

# The burden of neurological diseases in Europe: an analysis for the Global Burden of Disease Study 2017

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## Summary

**Background** Neurological disorders account for a large and increasing health burden worldwide, as shown in the Global Burden of Diseases (GBD) Study 2016. Unpacking how this burden varies regionally and nationally is important to inform public health policy and prevention strategies. The population in the EU is older than that of the WHO European region (western, central, and eastern Europe) and even older than the global population, suggesting that it might be particularly vulnerable to an increasing burden of age-related neurological disorders. We aimed to compare the burden of neurological disorders in the EU between 1990 and 2017 with those of the WHO European region and worldwide.

**Methods** The burden of neurological disorders was calculated for the year 2017 as incidence, prevalence, mortality, disability-adjusted life-years (DALYs), years of life lost, and years lived with disability for the countries in the EU and the WHO European region, totally and, separately. Diseases analysed were Alzheimer's disease and other dementias, epilepsy, headache (migraine and tension-type headache), multiple sclerosis, Parkinson's disease, brain cancer, motor neuron diseases, neuroinfectious diseases, and stroke. Data are presented as totals and by sex, age, year, location and socio-demographic context, and shown as counts and rates.

**Findings** In 2017, the total number of DALYs attributable to neurological disorders was 21·0 million (95% uncertainty interval 18·5–23·9) in the EU and 41·1 million (36·7–45·9) in the WHO European region, and the total number of deaths was 1·1 million (1·09–1·14) in the EU and 1·97 million (1·95–2·01) in the WHO European region. In the EU, neurological disorders ranked third after cardiovascular diseases and cancers representing 13·3% (10·3–17·1) of total DALYs and 19·5% (18·0–21·3) of total deaths. Stroke, dementias, and headache were the three commonest causes of DALYs in the EU. Stroke was also the leading cause of DALYs in the WHO European region. During the study period we found a substantial increase in the all-age burden of neurodegenerative diseases, despite a substantial decrease in the rates of stroke and infections. The burden of neurological disorders in Europe was higher in men than in women, peaked in individuals aged 80–84 years, and varied substantially with WHO European region and country. All-age DALYs, deaths, and prevalence of neurological disorders increased in all-age measures, but decreased when using age-standardised measures in all but three countries (Azerbaijan, Turkmenistan, and Uzbekistan). The decrease was mostly attributed to the reduction of premature mortality despite an overall increase in the number of DALYs.

**Interpretation** Neurological disorders are the third most common cause of disability and premature death in the EU and their prevalence and burden will likely increase with the progressive ageing of the European population. Greater attention to neurological diseases must be paid by health authorities for prevention and care. The data presented here suggest different priorities for health service development and resource allocation in different countries.

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## Introduction

Neurological disorders are an important source of premature mortality and transient or permanent disability in survivors. According to WHO, in 2006, neurological disorders were identified as one of the greatest public health problems, accounting for 6·3% of total disability-adjusted life-years (DALYs).<sup>1</sup> A report by the Global Burden of Disease (GBD) 2015 Neurological Disorders Collaborator Group<sup>2</sup> estimated that neurological disorders have become a leading cause of disability in the world, with attributable DALYs reaching 11·6%, and ranked second in terms of deaths after cardiovascular diseases, reaching 16·5% of all

deaths.<sup>2</sup> Given that the prevalence of most neurological disorders increases with age, their burden is expected to rise in countries with with ageing populations. In Europe, the population is rapidly ageing, therefore, this region is an important setting for the assessment of the dynamics of the burden of neurological disorders. Some European countries continue to strengthen their cooperation around health strategy and so knowledge about the similarities and disparities in the burden of neurological diseases across European countries could inform political decision making.

We aimed to calculate incidence, prevalence, mortality, DALYs, and their components (years of life lost [YLLs],

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### Research in context

#### Evidence before this study

We searched PubMed on Feb 10, 2019, and March 3, 2020, for articles focusing on the incidence, prevalence, mortality, and overall burden of neurological disorders in Europe using the search terms (“Burden of disease” OR “Epidemiology” OR “Costs”) AND (“Neurology” OR “Neurologic disease”) AND (“Europe”) with no language or time restrictions. Despite the publication of numerous studies on the frequency and outcome of neurological disorders in Europe, we found only a few and older reports on the burden of disease on a national basis or in selected age groups, and virtually no studies comparing various clinical conditions and different countries. The economic costs of disorders of the brain in Europe were calculated by the European Brain Council, but the corresponding burden was not measured. Worldwide and country-specific data on the burden of neurological disorders, in general and by type, have been provided only by the Global Burden of Diseases (GBD) studies. During the period 1990–2016, neurological disorders accounted for an increasing number of disability-adjusted life-years (DALYs). However, no data from the 27 EU countries plus the UK (EU28) were highlighted. As the population of EU28 is ageing and the prevalence of neurological disorders increases with age, the corresponding burden will increase even further in that area. Additionally, because the growth, ageing, and sociodemographic characteristics of the European population differ between the EU28 and the other countries in the larger WHO region, the burden of neurological disorders is expected to differ.

#### Added value of this study

This study updates the previous investigation covering the period 1990–2016 by providing data for the year 2017.

In the EU28, neurological disorders were the third most common source of DALYs, after cardiovascular diseases and cancer. Comparing data from the EU28 to the three WHO European regions (western, central, and eastern Europe), substantial differences were found across countries in the burden of neurological disorders, which might reflect not only the structure of the underlying populations, but also the different approaches of the local health-care facilities in the introduction of prevention and treatment measures. This difference was particularly true for neurodegenerative diseases for which robust preventive measures and disease-modifying treatments are not available. During the study period, the burden of headache was unchanged, whereas the burden of stroke decreased and the burden of Alzheimer’s disease and other dementias increased, but at different paces in the EU28 and the WHO European regions.

#### Implications of the available evidence

Because of the ageing of the European population, the EU28 is facing an increasing burden attributable to neurological disorders, presenting the governments with an increasing demand for acute care, rehabilitation, and support services. The growing numbers of affected individuals and the corresponding numbers of neurological DALYs and deaths should call for action to address these diseases, particularly neurodegenerative diseases that are still in need of robust preventive measures and disease-modifying treatments. The results of this study can be used by the health authorities and local governments to consider the burden of neurological disorders that could be addressed with preventive and therapeutic measures.

and years lived with disability [YLDs]) as absolute numbers and rates attributable to neurological disorders in the so-called EU28 (the 27 member countries of the EU plus the UK), in general and by age, sex, year, and location for the period 1990–2017, and to compare the EU28 with the WHO Europe region and the rest of the world.

## Methods

### Overview

The GBD Collaborator Group is a network of scientists producing estimates of the burden of 328 diseases and injuries including 13 neurological disorders (stroke, Alzheimer’s disease and other dementias, Parkinson’s disease, epilepsy, multiple sclerosis, brain and nervous system cancer, encephalitis, meningitis, tetanus, headache [migraine and tension-type headache], motor neuron diseases, and other neurological diseases). The guiding principle of GBD is to assess health loss due to premature mortality and disability, intended as any deviation from full health. The methodological basis of the GBD studies is described in detail elsewhere.<sup>2–4</sup>

The analyses in the current study follow the GBD methodology. The data provided here are for 1990–2017.

Population-based measures are taken from the related GBD standards.<sup>5</sup> For the purposes of this study, data on GBD outcomes from the countries included in the EU28 were isolated and compared with the WHO European region and with global data. According to WHO, the European region includes, in addition to the 28 European countries that were part of the EU28 in 2017, another 22 countries (appendix p 9). WHO divides the European region into three subregions: western, central, and eastern Europe (appendix p 15).

In our analysis we included 13 neurological disorders. We chose to exclude traumatic brain injury and spinal cord injury because post-traumatic disorders were outside the scope of our investigation. The burden of neurological disorders was calculated as incidence, prevalence, mortality, DALYs, YLLs, and YLDs, totally and, separately, for each disorder.

### DALYs, YLLs, and YLDs

To compare deaths and non-fatal outcomes within and between diseases, a unique indicator was used, DALYs, which is the sum of YLLs and YLDs. YLLs are the product of the number of deaths multiplied by the standard life

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expectancy at the age of death. Standard life expectancy is obtained from the lowest observed age-specific rates of mortality among populations in the world greater than 5 million.<sup>6</sup> YLDs are the product of the prevalence of individual sequelae of each disease multiplied by a disability weight, quantifying the severity of a sequela as a number between 0 (representing full health) and 1 (representing death). Disability weights were estimated in nine US population surveys and an open-access internet survey in which respondents were asked to choose the healthier option between random pairs of health states that were presented with a short description of the main features.<sup>7</sup>

Non-fatal estimates were obtained from systematic reviews of published and unpublished reports, records of health claims, survey microdata, registries, and disease surveillance systems. These data have been included in a single repository, the Global Health Data Exchange.<sup>8</sup> Non-fatal data were analysed using DisMod-MR 2.1, a Bayesian meta-regression tool that adjusts datapoints for variations in study methods among different sources and enforces consistency between data obtained from different measures, such as incidence and prevalence. A different model was used for brain cancer according to the respective GBD analysis.<sup>9</sup> For each disease, a parsimonious set of sequelae was selected that best described the different aspects of the various consequences of a disease. Each sequela is estimated separately (appendix).

### Mortality estimates

All-cause mortality rates were estimated from vital registration data in countries with complete coverage. For other countries, the probabilities of death before age 5 years and between ages 15 years and 60 years were estimated from verbal autopsies. Using model life tables, these probabilities were converted into age-specific death rates by sex, year, and location. The cause of death was classified using subsequent versions of the International Classification of Diseases (or bespoke classifications in some countries). When coded data were less informative or in the absence of a specific diagnosis, causes of death were redistributed to more precise codes.<sup>6</sup> Each death was assigned a single underlying cause. The mortality attributable to each cause is the product of the attributable fraction and the mortality due to the underlying disease. Causes of death were analysed using the cause of death ensemble model, a highly systematised tool running many different models on the same data and choosing an ensemble of models that best reflects the available input data. To enforce consistency, the sum of all cause-specific mortality rates was scaled to all-cause mortality rates in each category defined by age, sex, location, and year.

### Socio-demographic index

Data were stratified by socio-demographic index (SDI), a composite indicator of sociodemographic development

reflecting the geometric mean of normalised values of a location's income per capita, the average number of years of educational attainment in the population aged 15 years and over, and the total fertility rate. Countries and territories were grouped into five quintiles of high, high-middle, middle, low-middle, and low SDI on the basis of their 2017 values.<sup>6</sup>

DALYs were computed for the entire world, for the EU28, WHO Europe, the three WHO European subregions, and for each European country separately, and analysed by age, sex, and year.

