Health claims databases offer opportunities for studies on large populations of patients with kidney disease and health outcomes in a non-experimental setting. Among others, their unique features enable studies on healthcare costs or on longitudinal, epidemiological data with nationwide coverage. However, health claims databases also have several limitations. Because clinical data and information on renal function are often lacking, the identification of patients with kidney disease depends on the actual presence of diagnosis codes only. Investigating the validity of these data is therefore crucial to assess whether outcomes derived from health claims data are truly meaningful. Also, one should take into account the coverage and content of a health claims database, especially when making international comparisons. In this article, an overview is provided of international health claims databases and their main publications in the area of nephrology. The structure and contents of the Dutch health claims database will be described, as well as an initiative to use the outcomes for research and the development of the Dutch Kidney Atlas. Finally, we will discuss to what extent one might be able to identify patients with kidney disease using health claims databases, as well as their strengths and limitations.

Keywords: CKD, dialysis, epidemiology, health claims data, health claims database, kidney transplantation
longitudinal data on medical diagnoses and procedures, pharmacological treatment and costs. Most of these databases have nationwide coverage, include all age categories and offer data that reflect day-to-day clinical practice. However, when using administrative data for healthcare research, it is important to recognize their unique opportunities as well as their limitations. Furthermore, for international comparisons, one should take into consideration the differences in database characteristics when analysing and interpreting these data [1]. Nowadays, several health claims databases of national and regional healthcare systems are available and are being used for kidney research.

In this article we provide a worldwide overview of health claims databases and summarize their main publications in the area of nephrology. Next, we will introduce the Dutch health claims database and a new initiative to use it for research and the development of a Dutch Kidney Atlas. Finally, we will discuss to what extent they allow identification of patients with kidney disease as well as the strengths and limitations of such databases.

WORLDWIDE HEALTH INSURANCE CLAIMS DATABASES IN KIDNEY RESEARCH

We selected papers in which health claims data, whether or not combined with other administrative databases, were used for research on patients with kidney disease. A systematic literature search was not possible because the use of health claims data is often not clearly mentioned in the articles. We performed an Internet and PubMed search in August 2019 using different search terms (e.g. health claims data, health insurance claims data, healthcare claims data, administrative data in combination with terms to search for research on patients with kidney disease such as CKD, dialysis or kidney transplantation). We selected suitable papers based on the abstract and methods section. In addition, we checked the references of selected papers and checked other published papers by the main authors.

We identified 13 health claims databases in 10 countries (Canada, China, the UK, France, India, Japan, South Korea, Taiwan, The Netherlands and the USA) that were used for research on patients with kidney disease and resulted at least one publication in English in a scientific journal (Table 1). Other countries, such as Austria and Germany, have a health claims database that is used for clinical research. However, we were unable to find papers published in scientific journals specifically focusing on kidney disease patients from these countries [2]. In some countries in Southeast Asia, health claims databases have been recently established and so far their use for research purposes has been rare [3]. The use of health claims databases for kidney research worldwide may therefore be expanded.

The organization of healthcare systems, healthcare financing and socio-economic settings differ importantly between countries [4]. This results in differences in the coverage, the size and the content of the health claims databases. A recent study presented a comprehensive overview of existing systems and financing for kidney care and demonstrated significant heterogeneity [5]. In countries where health insurance is obligatory or universally accessible (such as Canada, France, Japan, South Korea, The Netherlands and Taiwan), health claims databases have (almost) complete coverage of all country or province inhabitants (Table 1). The coverage of the Chinese claims databases, in contrast, is much less complete. Although they comprise a large number of individuals, the content of these is limited to the ones with access to healthcare insurance. The USA does not have universal healthcare coverage and the Medicare program only provides health insurance for American citizens ≥65 years of age as well as for people of all ages with severe diseases such as those with end-stage kidney disease. Therefore, the Medicare claims database covers a selection of the US population. Furthermore, the Rajiv Aarogyaari Community Health Insurance Scheme is an Indian state government program providing free hospital care to poor individuals. Noteworthy is the Hospital Episode Statistics (HES) database, which provides admission data for National Health Service (NHS) hospitals in the UK.

