case – year of birth.

For epilepsy related factors (ie. Relating to seizures and treatment) and lifestyle factors, data from the CPRD was used from the date of death of the matched case up to 12 months prior. These variables are listed below:

- Emergency department visit or emergency admission: Emergency admissions or visits to emergency centres were identified from "Clinical data" from CPRD. These included keywords: seen by emergency clinician, in accident and emergency, emergency admission. A full list is available in Supplemental File 2.
- Number of antiepileptic drugs (AEDs): Number of different types of AEDs prescribed in 12 months were grouped into categories: 0, 1, 2, 3 and "4 or more". Product codes ("Prodcode") were used from "Therapy data" to identify AEDs in the CPRD. A full list of Prodcodes and corresponding definitions is available in Supplemental File 2.
- **Status epilepticus**: recorded status epilepticus code in previous 12 months (medcode: 4093; readcode: F253.11) clinical CPRD data.¹
- Depression: the variable was coded as either a documented diagnostic medcode related to "depression" from "Clinical data" or a prescription of antidepressants from "Therapy data" in the previous 12 months.¹ Antidepressants were identified through their product codes (prodcode) from therapy data. A list of medcodes for depression and prodcodes for antidepressants can be found in Supplemental File 2.
- Smoking status: Data was coded from medcodes related to smoking or smoking cessation interventions from "Clinical data" and overall smoking status if available in the CPRD "Additional Files". For any patients that did not have a smoking status recording in the 12-month data period, the last known recording from clinical data was used, as previously described by Booth et al.² The list of medcodes and readcodes used to identify smoking status can be found in Supplemental File 2.
- Injury: any injury documented in "Clinical data" was used. A list of medcodes from clinical CPRD data included: injuries, fractures, dislocations, burns, bruising, contusions, lacerations, abrasions, concussions and wounds.¹ A full list of medcodes and corresponding Read codes can be found in Supplemental File 2.
- Seizure freedom: recoded seizure freedom for > 12 months (medcode: 11015; readcode: 667F.00) in "Clinical data" from CPRD.¹

For co-morbidity, data coded for patient's lifetime (within start and end dates with acceptable quality data recording):

• **Dementia**: code list included any medcodes for "dementia" and "Alzheimer's" in patient history from "Clinical data" in the CPRD. The list of medcodes and corresponding Read codes can be found in Supplemental File 2.

• **Stroke**: code list included stroke, cerebral ischemia/arterial occlusion/haemorrhage/infarction. embolism in patient history from "Clinical data" in the CPRD. The list of medcodes and corresponding Read codes can be found in Supplemental File 2.

ICD-10 category (ICD-10 code)	2005	2000	2007	2008	2009	2010	2011	2012	2015
Frileney (CAO CA1)	(%)	(%)	(%)	(%)	(%) 5.7	(%) 6.2	(%) 5.2	(%)	(%) 5.2
Epilepsy (G40-G41)	6.3	6.3	5.9	6.1				7.1	
Certain infectious and parasitic diseases (A00-B99)	1.0	1.6	1.1	1.0	0.7	0.9	0.5	1.2	0.5
Neoplasms (C00-D48)	20.9	19.4	17.6	19.6	19.0	18.1	21.8	21.4	20.9
Diseases of the blood and blood-forming organs (D50- D89)	0.6	0.0	0.4	0.3	0.1	0.0	0.2	0.3	0.3
Endocrine, nutritional and metabolic diseases (E00-E90)	1.5	1.7	1.7	1.1	1.6	1.7	1.1	1.0	1.1
Mental and behavioural disorders (F00-F99)	4.6	3.8	5.8	5.0	6.6	4.5	10.7	10.9	10.5
Diseases of the nervous system (G00-G37, G43-G99)	4.6	5.1	5.1	6.0	7.5	5.5	7.3	5.6	7.2
Diseases of the circulatory system (I00-I99)	33.8	35.1	34.0	34.0	31.3	33.1	28.3	26.8	28.1
Diseases of the respiratory system (J00-J99)	12.9	16.3	14.7	13.6	13.0	14.5	12.6	12.7	12.5
Diseases of the digestive system (K00-KK93)	3.8	3.3	4.3	4.6	4.1	4.8	4.9	4.9	5.0
Diseases of the skin and subcutaneous tissue (L00-L99)	0.0	0.1	0.5	0.3	0.4	0.3	0.1	0.0	0.1
Diseases of the musculoskeletal system and connective tissue (M00-M99)	0.4	0.0	0.8	0.3	1.2	0.2	0.7	0.4	0.8
Diseases of the genitourinary system (N00-N99)	2.9	2.5	2.4	1.8	2.8	3.1	0.7	2.1	1.2
Congenital malformations, deformations and chromosomal abnormalities (Q00-Q99)	1.5	1.5	1.1	1.5	1.6	0.9	0.8	0.8	1.1
Symptoms, signs and abnormal clinical and laboratory findings, not elsewhere classified (R00- R99)	1.5	1.3	1.8	1.7	1.0	2.3	1.7	1.8	1.0
External causes of morbidity and mortality (V01-X59, X85- Y98)	3.3	1.6	2.6	2.8	3.1	3.1	3.2	2.6	3.5
External causes of morbidity and mortality (intentional) (X60-X84)	0.4	0.4	0.1	0.4	0.4	0.8	0.2	0.3	1.0
Total (n)	521	687	761	785	836	874	886	956	987

Table S2. Main cause of death of PWE registered in the CPRD between 2005 – 2013 by year

References

 Ridsdale L, Charlton J, Ashworth M, et al. Epilepsy mortality and risk factors for death in epilepsy: a population-based study. *Br J Gen Pract* 2011;61(586):e271-8. doi: 10.3399/bjgp11X572463
Booth HP, Prevost AT, Gulliford MC. Validity of smoking prevalence estimates from primary care electronic health records compared with national population survey data for England, 2007 to 2011. *Pharmacoepidemiol Drug Saf* 2013;22(12):1357-61. doi: 10.1002/pds.3537