With reference to the measure of sociodemographic development, European countries were stratified into two groups: high-SDI countries (Andorra, Austria, Belgium, Croatia, Cyprus, Czech Republic, Denmark, Estonia, Finland, France, Germany, Greece, Iceland, Ireland, Italy, Latvia, Lithuania, Luxembourg, Malta, the Netherlands, Norway, Poland, Russia, Slovakia, Slovenia, Sweden, Switzerland, and the UK) and high-middle-SDI countries (Armenia, Azerbaijan, Belarus, Bulgaria, Georgia, Hungary, Israel, Kazakhstan, Macedonia, Montenegro, Portugal, Romania, Serbia, Spain, Turkey, and Ukraine). Only Moldova was classified as a middle-SDI country.

### Standardisation and uncertainty measures

Unless otherwise specified, all rates were age-standardised using the GBD standard (appendix p 45). The uncertainty of all estimates was propagated throughout all the calculations by creating 1000 values for each estimate of the burden and aggregating across causes and locations at the level of each of the 1000 values for all intermediate steps in the calculation. The lower and upper bounds of the 95% uncertainty intervals (UIs) were the 25th and 975th values of the ordered 1000 values. Significant differences were established if 975 or more of the ordered 1000 values of difference were on either side of zero. Definitions, imputation procedures and modelling strategies for major neurological disorders are described in the appendix (pp 47–147).

### Role of the funding source

The funder of the study had no role in study design, data collection, data analysis, data interpretation, or writing of the report. EB and GD had access to all raw data. The corresponding author had full access to all the data in the study and had final responsibility for the decision to submit for publication.

### Results

In 2017, the total population in the EU28 was 512.4 million and the population of WHO Europe region was 925.6 million. In the same year, 307.9 million neurological diseases were counted in the EU28, 74.5 million of which were newly diagnosed (table 1). In the WHO Europe region 540.3 million neurological diseases were counted, 133.0 million of which were

newly diagnosed (table 1). The total number of DALYs in patients with neurological disorders was 21·0 million (95% UI 18·5–23·9) in the EU28 and 41·1 million (36·7–45·9) in the WHO European region and the total number of deaths was 1·1 million (1·09–1·14) in

the EU28 and 1·97 million (1·95–2·01) in the WHO European region (table 1). In 2017, the worldwide burden of neurological disorders accounted for 280·0 million (251·4–312·2) DALYs. In the EU28, neurological disorders ranked third for DALYs (after cancer and

	EU28		WHO European region		Global	
	Total	Age standardised rate per 100 000	Total	Age-standardised rate per 100 000	Total	Age-standardised rate per 100 000
<b>All neurological diseases</b>						
DALYs	21 046 899 (18 499 198–23 863 193)	190 (160–225)	41 103 486 (36 655 558–45 935 037)	232 (201–266)	279 895 042 (251 440 223–312 210 489)	278 (250–310)
Deaths	1 116 038 (1 094 328–1 142 613)	8 (8–8)	1 974 840 (1 947 870–2 012 204)	10 (10–10)	9 927 123 (9 771 333–10 100 744)	12 (12–12)
Prevalence	307 859 199 (291 599 989–327 023 248)	4 137 (3 900–4 395)	540 277 716 (510 725 114–573 264 505)	4 089 (3 856–4 350)	3 868 780 576 (3 644 294 271–4 129 974 156)	3 791 (3 569–4 049)
YLDs	8 622 940 (6 072 621–11 333 906)	99 (69–133)	15 213 540 (10 755 558–20 029 063)	101 (71–135)	93 483 900 (65 716 638–125 511 962)	92 (64–123)
YLLs	12 423 960 (12 149 237–12 714 922)	111 (109–114)	25 889 946 (25 502 381–26 430 347)	159 (157–163)	186 411 142 (182 252 461–190 710 446)	237 (231–244)
<b>Stroke</b>						
DALYs	7 314 978 (6 833 270–7 768 518)	691 (643–737)	18 831 007 (17 874 242–19 733 892)	1198 (1136–1256)	132 051 366 (126 498 600–137 350 169)	1657 (1587–1724)
Deaths	476 903 (463 602–504 651)	38 (37–40)	1 078 620 (1 058 760–1 117 505)	61 (60–63)	6 167 292 (6 044 260–6 327 598)	80 (79–83)
Prevalence	9 531 098 (8 958 668–10 024 854)	971 (921–1 021)	18 254 959 (17 183 892–19 283 541)	1196 (1 129–1 262)	104 178 740 (98 453 978–110 124 977)	1 301 (1 229–1 375)
Incidence	1 102 677 (1 030 004–1 178 489)	107 (101–115)	2 060 660 (1 916 873–2 206 398)	131 (123–141)	11 931 061 (11 118 416–12 825 770)	151 (140–162)
YLDs	1 621 402 (1 191 963–2 036 972)	162 (118–204)	3 304 779 (2 428 732–4 165 663)	215 (157–270)	18 695 419 (13 574 330–23 686 932)	236 (171–299)
YLLs	5 693 577 (5 543 620–5 958 058)	530 (516–552)	15 526 228 (15 235 982–16 036 397)	983 (965–1 016)	113 355 949 (110 957 795–116 180 577)	1 423 (1 392–1 458)
<b>Alzheimer's disease and other dementias</b>						
DALYs	5 271 654 (4 915 486 to 5 641 197)	399 (372–427)	7 686 392 (7 175 157–8 205 020)	410 (383–438)	30 521 481 (28 529 663–32 557 921)	413 (386–440)
Deaths	501 678 (489 118–512 705)	35 (34–36)	700 353 (685 974–712 101)	36 (35–36)	2 514 619 (2 470 512–2 550 346)	35 (35–36)
Prevalence	7 740 481 (6 896 189–8 583 917)	594 (531–657)	11 375 859 (10 038 137–12 683 497)	612 (541–681)	44 988 790 (39 716 207–50 377 809)	605 (533–676)
Incidence	1 249 178 (1 118 954–1 390 849)	98 (88–108)	1 851 062 (1 653 033–2 065 251)	101 (90–112)	7 300 645 (6 515 684–8 133 403)	97 (87–109)
YLDs	1 165 391 (832 669–1 516 059)	88 (63–114)	1 694 270 (1 209 023–2 212 896)	90 (64–118)	6 570 378 (4 678 144–8 588 469)	89 (64–117)
YLLs	4 106 264 (3 982 651–4 213 014)	311 (301–320)	5 992 121 (5 844 318–6 113 018)	320 (312–327)	23 951 105 (23 523 645–24 326 849)	324 (318–329)
<b>Parkinson's disease</b>						
DALYs	839 916 (766 256–940 813)	70 (63–78)	1 226 579 (1 126 744–1 363 923)	70 (64–78)	5 580 212 (5 151 586–6 084 655)	72 (67–79)
Deaths	60 812 (55 723–67 973)	5 (4–5)	84 599 (80 948–93 079)	5 (4–5)	340 639 (324 378–355 089)	5 (4–5)
Prevalence	1 215 045 (996 237–1 457 870)	105 (86–125)	1 838 098 (1 506 704–2 220 458)	107 (88–128)	8 525 404 (7 037 337–10 185 559)	109 (90–131)
incidence	150 667 (124 575–180 761)	13 (11–16)	225 529 (185 864–271 669)	13 (11–16)	1 025 939 (854 090–1 229 873)	13 (11–16)
YLDs	1 709 916 (1 176 42–2 333 979)	15 (10–20)	257 986 (178 581–354 467)	15 (10–21)	1 218 972 (823 711–1 662 213)	16 (11–22)
YLLs	669 000 (612 867–764 328)	55 (51–63)	968 593 (924 800–1 088 287)	55 (52–62)	4 361 241 (4 182 792–4 578 722)	57 (55–60)

(Table 1 continues on next page)

	EU28		WHO European region		Global	
	Total	Age-standardised rate per 100 000	Total	Age-standardised rate per 100 000	Total	Age-standardised rate per 100 000
(Continued from previous page)						
<b>Motor neuron disease</b>						
DALYs	252 987 (240 153–264 173)	31 (29–32)	314 523 (298 988–348 088)	24 (23–26)	878 518 (841 386–966 073)	11 (10–12)
Deaths	11 495 (10 900–12 020)	1 (1–1)	13 410 (12 728–14 790)	1 (1–1)	34 068 (32 796–37 053)	0 (0–0)
Prevalence	42 937 (38 970–47 145)	6 (5–7)	51 711 (46 991–56 707)	4 (4–5)	237 053 (211 191–264 106)	3 (3–3)
Incidence	13 010 (12 311–13 767)	1 (1–2)	15 728 (14 766–16 723)	1 (1–1)	67 322 (60 666–74 311)	1 (1–1)
YLDs	9131 (6492–12 227)	1 (1–2)	10 998 (7 822–14 720)	1 (1–1)	50 416 (35 744–67 546)	1 (1–1)
YLLs	243 856 (231 168–254 703)	29 (28–31)	303 526 (287 735–337 104)	23 (22–25)	828 102 (796 699–917 135)	11 (10–12)
<b>Multiple sclerosis</b>						
DALYs	282 765 (233 418–328 839)	39 (32–46)	391 078 (327 333–453 473)	32 (26–37)	1 084 757 (942 878–1 237 344)	13 (12–15)
Deaths	5 601 (3954–6056)	1 (0–1)	7 413 (5673–8395)	1 (0–1)	20 655 (17 721–22 238)	0 (0–0)
Prevalence	522 674 (473 413–579 422)	77 (69–85)	699 238 (635 288–772 355)	59 (53–65)	1 761 078 (1 598 226–1 947 909)	22 (20–24)
Incidence	11 847 (10 861–13 042)	3 (2–3)	17 544 (16 068–19 308)	2 (2–2)	54 895 (50 054–60 812)	1 (1–1)
YLDs	132 159 (95 038–171 558)	20 (14–26)	178 020 (127 760–230 616)	15 (11–20)	456 556 (327 739–595 013)	6 (5–8)
YLLs	150 607 (110 315–166 542)	19 (14–22)	213 059 (169 517–254 322)	17 (14–20)	628 202 (563 020–682 386)	8 (7–9)
<b>Brain and nervous system cancer</b>						
DALYs	964 964 (789 495–1 033 050)	140 (117–153)	1 668 047 (1 439 524–1 799 509)	147 (130–163)	8 744 765 (7 652 109–9 554 150)	112 (98–123)
Deaths	35 917 (27 137–38 316)	4 (3–4)	56 246 (45 620–59 326)	4 (3–4)	247 143 (212 969–265 001)	3 (3–3)
Prevalence	330 074 (270 005–360 630)	55 (46–62)	462 982 (398 536–511 728)	45 (40–52)	1 705 702 (1 470 989–1 894 771)	22 (19–24)
Incidence	70 026 (54 549–75 620)	10 (8–11)	102 219 (86 050–109 460)	9 (8–10)	405 218 (351 030–442 624)	5 (4–6)
YLDs	30 368 (21 160–40 887)	5 (3–6)	43 489 (31 024–57 873)	4 (3–5)	166 947 (117 460–222 977)	3 (2–3)
YLLs	934 596 (766 157–997 931)	135 (113–148)	1 624 558 (1 404 505–1 753 504)	143 (127–158)	8 577 818 (7 527 040–9 359 303)	110 (97–121)
<b>Meningitis</b>						
DALYs	84 550 (77 767–97 340)	20 (18–24)	288 614 (267 659–324 772)	38 (35–43)	20 370 870 (17 800 140–23 356 670)	293 (254–337)
Deaths	2033 (1947–2224)	0 (0–0)	5 483 (5 253–6 125)	1 (1–1)	288 021 (254 287–333 240)	4 (4–5)
Prevalence	183 151 (153 769–216 836)	33 (28–39)	505 430 (430 502–591 790)	53 (45–62)	10 572 886 (8 836 735–12 552 236)	139 (116–165)
Incidence	92 542 (82 041–105 206)	22 (19–25)	232 423 (205 684–266 898)	30 (26–35)	5 045 411 (4 435 073–5 877 841)	71 (62–83)
YLDs	17 079 (12 032–22 999)	4 (3–5)	45 301 (31 738–60 745)	5 (4–7)	933 935 (652 964–1 255 126)	13 (9–17)
YLLs	67 471 (64 007–80 231)	17 (16–21)	243 313 (230 129–280 048)	33 (31–39)	19 436 936 (16 935 143–22 335 832)	281 (244–324)