Within health claims data, kidney disease patients are generally identified with International Classification of Disease codes in combination with medical procedure codes. In Japan, patients are identified with Diagnosis Procedure Codes (DPC), whereas in The Netherlands patients are identified using diagnosis treatment combinations, a system similar to the DPC system. When evaluating studies using health claims data, it is essential to consider the coverage (such as age distribution, health system characteristics, insurance coverage and percentage of the population with healthcare insurance) and the specific procedure of patient selection to correctly interpret its results. This is of special importance when making international comparisons.

Accessibility of health claims databases

Several barriers for the secondary use of administrative data in general have been identified. The impact of these barriers differs considerably between countries and influences the availability and utility of health claims data for research per country [6]. In general, the complexity and the enormous amount of unprocessed raw data make it difficult and laborious to work with health claims databases. The pre-processing of data is time-consuming and requires experience with working with big data. In addition, extensive application processes as well as data processing fees are usually needed to access the data.

Data accessibility differs across countries. For example, the National Health Insurance Research Database (NHIRD) of Taiwan is known for its high accessibility. The database is publicly available and any researcher may apply for data that are provided for a small processing fee [7]. In other countries (e.g. South Korea, Japan, France, The Netherlands, Canada, the UK and the USA), researchers have to go through more extensive data request procedures [8–14]. Physical access to extracts of de-identified data is sometimes only possible in a designated secure environment (The Netherlands) or researchers are required to have a secured physical environment at their institution (Japan) [9, 15]. In France, data are supplied on a secure, electronic medium while in Canada access is provided via a secure, online research environment [10, 12]. Information about the accessibility of the databases of China and India was not available on public websites or in publications [8].

The processing fees are determined by fees for data extraction, the complexity of the data request and possible extra delivered services such as professional assistance in working with the complex databases. The level of the fees is therefore difficult to determine, but they are often described as costly. The extra financing needed for the use of health claims databases as well as the expertise needed for a study on health claims data (e.g. data analyst specialized in big data, nephrologist, epidemiologist and PhD student) may be challenging, especially for...
grant applications that use a fixed amount and usually mainly reimburse the salary costs of the researcher.

**Studies using health claims data**

Health claims data are being used to answer a wide variety of healthcare-related questions in different fields of research. Several countries have the ability to link their health claims database to other administrative databases such as national dialysis and transplantation registries, national vital statistics or clinical laboratory data (Table 1). This linkage capacity greatly expands its research possibilities [16].

In general, studies using health claims data mainly focus on the epidemiology of a disease, including patients’ morbidity and

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### Table 1. Overview of health claims databases in the world used for kidney research

| Country or province | Number of inhabitants in 2018 | Health claims database | Coverage | Linkage to other (administrative) databases |
|---------------------|------------------------------|------------------------|----------|-------------------------------------------|
| Canada - Alberta    | 4.3 million                  | Alberta Provincial Physician Claims database | >99% of inhabitants | Part of the Alberta Kidney Disease Network database; linkage to the Northern and Southern Alberta Renal Programs and clinical laboratory data |
| Canada - Manitoba   | 1.4 million                  | Manitoba Health Physician Claims database | >99% of inhabitants | Linkage to Manitoba Renal Program Dialysis Registry |
| Canada - Ontario    | 14.3 million                 | Ontario Health Insurance Plan database (OHIP) | >99% of inhabitants | Linkage to Ontario’s central organ and tissue donation agency, Canadian Institute for Health Information Discharge Abstract Database (CIHI-DAD), National Ambulatory Care Reporting System (data on emergency room visits), Ontario Registered Persons Database Information (demographics and vital status), Ontario Drug Benefit Plan (outpatient prescription drug usage for individuals >65 years) |
| Canada - Quebec     | 8.4 million                  | Régie de l’assurance maladie du Québec (RAMQ) | >99% of inhabitants | Linkage to Canadian Organ Replacement Register (CORR) |
| China               | 1.4 billion                  | China Health Insurance Research Association database (CHIRA) | 977 million insured people in 2015 | |
| UK                  | 55.3 million                 | Hospital Episode Statistics (HES) | 60 million customers in 2015 | Linkage to the Office for National Statistics (ONS) for mortality data |
| France              | 66.9 million                 | Système national d’information interrégimes de l’Assurance Maladie (Snírir) | 96% of inhabitants | Linkage to French Renal Epidemiology and Information Network (REIN) registry, French national hospital computerized medical information system (PMSI) |
| India - Andhra Pradesh | 1.3 billion                 | Rajiv Aarogyasri Community Health Insurance Scheme | 81% of inhabitants | |
| Japan               | 127.2 million                | National Database of Health Insurance Claims and Specific Health Checkups of Japan (NDB) | 90% of inhabitants | |
| South Korea         | 51.1 million                 | Health Insurance and Review Assessment Service (HIRA) | 98% of inhabitants | Linkage to a national health screening program (including 10 million Koreans) providing information on serum creatinine and urine albumin measurements |
| Taiwan              | 23.8 million                 | National Health Insurance Administration Research Database (NHIRD) | >99% of inhabitants | Linkage to e.g. death registry, cancer registry, reportable infectious disease registry |
| The Netherlands USA  | 17.1 million                 | Vektis database | 98% of inhabitants | All patients on RRT aged >65 years and patients with end-stage renal disease (RRT) |
| USA                 | 327.2 million                | Medicare Services |                    | Linkage to the United States Renal Data System (USRDS), Scientific registry of transplant recipients (SRTRS), National Health and Nutrition Examination Survey data (NHANES) |
survival, evaluating healthcare costs and the delivery of health-care services, describing prescription patterns and the effectiveness of pharmacological therapies and exploring clinical outcomes [14]. Table 2 provides a selection of papers on patients with kidney disease based on health claims data (or a combination of health claims data and other linked administrative data) published in scientific journals. We divided these articles into three main fields of research, i.e. validation studies (see Table 3), cost studies and descriptive and outcome studies, to provide insights into the variety of study questions.