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	EU28		WHO European region		Global	
	Total	Age-standardised rate per 100 000	Total	Age-standardised rate per 100 000	Total	Age-standardised rate per 100 000
(Continued from previous page)						
<b>Encephalitis</b>						
DALYs	57 720 (50 302–63 183)	11 (10–12)	270 736 (236 953–292 768)	34 (30–38)	5 112 280 (4 541 340–5 764 967)	71 (63–79)
Deaths	1 729 (1 337–1 829)	0 (0–0)	5 953 (4 631–6 266)	1 (0–1)	92 370 (83 134–107 936)	1 (1–1)
Prevalence	142 435 (74 374–234 773)	22 (12–35)	315 202 (173 810–507 091)	29 (17–46)	6 724 882 (3 731 161–10 760 385)	86 (48–137)
Incidence	47 411 (46 409–48 414)	9 (9–9)	110 769 (108 844–112 873)	12 (12–12)	2 220 535 (2 289 100–2 225 245)	30 (29–30)
YLDs	9 372 (6 386–12 593)	2 (1–2)	22 773 (15 651–30 309)	3 (2–3)	524 114 (365 488–691 266)	7 (5–9)
YLLs	48 348 (41 387–51 476)	10 (8–11)	247 963 (214 185–269 172)	33 (28–36)	4 588 167 (4 059 515–5 230 709)	65 (57–73)
<b>Tetanus</b>						
DALYs	649 (558–755)	0 (0–0)	2 127 (1 860–2 770)	0 (0–0)	2 449 433 (1 736 526–3 201 172)	35 (25–46)
Deaths	39 (33–46)	0 (0–0)	82 (72–100)	0 (0–0)	38 134 (25 893–48 771)	1 (0–1)
Prevalence	4 (3–5)	0 (0–0)	160 (133–187)	0 (0–0)	59 583 (56 726–62 572)	1 (1–1)
Incidence	68 (58–82)	0 (0–0)	164 (140–205)	0 (0–0)	79 192 (53 360–105 261)	1 (1–1)
YLDs	1 (0–1)	0 (0–0)	4 (3–7)	0 (0–0)	1 695 (1 065–2 541)	1 (1–1)
YLLs	649 (558–754)	0 (0–0)	2 122 (1 856–2 765)	0 (0–0)	2 447 740 (1 734 885–3 199 044)	36 (25–47)
<b>Epilepsy</b>						
DALYs	781 549 (512 4583–1 168 936)	140 (90–211)	1 479 134 (1 063 428–2 050 641)	158 (113–220)	14 793 945 (11 417 678–18 991 487)	196 (151–252)
Deaths	9 526 (7 286–10 097)	1 (1–1)	15 370 (12 663–16 443)	1 (1–1)	130 237 (116 998–150 774)	2 (2–2)
Prevalence	2 244 668 (1 496 976–2 977 225)	399 (266–529)	3 742 828 (2 718 716–4 823 770)	385 (280–497)	27 288 268 (21 576 010–33 443 826)	359 (284–441)
Incidence	184 947 (124 646–244 171)	37 (25–50)	309 286 (221 551–399 256)	35 (25–46)	2 470 759 (1 905 450–3 062 903)	33 (26–41)
YLDs	523 003 (259 741–908 273)	96 (48–166)	942 629 (528 597–1 512 164)	100 (57–160)	8 561 880 (5 380 607–12 551 498)	114 (72–166)
YLLs	258 546 (219 592–274 228)	44 (40–47)	536 505 (490 237–585 052)	58 (53–64)	6 232 066 (5 709 753–7 289 706)	83 (76–97)
<b>Migraine</b>						
DALYs	4 231 019 (2 740 462–6 048 715)	770 (495–1115)	7 401 471 (4 810 162–10 539 348)	735 (473–1056)	47 245 390 (29 986 692–68 669 318)	597 (378–866)
Prevalence	112 207 672 (104 739 016–119 973 469)	20 646 (19 234–22 225)	195 794 407 (182 677 431–209 186 190)	19 650 (18 270–21 082)	1 331 364 642 (1 237 219 585–1 433 357 249)	16 828 (15 638–18 119)
Incidence	7 506 784 (6 829 187–8 173 362)	1 724 (1 568–1 883)	13 669 490 (12 467 555–14 861 052)	1 641 (1 493–1 789)	112 933 549 (102 829 920–122 899 843)	1 477 (1 344–1 607)
YLDs	4 231 019 (2 740 461–6 048 715)	770 (495–1115)	7 401 471 (4 810 162–10 539 348)	735 (473–1056)	47 245 391 (29 986 693–68 669 319)	597 (379–866)

(Table 1 continues on next page)

cardiovascular diseases) and deaths (after cardiovascular diseases and cancer). These diseases comprised 13.4% (10.3–17.1) of global DALYs and 19.5% (18.0–21.3) of deaths. In the WHO European region, neurological disorders ranked third for DALYs and deaths (after cardiovascular diseases and neoplasms). The proportion

of total DALYs attributable to neurological disorders was 13.1% (12.8–13.15) and the proportion of deaths was 19.0% (19.0–19.1). The percentage of DALYs attributable to neurological disorders was lowest in western Europe (14.0% [12.7–15.2]) and highest in central Europe (15.1% [13.9–16.2]). The proportion of

	EU28		WHO European region		Global	
	Total	Age-standardised rate per 100 000	Total	Age-standardised rate per 100 000	Total	Age-standardised rate per 100 000
(Continued from previous page)						
<b>Tension-type headache</b>						
DALYs	600 667 (343 433–936 239)	101 (57–159)	1 163 732 (664 951–1 799 904)	110 (63–173)	7 096 415 (4 044 590–11 213 525)	90 (51–142)
Prevalence	173 696 347 (158 485 543–190 762 959)	30 871 (28 014–33 930)	309 890 284 (281 758 135–340 323 834)	31 022 (28 086–34 097)	2 331 334 677 (2 110 373 311–2 575 461 450)	29 810 (27 057–32 943)
Incidence	63 611 672 (57 147 715–69 565 055)	11 725 (10 423–12 971)	114 411 512 (102 667 592–125 554 761)	11 805 (10 490–13 036)	882 441 388 (783 241 228–975 064 340)	11 371 (10 058–12 591)
YLDs	600 667 (343 433–936 239)	101 (57–159)	1 163 732 (664 951–1 799 904)	110 (63–173)	7 096 415 (4 044 591–11 213 526)	90 (51–143)
<b>Other neurological disorders</b>						
DALYs	363 480 (314 552–426 877)	65 (54–79)	599 059 (517 397–698 284)	63 (54–76)	3 965 609 (3 265 588–4 845 474)	53 (43–65)
Deaths	10 303 (9 219–10 814)	1 (1–1)	14 052 (12 762–14 637)	1 (1–1)	53 946 (51 552–59 030)	1 (1–1)
Prevalence	2613 (1 736–3 517)	0 (0–1)	4364 (2 939–5 868)	0 (0–1)	38 870 (25 596–53 211)	1 (0–1)
Incidence	..	0 (0–0)	..	0 (0–0)	..	0 (0–0)
YLDs	112 433 (68 271–174 266)	23 (13–37)	206 737 (130 672–304 277)	24 (15–36)	1 961 790 (1 276 347–2 823 819)	27 (17–38)
YLLs	251 047 (232 719–264 643)	42 (39–45)	392 321 (369 776–410 145)	40 (37–42)	2 003 820 (1 856 755–2 269 519)	27 (25–31)

Data are n (95% UI). DALYs=disability-adjusted life-years. EU28=the 27 EU countries plus the UK. YLDs=years lived with disability. YLLs= years of life lost. UI=uncertainty interval.

**Table 1: Comparison of DALYs, deaths, prevalence, YLDs, and YLLs for the EU28, WHO European region, and globally for all neurological diseases and each subcategory separately in 2017**

deaths attributable to neurological disorders was 21·8% (21·6–22·1) in eastern Europe, 22·6% (22·3–23·0) in central Europe, and 21·46% (21·2–21·9) in western Europe (appendix p 16).

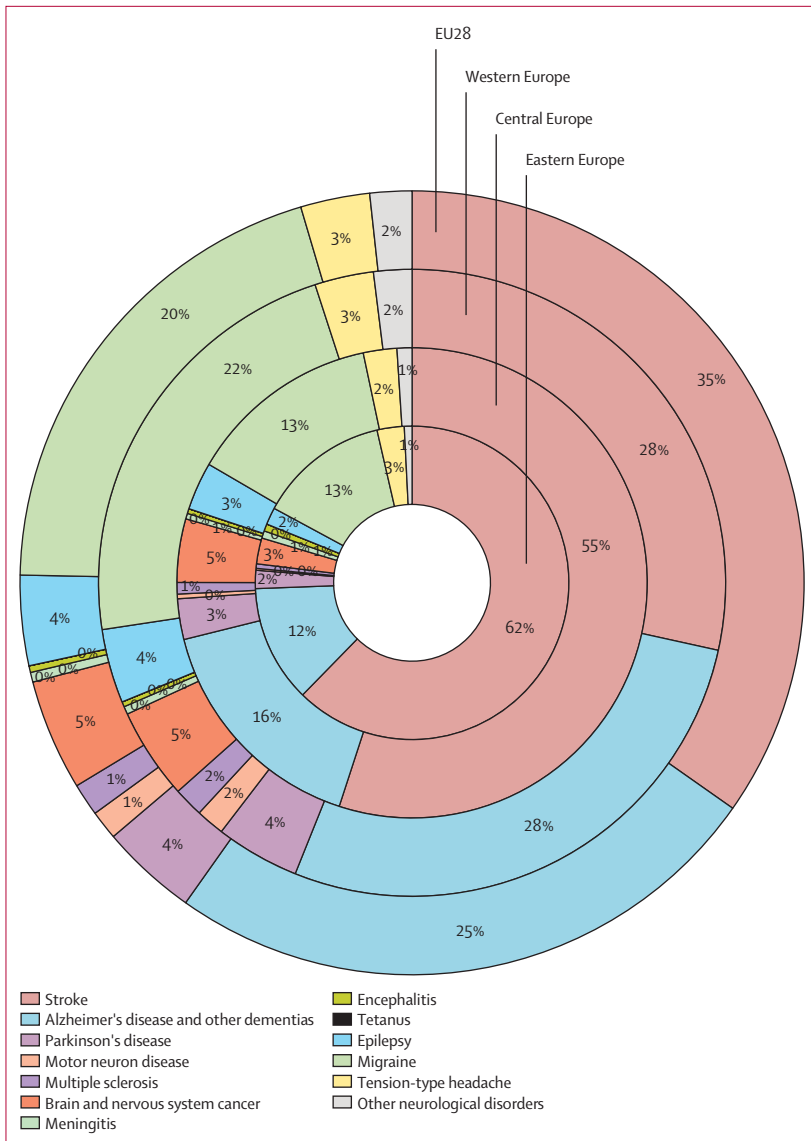
Stroke (7·3 million [95% UI 6·7–7·9]; 35% [35–38]), dementia (5·3 million [4·9–5·6]; 25% [23–28]), and headache (migraine and tension-type headache combined; 4·8 million [3·1–7·1]; 23% [19–29]) were, in decreasing order, the three commonest sources of DALYs in the EU28 (table 1; figure 1; appendix p 17). Stroke also was the leading source of burden attributable to neurological disorders in the WHO European region, accounting for the largest proportion of DALYs (18·3 million [17·9–19·7]; 45·6% [42·0–49·0]), followed by headache (8·6 million [5·4–12·6]; 20·6% [14·7–27·2]) and dementia (7·7 million, 7·1–8·3); 18·7% [17·3–20·2]). In western Europe, the leading contributor was stroke (28% [25·5–31·5]; figure 1), which accounted for 4·7 million DALYs (4·4–5·0). Stroke also predominated in central Europe (55% [50·3–59·5]) with 3·3 million DALYs (3·1–3·5) and eastern Europe (62% [57·3–66·7]) with 8·2 million (7·8–8·5). All the other neurological disorders accounted for lower numbers and rates (appendix p 17).

In the EU28, the total number of DALYs was higher in women (12·0 million [9·9–14·5]) than in men (9·1 million [7·6–10·8]). The major contributors to the burden of

neurological disorders were dementia and stroke, the attributable proportions of which increased substantially with age both for the number of DALYs and for DALY rates (figure 2). We found substantial sex differences in the number of DALYs and in age-standardised DALY rates in all age groups for dementia, migraine, and multiple sclerosis (that were higher in women) and for stroke and Parkinson's disease (that were higher in men; appendix p 10). DALYs and DALY rates for epilepsy and brain cancer were also higher in men than in women. Findings were similar in the WHO European region for both sexes and for women and men separately (appendix p 12). In the EU28 the peak in the number of DALYs and DALY rates was in people aged 80–84 years for both sexes. By contrast, in the WHO European region the burden peaked in women aged 80–84 years and in men aged 75–79 years.

Dementia, stroke, and headache differed when comparing findings in western, central, and eastern Europe. In western Europe the burden was predominantly represented by dementia and stroke after age 70 years in both sexes, whereas stroke was the strongest contributor in those aged 40–89 years in central Europe and 50–89 years in eastern Europe (appendix p 12).

The changes in all-age and age-standardised DALY rates of neurological disorders in the EU28 and WHO European region between 1990 and 2017 are shown in



**Figure 1: Contribution of each disease to the overall burden of neurological disorders in the EU28, western, central, and eastern Europe in 2017**  
 Percentages represent proportion of DALYs. DALYs=disability-adjusted life-years. EU28=the 27 countries in the EU plus the UK.

table 2, and for global comparisons, in the appendix (p 19). In the EU28, dementia accounted for a 63% (95% UI 58 to 66) increase in DALYs and a 93% (88 to 97) increase in deaths. Similar changes were found for Parkinson's disease (61% increase [31 to 69]) and motor neuron disease (60% increase [49 to 69]), for which we found a 91% (75 to 102) increase in deaths. Modest increases in DALYs and other measures were observed for multiple sclerosis and, to a lesser extent, migraine and tension-type headache (appendix p 19). By contrast, a 30% decrease (−27 to −33) in stroke DALYs was observed despite a 25% (20 to 30) increase in prevalence and a 28% (22 to 34) increase in YLDs. A significant decrease in all measures

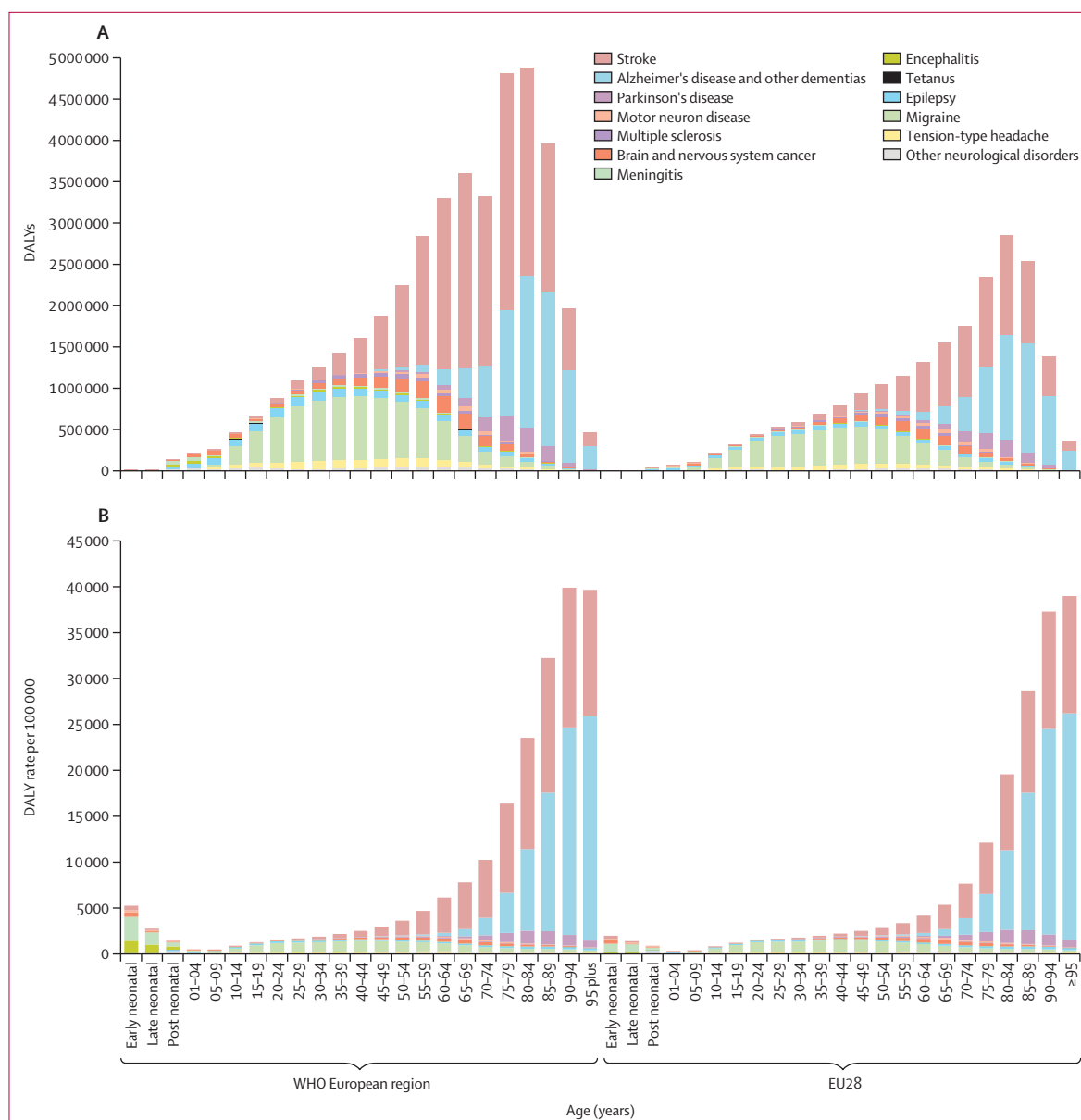
of burden was also observed for encephalitis, meningitis, and tetanus (appendix p 19). We found a 53% increase (3 to 64) in deaths for epilepsy and 43% (0 to 54) in deaths for brain cancers. Brain cancers showed a 161% (98 to 192) increase in prevalence. A 41% (39 to 44) increased prevalence was also found for multiple sclerosis. With some exceptions, in the EU28 all measures differed only slightly from those in the WHO European region but significantly from the global estimates (appendix p 19).

In the EU28 and WHO European region, age-standardised DALY rates showed a significant decrease for stroke and infections and a modest, but still significant, decrease for dementia. We found a significant increase only for motor neuron disease in the EU28 (table 2). Findings were similar for deaths (appendix p 19), whereas prevalence (all age and age standardised) increased for all clinical conditions apart from headache, although at a variable speed between regions (appendix p 19).