Most papers reported on patients receiving dialysis treatment and significantly less on patients with kidney transplantation or CKD patients not treated with RRT. Interestingly, health claims data can also provide opportunities to report on specific kidney diseases such as polycystic kidney disease [54]. Several papers focused on the validity of health claims data to identify patients with CKD [57–59], dialysis [60–62] and kidney transplantation [63]. The results of these validity studies are presented in Table 3 and will be discussed later.

As health claims databases include reimbursement data, it is possible to study healthcare costs. Several studies provided cost estimations for dialysis and kidney transplantation in different countries [17–21, 23–26]. Estimations of healthcare costs of CKD patients not treated with RRT were less common and only provided in two studies [15, 22]. Since health claims usually contain longitudinal data with nationwide coverage, studies frequently report epidemiological data such as incidence and prevalence of kidney diseases, the occurrence of kidney disease-related risk factors/comorbidities, mortality rates and their trends over time. A wide range of outcomes have been described using health claims data. Frequently studied outcomes include those that are cardiovascular related, such as cardiovascular disease in transplant patients, stroke, atrial fibrillation and major cardiovascular events. The comprehensive drug delivery data of pharmacies provide unique opportunities for pharmaco-epidemiological research to study the use and effects of medication in CKD patients. This results, for example, in studies reporting the use of warfarin, angiotensin-converting enzyme inhibitors/angiotensin receptor blockers or β-blockers in dialysis patients or metformin use in CKD patients. With pharmacological data, it is important to realize that non-reimbursable drugs or over-the-counter drugs remain undetected in health claims data. Other less common outcomes reported include depressive disorder in kidney transplantation, fracture risk and associated mortality in kidney transplantation, the peptic ulcer rebleeding risk in dialysis patients or the risk of end-stage renal disease after hypertensive disorders in pregnancy (see Table 2).

The number of published papers on kidney disease patients differs widely by database, with Taiwan being the leading country in utilizing claims data for clinical research. Data from Taiwan's NHIRD is available for any researcher in Taiwan for a small processing fee, which has been shown to increase the publication rate dramatically. This emphasizes the importance of keeping the financial and technical barriers for the reuse of health claims data for research purposes as low as possible [64].

The Dutch Kidney Atlas

Recently, Dutch health claims data have been processed to study patients with kidney disease in the Dutch Kidney Atlas project. Box 1 provides a detailed description of the Dutch health claims database (called Vektis) and the related healthcare system. The Dutch Kidney Atlas project provides information on the number of CKD patients (CKD Stage G4–G5 [estimated glomerular filtration rate (eGFR) < 30 mL/min/1.73 m²]) not treated with RRT, those on dialysis and kidney transplant patients, their healthcare costs, prescribed medication, outcomes (such as the number of hospital visits, intensive care unit admittance and mortality) and comorbid disorders like diabetes mellitus, hypertension and cardiovascular diseases. Data are published on a website (www.nieratlas.nl) and are reported by age group and sex and are compared with a reference group from the general population. Figure 1 shows several graphs presented in the Dutch Kidney Atlas. Data are presented on a national level as well as on a regional level to demonstrate potential geographic variation. Furthermore, the website includes data since 2012 and will be updated on an annual basis. The website was designed for public use by healthcare professionals, policymakers, researchers and insurance companies, as well as patients with CKD.