In 1990, in high-SDI European countries stroke was the leading disease, accounting for 1286 age-standardised DALYs per 100 000 (95% UI 1228 to 1341), followed by migraine (676 [433 to 975]), and dementia (430 [403 to 458]; tables 3). Migraine, stroke, and dementia ranked first, second, and third in 2017, but while the age-standardised DALY rate for stroke was substantially lower in comparison to 1990 levels, at 650 (597 to 703; 49% decrease [−52 to −47]), the age-standardised DALY rate for migraine and dementia remained similar at 674 (431 to 970; unchanged) and 399 (373 to 426; 7% decrease; −9 to −5; table 3), respectively. In high-middle-SDI countries, stroke and dementia ranked first and third both in 1990 and in 2017, but the age-standardised rate of stroke decreased by 37% (−40 to −35) from 3196 (3097 to 3294) in 1990, while the decrease in the age-standardised rate of dementia was only 6% (−8 to −3; table 3). Epilepsy showed a 6% (−23 to 14) non-significant decrease in high-SDI countries and a 27% (−42 to −9) decrease in high-middle-SDI countries. Notably, the disease which showed the largest increase of prevalence and DALY rates, but only in high-SDI countries, was motor neuron disease (11% [6 to 16]). Parkinson's disease was next, showing a non-significant increase (6% [−16 to 11]) in high-SDI countries but a significant decrease (−4% [−10 to 0]) in high-middle SDI countries).

During the study period, all-age DALY numbers, deaths, and prevalence of neurological disorders varied across countries (appendix p 27), with increases in Albania, Andorra, Armenia, Azerbaijan, Belarus, Bosnia and Herzegovina, Bulgaria, Georgia, Greece, Lithuania, Macedonia, Malta, Moldova, Montenegro, Romania, Russia, Serbia, Turkmenistan, Ukraine, and Uzbekistan, and no change or decrease in the other countries. By contrast, age-standardised DALY and death rates decreased in all WHO European region countries, except for Azerbaijan, Turkmenistan, and Uzbekistan, where both measures increased, and Albania and Bosnia and





**Figure 2: EU28 and WHO European region DALYs**

Total number of DALYs (A) and age-standardised DALY rates (B) by age group including both sexes in 2017. DALYs=disability-adjusted life-years. EU28=the 27 countries in the EU plus the UK.

Herzegovina, where death rates were unchanged. We found a modest increase in age-standardised prevalence rates of neurological disorders in Azerbaijan, Bosnia and Herzegovina, Lithuania, Montenegro, the Netherlands, and Switzerland. We found an almost universal decrease in standardised DALY rates in all countries (maximally represented by a reduction in YLLs and, only to a lesser extent, by a reduction in YLDs), the only exceptions being Azerbaijan, Turkmenistan, and Uzbekistan. By contrast, except for Estonia, Latvia and Ukraine, where a decrease of all raw numbers was also observed, prevalent cases, deaths, DALYs, YLLs, and YLDs all increased in

Azerbaijan, Cyprus, Finland, France, Greece, Iceland, Israel, Kazakhstan, Macedonia, Malta, Montenegro, the Netherlands, Tajikistan, Turkmenistan, and Uzbekistan (appendix p 27).

## Discussion

In 2017, neurological disorders were among the most common sources of health loss in the EU28 and the WHO European region. Although globally neurological disorders were the second largest contributor to DALYs and the second largest cause of death, in both European regions neurological disorders ranked third for DALYs and deaths.

	EU28			WHO European region								
	Age-standardised rate, 1990	Age-standardised rate, 2017	Change from 1990 to 2017	Total, 1990	Total, 2017	Change from 1990 to 2017	Age-standardised rate, 1990	Age-standardised rate, 2017	Change from 1990 to 2017	Total, 1990	Total, 2017	Change from 1990 to 2017
<b>Stroke</b>												
DALYs	1489 (1434 to 1538)	691 (643 to 737)	-54% (-56 to -52)	10 446 307 (10 075 476 to 10 789 349)	7 314 978 (6 833 270 to 7 768 518)	-30% (-33 to -27)	2012 (1946 to 2075)	1198 (1136 to 1256)	-40% (-42 to -39)	21 991 312 (21 276 861 to 22 669 254)	18 831 007 (17 874 242 to 19 733 892)	-14% (-17 to -12)
Deaths	86 (85 to 87)	38 (37 to 40)	-57% (-58 to -54)	619 264 (611 400 to 627 766)	476 903 (463 602 to 504 651)	-23% (-25 to -19)	112 (111 to 113)	61 (60 to 63)	-45% (-47 to -44)	1 207 823 (1 195 003 to 1 225 524)	1 078 620 (1 058 760 to 1 117 505)	-11% (-13 to -7)
Prevalence	1117 (1073 to 1167)	971 (921 to 1021)	-13% (-16 to -10)	7 603 547 (7 294 082 to 7 947 767)	9 531 098 (8 958 668 to 10 024 854)	25% (20 to 30)	1285 (1236 to 1345)	1196 (1129 to 1262)	-7% (-10 to -4)	13 870 323 (13 323 695 to 14 529 934)	18 254 959 (17 183 892 to 19 283 541)	32% (27 to 36)
YLDs	185 (135 to 232)	162 (118 to 204)	-13% (-16 to -9)	1 266 025 (925 645 to 1 584 998)	1 621 402 (1 191 963 to 2 036 972)	28% (22 to 34)	229 (167 to 287)	215 (157 to 270)	-6% (-9 to -4)	2 475 049 (1 797 028 to 3 105 300)	3 304 779 (2 428 732 to 4 165 663)	34% (29 to 38)
YLLs	1304 (1284 to 1321)	530 (516 to 552)	-59% (-61 to -58)	9 180 282 (9 044 047 to 9 307 537)	5 693 577 (5 543 620 to 5 958 058)	-38% (-40 to -35)	1783 (1761 to 1813)	983 (965 to 1016)	-45% (-46 to -43)	19 516 263 (19 283 622 to 19 856 420)	15 526 228 (15 235 982 to 16 036 397)	-20% (-22 to -18)
<b>Alzheimer's disease and other dementias</b>												
DALYs	444 (416 to 473)	399 (372 to 427)	-10% (-12 to -8)	3 240 468 (3 038 789 to 3 453 552)	5 271 654 (4 915 486 to 5 641 197)	63% (58 to 66)	446 (418 to 476)	410 (383 to 438)	-8% (-10 to -7)	4 774 751 (4 471 802 to 5 088 817)	7 686 392 (7 175 157 to 8 205 020)	61% (58 to 63)
Deaths	38 (38 to 39)	35 (34 to 36)	-8% (-11 to -6)	260 244 (258 513 to 262 203)	501 678 (489 118 to 512 705)	93% (88 to 97)	38 (38 to 39)	36 (35 to 36)	-7% (-9 to -6)	376 119 (372 115 to 379 282)	700 353 (685 974 to 712 101)	86% (83 to 89)
Prevalence	654 (580 to 727)	594 (531 to 657)	-9% (-11 to -8)	4 788 372 (4 216 845 to 5 360 285)	7 740 481 (6 896 189 to 8 583 917)	62% (57 to 67)	658 (580 to 734)	612 (541 to 681)	-7% (-8 to -6)	7 074 259 (6 222 667 to 7 942 113)	11 375 859 (10 038 137 to 12 683 497)	61% (57 to 65)
YLDs	97 (69 to 126)	88 (63 to 114)	-9% (-9 to -9)	706 265 (504 015 to 919 170)	1 165 391 (832 669 to 1 516 059)	65% (59 to 71)	97 (69 to 126)	90 (64 to 118)	-7% (-8 to -5)	1 033 646 (741 294 to 1 345 291)	1 694 270 (1 209 023 to 2 212 896)	64% (59 to 69)
YLLs	347 (345 to 349)	311 (301 to 320)	-10% (-13 to -9)	2 534 204 (2 517 343 to 2 553 205)	4 106 264 (3 982 651 to 4 213 014)	62% (57 to 66)	349 (346 to 352)	320 (312 to 327)	-8% (-10 to -7)	3 741 105 (3 702 678 to 3 772 727)	5 992 121 (5 844 318 to 6 113 018)	60% (57 to 63)
<b>Parkinson's disease</b>												
DALYs	68 (63 to 82)	70 (63 to 78)	2% (-17 to 7)	522 267 (482 466 to 625 465)	839 916 (766 256 to 940 813)	61% (31 to 69)	69 (63 to 80)	70 (64 to 78)	2% (-10 to 5)	781 347 (720 120 to 908 025)	1 226 579 (1 126 744 to 1 363 923)	57% (38 to 62)
Deaths	5 (4 to 6)	5 (4 to 5)	2% (-19 to 8)	34 236 (33 270 to 41 800)	60 812 (55 723 to 67 973)	78% (40 to 88)	4 (4 to 5)	5 (4 to 5)	2% (-13 to 6)	49 050 (47 428 to 57 737)	84 599 (80 948 to 93 079)	72% (47 to 79)
Prevalence	94 (79 to 112)	105 (86 to 125)	11% (7 to 15)	704 533 (584 038 to 840 424)	1 215 045 (996 237 to 1 457 870)	72% (65 to 80)	100 (82 to 119)	107 (88 to 128)	7% (5 to 10)	1 123 483 (925 451 to 1 344 223)	1 838 098 (1 506 704 to 2 220 458)	64% (59 to 69)
YLDs	13 (9 to 18)	15 (10 to 20)	11% (7 to 16)	99 431 (68 796 to 134 551)	170 916 (117 642 to 233 979)	72% (64 to 80)	14 (10 to 19)	15 (10 to 21)	8% (5 to 11)	158 046 (107 768 to 215 467)	257 986 (178 581 to 354 467)	63% (57 to 69)
YLLs	55 (53 to 69)	55 (51 to 63)	-1% (-2 to 6)	422 835 (409 771 to 526 346)	669 000 (612 867 to 764 328)	58% (23 to 68)	55 (53 to 65)	55 (52 to 62)	0% (-14 to 4)	623 301 (601 534 to 744 574)	968 593 (924 800 to 1 088 287)	55% (33 to 62)
<b>Motor neuron disease</b>												
DALYs	27 (26 to 29)	31 (29 to 32)	12% (4 to 20)	157 904 (151 812 to 164 075)	252 987 (240 153 to 264 173)	60% (49 to 69)	22 (19 to 26)	24 (23 to 26)	11% (-5 to 27)	203 830 (188 169 to 229 421)	314 523 (298 988 to 348 088)	54% (38 to 75)
Deaths	1 (1 to 1)	1 (1 to 1)	29% (19 to 37)	6034 (5821 to 6282)	11 495 (10 900 to 12 020)	91% (75 to 102)	1 (1 to 1)	1 (1 to 1)	31% (20 to 43)	7 076 (6 773 to 7 536)	13 410 (12 728 to 14 790)	90% (76 to 103)

(Table 2 continues on next page)

	EU28			WHO European region								
	Age-standardised rate, 1990	Age-standardised rate, 2017	Change from 1990 to 2017	Total, 1990	Total, 2017	Change from 1990 to 2017	Age-standardised rate, 1990	Age-standardised rate, 2017	Change from 1990 to 2017	Total, 1990	Total, 2017	Change from 1990 to 2017
(Continued from previous page)												
Prevalence	4 (4 to 5)	6 (5 to 7)	42% (39 to 45)	23 986 (21 913 to 26 169)	42 937 (38 970 to 47 145)	79% (74 to 85)	3 (3 to 4)	4 (4 to 5)	35% (32 to 37)	30 603 (27 763 to 33 544)	51 711 (46 991 to 56 707)	69% (64 to 74)
YLDs	1 (1 to 1)	1 (1 to 2)	42% (39 to 45)	5101 (3618 to 6809)	9131 (6492 to 12 227)	79% (74 to 85)	1 (0 to 1)	1 (1 to 1)	35% (32 to 37)	6509 (4612 to 8693)	10 998 (7 822 to 14 720)	69% (64 to 74)
YLLs	27 (25 to 28)	29 (28 to 31)	11% (3 to 19)	152 803 (146 985 to 158 391)	243 856 (231 168 to 254 703)	60% (48 to 69)	21 (19 to 25)	23 (22 to 25)	10% (-7 to 27)	197 322 (181 684 to 224 050)	303 526 (287 735 to 337 104)	54% (37 to 75)
<b>Multiple sclerosis</b>												
DALYs	39 (33 to 45)	39 (32 to 46)	0% (-16 to 12)	226 945 (195 825 to 260 163)	282 765 (233 418 to 328 839)	25% (1 to 37)	33 (28 to 38)	32 (26 to 37)	-5% (-15 to 11)	326 058 (279 079 to 372 431)	391 078 (327 333 to 453 473)	20% (5 to 38)
Deaths	1 (1 to 1)	1 (0 to 1)	-6% (-40 to 6)	4 344 (3 781 to 5 021)	5 601 (3 954 to 6 056)	29% (-21 to 41)	1 (1 to 1)	1 (0 to 1)	-11% (-35 to 8)	6104 (5 202 to 6 645)	7 413 (5 673 to 8 395)	21% (-14 to 43)
Prevalence	65 (59 to 71)	77 (69 to 85)	18% (16 to 21)	369 975 (336 977 to 407 130)	522 674 (473 413 to 579 422)	41% (39 to 44)	52 (47 to 58)	59 (53 to 65)	13% (11 to 14)	504 285 (457 958 to 555 895)	699 238 (635 288 to 772 355)	39% (37 to 41)
YLDs	17 (12 to 22)	20 (14 to 26)	18% (13 to 23)	94 374 (67 351 to 123 083)	132 159 (95 038 to 171 558)	40% (34 to 46)	13 (10 to 18)	15 (11 to 20)	12% (9 to 16)	129 438 (92 358 to 168 926)	178 020 (127 760 to 230 616)	38% (33 to 42)
YLLs	22 (18 to 25)	19 (14 to 22)	-13% (-40 to 6)	132 570 (111 714 to 148 143)	150 607 (110 315 to 166 542)	14% (-25 to 34)	20 (16 to 21)	17 (14 to 20)	-16% (-34 to 10)	196 620 (162 235 to 206 779)	213 059 (169 517 to 254 322)	8% (-17 to 37)
<b>Brain and nervous system cancer</b>												
DALYs	153 (136 to 171)	140 (117 to 153)	-9% (-28 to 5)	823 636 (748 909 to 932 285)	964 964 (789 495 to 1 033 050)	17% (-11 to 28)	158 (141 to 175)	147 (130 to 163)	-7% (-20 to 5)	1 435 491 (1 289 242 to 1 584 211)	1 668 047 (1 439 524 to 1 799 509)	16% (-1 to 28)
Deaths	4 (4 to 5)	4 (3 to 4)	2% (-25 to 9)	25 123 (23 238 to 28 753)	35 917 (27 137 to 38 316)	43% (0 to 54)	4 (4 to 4)	4 (3 to 4)	5% (-15 to 14)	39 568 (36 090 to 44 242)	56 246 (45 620 to 59 326)	42% (12 to 55)
Prevalence	24 (22 to 27)	55 (46 to 62)	127% (76 to 168)	126 363 (114 687 to 142 588)	330 074 (270 005 to 360 630)	161% (98 to 192)	21 (19 to 24)	45 (40 to 52)	113% (76 to 150)	190 647 (174 163 to 212 101)	462 982 (398 536 to 511 728)	143% (97 to 176)
YLDs	3 (2 to 3)	5 (3 to 6)	84% (41 to 112)	13 971 (9 980 to 18 627)	30 368 (21 160 to 40 887)	117% (61 to 142)	2 (2 to 3)	4 (3 to 5)	74% (43 to 101)	21 212 (15 112 to 28 018)	43 489 (31 024 to 57 873)	105% (64 to 128)
YLLs	150 (134 to 169)	135 (113 to 148)	-10% (-30 to 3)	809 666 (735 221 to 917 160)	934 596 (766 157 to 997 931)	15% (-12 to 26)	156 (139 to 172)	143 (127 to 158)	-8% (-21 to 4)	1 414 279 (1 271 685 to 1 561 721)	1 624 558 (1 404 505 to 1 753 504)	15% (-2 to 27)
<b>Meningitis</b>												
DALYs	75 (71 to 81)	20 (18 to 24)	-74% (-76 to -69)	278 620 (261 680 to 296 452)	84 550 (77 767 to 97 340)	-70% (-72 to -65)	172 (115 to 203)	38 (35 to 43)	-78% (-81 to -64)	1 182 458 (811 688 to 1 378 097)	288 614 (267 659 to 324 772)	-76% (-79 to -61)
Deaths	1 (1 to 1)	0 (0 to 0)	-74% (-76 to -69)	4 937 (4 360 to 5 147)	2 033 (1 947 to 2 224)	-59% (-61 to -51)	2 (2 to 3)	1 (1 to 1)	-75% (-78 to -60)	16 633 (11 721 to 18 979)	5 483 (5 253 to 6 125)	-67% (-71 to -50)
Prevalence	67 (59 to 76)	33 (28 to 39)	-51% (-54 to -47)	336 120 (292 854 to 384 061)	183 151 (153 769 to 216 836)	-46% (-50 to -41)	98 (85 to 112)	53 (45 to 62)	-46% (-49 to -42)	857 183 (742 594 to 983 424)	505 430 (430 502 to 591 790)	-41% (-45 to -37)
YLDs	7 (5 to 9)	3 (2 to 5)	-49% (-52 to -46)	31 197 (21 873 to 41 791)	17 079 (12 032 to 22 999)	-45% (-49 to -42)	9 (6 to 12)	5 (4 to 7)	-44% (-46 to -41)	76 367 (53 464 to 102 765)	45 301 (31 738 to 60 745)	-41% (-43 to -38)

(Table 2 continues on next page)

	EU28					WHO European region						
	Age-standardised rate, 1990	Age-standardised rate, 2017	Change from 1990 to 2017	Total, 1990	Total, 2017	Change from 1990 to 2017	Age-standardised rate, 1990	Age-standardised rate, 2017	Change from 1990 to 2017	Total, 1990	Total, 2017	Change from 1990 to 2017
(Continued from previous page)												
YLLs	69 (65 to 75)	16 (15 to 20)	-76% (-79 to -71)	247 423 (231 164 to 263 632)	67 471 (64 007 to 80 231)	-73% (-75 to -67)	163 (107 to 193)	33 (31 to 39)	-80% (-83 to -66)	1 106 091 (733 867 to 1 300 227)	243 313 (230 129 to 280 048)	-78% (-81 to -64)
<b>Encephalitis</b>												
DALYs	20 (14 to 21)	11 (10 to 12)	-44% (-50 to -16)	81 788 (62 092 to 87 719)	57 720 (50 302 to 63 183)	-29% (-36 to -6)	42 (38 to 48)	34 (30 to 38)	-19% (-34 to -4)	321 230 (291 041 to 359 673)	270 736 (236 953 to 292 768)	-16% (-29 to -4)
Deaths	0 (0 to 0)	0 (0 to 0)	-30% (-36 to -9)	1 581 (1 338 to 1 648)	1 729 (1 437 to 1 829)	9% (1 to 24)	1 (1 to 1)	1 (0 to 1)	-11% (-24 to -2)	5 433 (4 981 to 5 885)	5 953 (4 931 to 6 266)	10% (-5 to 17)
Prevalence	26 (14 to 41)	22 (12 to 35)	-16% (-18 to -15)	141 844 (76 154 to 230 446)	142 435 (74 374 to 234 773)	0% (-2 to 3)	34 (19 to 53)	29 (17 to 46)	-14% (-15 to -13)	311 615 (175 514 to 494 479)	315 202 (173 810 to 507 091)	1% (-1 to 3)
YLDs	2 (1 to 3)	2 (1 to 2)	-18% (-20 to -17)	9 989 (6 836 to 13 303)	9 372 (6 386 to 12 593)	-6% (-8 to -4)	3 (2 to 4)	2 (2 to 3)	-15% (-17 to -14)	23 932 (16 651 to 31 792)	22 773 (15 651 to 30 309)	-5% (-7 to -3)
YLLs	18 (12 to 19)	10 (8 to 11)	-47% (-53 to -15)	71 799 (52 152 to 76 731)	48 348 (41 387 to 51 476)	-33% (-39 to -6)	40 (36 to 46)	32 (28 to 36)	-19% (-35 to -4)	297 298 (266 649 to 334 872)	247 963 (214 185 to 269 172)	-17% (-30 to -4)
<b>Tetanus</b>												
DALYs	1 (1 to 1)	0 (0 to 0)	-90% (-92 to -88)	4809 (3793 to 5134)	649 (558 to 755)	-86% (-88 to -84)	11 (4 to 26)	0 (0 to 0)	-98% (-99 to -95)	69 067 (28 864 to 158 779)	2127 (1860 to 2770)	-97% (-99 to -93)
Deaths	0 (0 to 0)	0 (0 to 0)	-90% (-91 to -88)	240 (197 to 256)	39 (33 to 46)	-84% (-86 to -81)	0 (0 to 0)	0 (0 to 0)	-96% (-98 to -92)	1 066 (612 to 2 088)	82 (72 to 100)	-92% (-96 to -87)
Prevalence	0 (0 to 0)	0 (0 to 0)	-88% (-90 to -85)	26 (20 to 30)	4 (3 to 5)	-83% (-85 to -80)	0 (0 to 0)	0 (0 to 0)	-33% (-55 to -11)	214 (166 to 296)	160 (133 to 187)	-25% (-46 to -7)
YLDs	0 (0 to 0)	0 (0 to 0)	-90% (-91 to -88)	3 (2 to 5)	1 (0 to 1)	-85% (-87 to -82)	0 (0 to 0)	0 (0 to 0)	-77% (-88 to -58)	15 (8 to 28)	4 (3 to 7)	-72% (-84 to -54)
YLLs	1 (1 to 1)	0 (0 to 0)	-90% (-92 to -88)	4806 (3790 to 5131)	649 (558 to 754)	-86% (-88 to -84)	11 (4 to 26)	0 (0 to 0)	-98% (-99 to -95)	69 051 (28 852 to 158 748)	2 122 (1 856 to 2 765)	-97% (-99 to -93)
<b>Epilepsy</b>												
DALYs	149 (105 to 208)	140 (90 to 211)	-7% (-30 to 24)	735 793 (514 448 to 1 023 814)	781 549 (512 458 to 1 168 936)	6% (-20 to 41)	179 (137 to 235)	158 (113 to 220)	-12% (-29 to 9)	1 507 756 (1 145 663 to 1 988 615)	1 479 134 (1 063 428 to 2 050 641)	-2% (-21 to 21)
Deaths	1 (1 to 1)	1 (1 to 1)	3% (-28 to 11)	6 223 (6 106 to 6 782)	9 526 (7 286 to 10 097)	53% (3 to 64)	1 (1 to 2)	1 (1 to 1)	-8% (-26 to 8)	12 528 (11 675 to 13 504)	15 370 (12 663 to 16 443)	23% (-4 to 38)
Prevalence	358 (241 to 470)	399 (266 to 529)	11% (-19 to 50)	1 794 858 (1 198 133 to 2 356 841)	2 244 668 (1 496 976 to 2 977 225)	25% (-9 to 69)	352 (259 to 453)	385 (280 to 497)	9% (-15 to 35)	3 082 147 (2 254 550 to 3 960 937)	3 742 828 (2 718 716 to 4 823 770)	21% (-6 to 50)
YLDs	98 (53 to 156)	96 (48 to 166)	-2% (-37 to 49)	482 243 (264 069 to 770 314)	523 003 (259 741 to 908 273)	8% (-31 to 64)	104 (63 to 159)	100 (57 to 160)	-3% (-32 to 30)	894 217 (541 420 to 1 373 670)	942 629 (528 597 to 1 512 164)	5% (-25 to 44)
YLLs	52 (51 to 59)	44 (40 to 47)	-15% (-40 to -8)	253 550 (247 508 to 285 858)	258 546 (219 592 to 274 228)	2% (-30 to 10)	75 (65 to 84)	58 (53 to 64)	-23% (-35 to -4)	613 539 (549 291 to 680 608)	536 505 (490 237 to 585 052)	-13% (-28 to 5)
<b>Migraine</b>												
DALYs	763 (491 to 1106)	770 (495 to 1115)	1% (0 to 2)	3 947 343 (2 552 335 to 5 707 835)	4 231 019 (2 740 462 to 6 048 715)	7% (5 to 9)	737 (475 to 1059)	735 (473 to 1056)	0% (-1 to 1)	6 702 168 (4 320 439 to 9 679 278)	7 401 471 (4 810 162 to 10 539 348)	10% (9 to 12)

(Table 2 continues on next page)

	EU28			WHO European region			WHO European region			WHO European region		
	Age-standardised rate, 1990	Age-standardised rate, 2017	Change from 1990 to 2017	Total, 1990	Total, 2017	Change from 1990 to 2017	Age-standardised rate, 1990	Age-standardised rate, 2017	Change from 1990 to 2017	Total, 1990	Total, 2017	Change from 1990 to 2017
(Continued from previous page)												
Prevalence	20543 (19061 to 22105)	20646 (19234 to 22225)	0% (0 to 1)	105746319 (98488927 to 113530086)	112207672 (104739016 to 119973469)	6% (4 to 8)	19697 (18304 to 21171)	19650 (18270 to 21082)	0% (-1 to 1)	178317032 (166023492 to 191281010)	195794407 (182677431 to 209186190)	10% (8 to 11)
YLDs	763 (491 to 1106)	770 (495 to 1115)	1% (0 to 2)	3947343 (2552335 to 5707835)	4231019 (2740462 to 6048715)	7% (5 to 9)	737 (475 to 1059)	735 (473 to 1056)	0% (-1 to 1)	6702168 (4320439 to 9679278)	7401471 (4810162 to 10539348)	10% (9 to 12)
<b>Tension-type headache</b>												
DALYs	102 (58 to 160)	101 (57 to 159)	0% (-2 to 1)	542038 (309633 to 850297)	600667 (343433 to 936239)	11% (8 to 14)	113 (64 to 176)	110 (63 to 173)	-2% (-3 to -1)	1043015 (596692 to 1625120)	1163732 (664951 to 1799904)	12% (9 to 14)
Prevalence	31256 (28329 to 34499)	30871 (28014 to 33930)	-1% (-2 to 0)	162144987 (147265221 to 178088317)	173696347 (158485543 to 190762959)	7% (5 to 10)	31858 (28899 to 35154)	31022 (28086 to 34097)	-3% (-3 to -2)	287861348 (261220814 to 316795771)	309890284 (281758135 to 340323834)	8% (6 to 10)
YLDs	102 (58 to 160)	101 (57 to 159)	0% (-2 to 1)	542038 (309633 to 850297)	600667 (343433 to 936239)	11% (8 to 14)	113 (64 to 176)	110 (63 to 173)	-2% (-3 to -1)	1043015 (596692 to 1625120)	1163732 (664951 to 1799904)	12% (9 to 14)
<b>Other neurological disorders</b>												
DALYs	66 (57 to 76)	65 (54 to 79)	-1% (-16 to 16)	313735 (277741 to 360083)	363480 (314552 to 426877)	16% (3 to 30)	64 (55 to 75)	63 (54 to 76)	-1% (-14 to 15)	531078 (460688 to 617442)	599059 (517397 to 698284)	13% (0 to 28)
Deaths	1 (1 to 1)	1 (1 to 1)	-2% (-11 to 6)	7205 (6931 to 7450)	10303 (9219 to 10814)	43% (26 to 52)	1 (1 to 1)	1 (1 to 1)	1% (-8 to 10)	9941 (9424 to 10402)	14052 (12762 to 14637)	41% (26 to 52)
Prevalence	0 (0 to 1)	0 (0 to 1)	7% (4 to 11)	2067 (1372 to 2855)	2613 (1736 to 3517)	26% (20 to 34)	0 (0 to 1)	0 (0 to 1)	4% (2 to 6)	3674 (2440 to 5020)	4364 (2939 to 5868)	19% (13 to 26)
YLDs	18 (11 to 28)	23 (13 to 37)	28% (-8 to 76)	84916 (54641 to 126893)	112433 (68271 to 174266)	32% (0 to 76)	19 (12 to 28)	24 (15 to 36)	24% (-6 to 61)	159965 (104111 to 235740)	206737 (130672 to 304277)	29% (1 to 63)
YLLs	48 (45 to 52)	42 (39 to 44)	-12% (-24 to -3)	228818 (218306 to 242385)	251047 (232719 to 264643)	10% (-2 to 20)	45 (39 to 50)	40 (37 to 42)	-12% (-24 to 3)	371114 (333652 to 403551)	392321 (369776 to 410145)	6% (-6 to 20)

Data are n (95% UI) or % (95% UI). DALYs=disability-adjusted life-years. EU=the 27 EU countries plus the UK. YLDs=years lived with disability. YLLs=years of life lost. UI=uncertainty interval.

**Table 2: DALYs, death, prevalence, YLDs, YLLs, and rates of neurological disorders by category in the EU28 and WHO European region, with changes from 1990 to 2017**

The high proportion of DALYs and deaths attributable to neurological disorders can be explained in part by the long life expectancy in Europe and, despite an overall decrease in age-standardised rates, by the increasing incidence and the increasingly long duration of ageing-related diseases.<sup>10</sup> Population growth could also be implicated. The total population of Europe was 721 million in 1990 and rose to 916 million by 2016.<sup>11</sup>

While the substantial sex differences observed in the burden of neurological disorders reflect the differing distribution of each clinical condition in men and women, the increasing number of DALYs attributable to the most common clinical conditions with age is most likely linked with the ageing of the European population. The peak of DALYs observed in the oldest age groups reflects the predominance of ageing-related diseases in these older age groups. The predominance of dementia

in the oldest groups in the EU28 and in western Europe can be similarly explained.

Stroke, migraine, and dementia were the major contributors to the overall burden of neurological disorders when comparing the EU28, western, central, and eastern Europe, but with different numbers according to age. Whereas in western Europe dementia and stroke were the largest contributors to the burden of neurological disorders after age 70 years in both sexes, stroke was the strongest contributor between age 40 years and 89 years in central Europe and between age 50 years and 89 years in eastern Europe. National and regional differences in population age structure (for dementia), implementation of preventive and therapeutic measures (for stroke), and attention to diseases and access to health-care facilities are possible explanations.

This study has shown differences among WHO subregions in the proportion of DALYs, deaths, and

	High-SDI countries			High-middle SDI countries		
	1990	2017	Change	1990	2017	Change
Stroke	1286 (1228 to 1341)	650 (597 to 703)	-49% (-52 to -47)	3196 (3097 to 3294)	1998 (1900 to 2104)	-37% (-40 to -35)
Alzheimer's disease and other dementias	430 (403 to 458)	399 (373 to 426)	-7% (-9 to -5)	466 (436 to 497)	439 (410 to 472)	-6% (-8 to -3)
Parkinson's disease	63 (58 to 74)	67 (59 to 73)	6% (-16 to 11)	78 (72 to 86)	74 (68 to 83)	-4% (-10 to 0)
Motor neuron disease	28 (27 to 29)	31 (30 to 32)	11% (6 to 16)	8 (7 to 12)	10 (9 to 11)	17% (-22 to 43)
Multiple Sclerosis	33 (28 to 28)	34 (29 to 40)	6% (-12 to 10)	14 (12 to 16)	11 (10 to 13)	-21% (-31 to -1)
Brain and nervous system cancer	125 (112 to 139)	114 (99 to 124)	-9% (-25 to 3)	148 (113 to 169)	132 (117 to 147)	-11% (-22 to 16)
Meningitis	62 (58 to 66)	19 (17 to 22)	-70% (-72 to -63)	201 (144 to 227)	46 (40 to 49)	-77% (-80 to -71)
Encephalitis	14 (13 to 15)	10 (9 to 12)	-27% (-34 to -12)	54 (46 to 64)	32 (28 to 38)	-41% (-54 to -16)
Tetanus	1 (1 to 1)	0 (0 to 0)	-89% (-90 to -86)	43 (33 to 54)	2 (1 to 3)	-95% (-97 to -91)
Epilepsy	125 (88 to 171)	118 (77 to 173)	-6% (-23 to 14)	175 (138 to 223)	127 (91 to 177)	-27% (-42 to -9)
Migraine	676 (433 to 975)	674 (431 to 970)	0% (-1 to -1)	550 (350 to 800)	552 (351 to 804)	0% (-1 to 2)
Tension-type headache	93 (53 to 147)	93 (53 to 148)	0% (-1 to 1)	97 (55 to 151)	92 (52 to 145)	-5% (-6 to -3)
Other neurological disorders	61 (54 to 70)	61 (52 to 72)	0% (-10 to -10)	46 (37 to 56)	45 (37 to 57)	-2% (-18 to 21)

Data are n (95% UI) or % (95% UI). DALY=disability-adjusted life-year. EU28=the 27 EU countries plus the UK. SDI=socio-demographic index. UI=uncertainty interval.

**Table 3: Age-standardised DALY rates by SDI in the EU28 and WHO European region from 1990 to 2017**

prevalence attributable to the commonest neurological disorders. Stroke was the largest contributor to the total DALYs in eastern Europe, followed by central and western Europe, whereas dementia showed the opposite gradient.

Stroke was the leading source of burden attributable to neurological disorders in the EU28 and the WHO European region. The age-standardised DALY rate for stroke was significantly higher in high-middle SDI than in high-SDI countries. Stroke is the overall leading contributor of DALYs and deaths worldwide.<sup>12</sup> However, the declining all-age and age-standardised DALYs attributable to stroke is an important finding. As the prevalence of stroke showed a substantial increase, possibly due to the ageing of the society and the increasing detection of less severe cases, we found a corresponding decrease in DALYs, both absolute and relative. Preventive medicine and lifestyle changes are the most likely explanations and underscore the effects of increasing control of preventable risk factors (blood pressure control and smoking cessation) affecting mainly the prevalence. Previous reports from the GBD collaborators showed that more than 90% of the stroke burden was attributable to modifiable risk factors.<sup>13-16</sup> Stroke unit care and thrombolysis have been proposed to have a role in ameliorating the outcome of ischaemic stroke<sup>17</sup> and can explain the significant reduction in deaths and DALYs per patient. In the USA, the rate of use of r-TPA began to increase in 2001 and progressed steadily in the subsequent years.<sup>18</sup> Similar trends are most likely present in Europe.

The increasing burden of Alzheimer's disease and other dementias observed since 1990 is in line with

increases in life expectancy, with burden increasing most rapidly in the population aged 60 years or older due to increasing longevity and declining birth rates.<sup>19</sup>

Another notable disease with a large increase in prevalence and DALYs was Parkinson's disease. The global burden of Parkinson's disease more than doubled during the study period.<sup>20</sup> The growth of the burden of Parkinson's disease surpassed that of Alzheimer's disease and other dementias. This increase has been attributed to the ageing of the European population, the reduction of rural populations, occupational exposures, and the declining smoking rates.<sup>21,22</sup> Although increased attention towards the disease might partially explain the increasing incidence observed, Parkinson's disease is still underdiagnosed, as shown in the USA<sup>23</sup> and by an online European survey that found that 40% of respondents had never seen a Parkinson's disease specialist.<sup>24,25</sup>

A substantial increase in all the measures of burden has been observed in Europe for motor neuron disease, another ageing-related disease. However, contrary to the other neurodegenerative diseases, we found a parallel increase of age-standardised rates, which can be explained by an increasing diagnostic accuracy rather than a true increase of the incidence of the disease.

Primary headache (migraine and tension-type headache) was the most prevalent neurological disorder and the second highest source of burden both in high-SDI and in high-middle SDI countries. Headache is a major public health concern worldwide. Based on the results of the GBD Study 2016,<sup>26</sup> almost 3 billion individuals were estimated to have repeated migraine or tension-type headache attacks, accounting for a total of 52.3 million YLDs. In the WHO European region, migraine and

tension-type headache were the second most common source of burden attributable to neurological disorders (the third in the EU28). In this study, headache was in line with the worldwide burden of the disease and, by definition, accounted for a substantial health loss in terms of YLDs, with premature mortality virtually absent. Despite the abundance of symptomatic drugs available to treat headache, we found a modest increase in the absolute measures of burden of headache, suggesting that interventions should be directed towards increasing public knowledge of and decreasing exposure to modifiable risk factors, such as obesity, smoking, air pollution, low physical activity, blood pressure, and stress.<sup>26</sup>

Contrary to other neurological disorders and in spite of a slight increase in prevalence and deaths, epilepsy showed a reduction in all-age and age-standardised DALYs in Europe. The increasing number of deaths might be due to the ageing of the European population. This finding is not unexpected because epilepsy incidence is age related, with a first peak in the young but a steady increase in the older population and thereby an increase of mortality in an ageing society.<sup>27</sup> The decreasing burden might be instead a reflection of improved management. The differences between high-SDI countries (showing a slight increase) and high-middle-SDI countries (showing a significant decrease) can be explained by the contrasting effects of ageing and quality of care, with a negative balance in countries with top economies (where the management of epilepsy could not further improve despite the increasing number of affected individuals) and a positive balance in growing economies (where the quality of care might be rapidly improving).

Multiple sclerosis is less frequent than other neurological disorders. The increasing prevalence, and the consequent burden, when comparing the first and the last year of the study period, could be due to the changing classifications of the disease. The disease can now be diagnosed earlier than was possible at the start of the study. Patients with clinically isolated syndromes showing one attack and evidence of two or more T2 or gadolinium-enhancing lesions have been included since 2010.<sup>28</sup> Therefore, the incidence of the disease might not be truly increasing, but the expansion of the diagnostic boundaries might be a source of increasing burden and, consequently, of expenditures for the propensity of the caring physicians to anticipate the onset of treatment with disease-modifying drugs in the attempt to improve the disease course.<sup>29</sup>

In line with the global decrease of absolute numbers and age-standardised rates,<sup>30</sup> CNS infections and tetanus showed a significant decrease in the EU28 and the WHO European region. The increase in prevalence and YLDs and, to a lesser extent, the increase of deaths attributable to brain tumours can be explained by better diagnostic ascertainment; the role of environmental risk factors is controversial.<sup>31,32</sup>

All the other neurological disorders accounted for a low proportion of the overall burden, due to low prevalence

or a less remarkable effect on life expectancy or functional abilities.

The standardised DALY rates for all neurological diseases varied across countries but, with few exceptions, we found an overall decrease, mostly driven by a reduction in premature mortality. Given the predominant role of stroke in terms of attributable proportion, we infer that the decrease in DALYs overall is in large part explained by the improved prevention of stroke deaths.

The major strength of the study is the collection of epidemiological data worldwide using the same methodology and modelling measures. This methodology allowed us to document major differences between the EU28 and the WHO European region and within European countries in the burden of neurological disorders. Our study was able to highlight geographical trends in the burden of stroke within Europe and suggests that prevention and treatment are still deficient in several European countries. These results can be helpful for local authorities to address the burden of preventable and treatable neurological disorders. Other strengths are the continuous refinement of the available data, adding new original sources and the use of more sophisticated statistical methods when available, which allowed us to compare the European situation with the spectrum of the global situation.

This study also has limitations. First, because original epidemiological data were not available for all countries, Bayesian statistical models were used to estimate deaths and disease prevalence for countries where information was missing. For this reason, the inclusion in future of new primary data from countries with little data could lead to more precise estimates that might vary from current predicted values. Second, the disability weights used for the calculation of YLDs might not be uniform across populations and sociodemographic strata. However, a systematic variation in disability weights across populations, or within the same population as a function of education, was not detected by population surveys.<sup>7,33</sup> Third, the 95% UIs are frequently wide, reflecting the low precision of the estimates and potentially limiting the ability to detect smaller differences across countries. Additional data on the proportions influencing the sequelae of neurological disorders might reduce uncertainties in future analyses. Fourth, neurological diseases can be correlated with other somatic and psychiatric comorbidities and injuries. Here, the correction for comorbidity was based on the assumption that diseases and their sequelae are independent. Future improvements of the GBD modelling should include dependent comorbidity. Fifth, disease rankings that are based on age-standardised rates can be influenced by the choice of the standard population. Sixth, the use of medical claims data could be biased because people who are not treated or are excluded from health insurance would not be counted. Seventh, the results reflect the study period and the

quality of the sources. Additionally, the variable quality and completeness of the information available can partly explain the differences found when comparing the different European countries. Eighth, the definition of causes of death reflects the quality and completeness of the sources. Finally, several highly prevalent neurological diseases are not yet included in the GBD database, such as sleep disorders,<sup>34</sup> essential tremor,<sup>35</sup> or restless legs syndrome.<sup>36</sup>

This study has shown the huge burden of neurological disorders in Europe. The numbers of citizens affected and the numbers of neurological DALYs and deaths call for action regarding this disease group. The major challenge for all countries is the increasing number of neurological diseases even if the DALYs per patient are decreasing for some conditions. Despite many similarities, the most prevalent diseases differ considerably in different regions or even countries and thus policy and actions needed might differ. Strategic planning in European countries might use this information to customise plans to efficiently deal with this burden. However, to improve the precision of our estimates, new studies are awaited to provide original data in countries where the information regarding the incidence, prevalence, and mortality of the target diseases is not yet available. The inclusion of these data in our models can correct the present findings and provide a more accurate estimate of the burden of neurological diseases and trends.

#### Contributors

EB, GD, and TV analysed the data and prepared the first draft. All other authors provided data, developed models, reviewed results, provided guidance on methodology, or reviewed and approved the final version of the manuscript.

#### Declaration of interests

GD reports grants from Medtronic; and personal fees from Boston Scientific, Cavion, Functional Neuromodulation, and Thieme publishers, outside of the submitted work. GD receives funding for his research from the German Research Council (SFB 1261, B5). EB reports grants from the Italian Ministry of Health and the Swedish Orphan Biovitrum, and American ALS Association; and personal fees from Arvelle Therapeutics, outside of the submitted work. All other authors declare no competing interests.

#### Data sharing

Data collected within the framework of this study are accessible to interested parties by contacting the corresponding author.

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