The Dutch Kidney Atlas project also involves scientific research on patients with CKD using the Dutch health claims database. So far, two studies on the healthcare costs of patients with CKD with and without RRT have been published [15, 25].

Box 1. Dutch health claims database

- In The Netherlands, healthcare provision and payment for healthcare and healthcare-related services through insurance is embedded within a social security system [65]. Because basic health insurance is obligatory for all Dutch residents, an estimated 99.8% [66] of the Dutch population of ~17 million people has healthcare insurance [67]. The Dutch healthcare system has a gatekeeping principle, which means that patients can easily contact a primary care provider (e.g. general practitioner, dentist, midwife and physiotherapist), but hospital care and specialist care require a referral from a primary care provider, with emergency care as an exception.
- Basic health insurance covers the main aspects of healthcare, including primary care, hospital care, medication, mental healthcare, maternity care and home nursing care. Care not covered by the basic insurance can be insured through voluntary health insurance. Health insurance companies pay the hospital based on Diagnose Behandelcombinatie (DBC)s, a system similar to the concept of diagnosis-related groups. A DBC contains information characterizing the delivered hospital care for a specific medical condition or complaint by type of specialization. The DBC comprises all medical activities needed, from establishing the diagnosis to the last check after treatment, and thereby describing a complete care episode. Every type of DBC has a fixed price, which is the sum of costs of all intermediate products, i.e. the activities, thereby covering all direct and indirect costs of a care episode [68].
- The health claims data of all Dutch health insurance companies are collected in the Vektis database, which covers (almost) all inhabitants of The Netherlands. For each health claim in the Vektis database, data are available on patient characteristics (year of birth, sex, area of residence, socio-economic status and date of death) and the costs involved [16]. Vektis complies with the Dutch law and the European General Data Protection Regulation. To ensure privacy while performing the present research, Vektis pseudonymized the persons’ national identification number and data access is only allowed in a physically secured environment designated by Vektis; only aggregated data are allowed to leave this secured environment. All contributing insurance companies provided permission for the use of these national data.

The Dutch Kidney Atlas project provides detailed information on the number of CKD patients (CKD Stage G4–G5 [estimated glomerular filtration rate (eGFR) < 30 mL/min/1.73 m²]) not treated with RRT, those on dialysis and kidney transplant patients, their healthcare costs, prescribed medication, outcomes (such as the number of hospital visits, intensive care unit admittance and mortality) and comorbid disorders like diabetes mellitus, hypertension and cardiovascular diseases. Data are published on a website (www.nieratlas.nl) and are reported by age group and sex and are compared with a reference group from the general population. Figure 1 shows several graphs presented in the Dutch Kidney Atlas. Data are presented on a national level as well as on a regional level to demonstrate potential geographic variation. Furthermore, the website includes data since 2012 and will be updated on an annual basis. The website was designed for public use by healthcare professionals, policymakers, researchers and insurance companies, as well as patients with CKD.

The Dutch Kidney Atlas project also involves scientific research on patients with CKD using the Dutch health claims database. So far, two studies on the healthcare costs of patients with CKD with and without RRT have been published [15, 25].
| Study types                          | Author                  | Journal               | Year | Country/region | Population | Title                                                                 |
|-------------------------------------|-------------------------|-----------------------|------|----------------|------------|----------------------------------------------------------------------|
| Cost studies                        | Chang et al. [17]       | Nephrology            | 2015 | Taiwan         | Dialysis   | Trends of cost and mortality of patients on haemodialysis with end stage renal disease |
|                                     | Chang et al. [18]       | Sci Rep               | 2016 | Taiwan         | Dialysis   | Cost-effectiveness of haemodialysis and peritoneal dialysis: a national cohort study with 14 years follow-up and matched for comorbidities and propensity score |
|                                     | Couchoud et al. [19]    | Nephrol Dial Transplant | 2015 | France         | Dialysis, transplantation | Economic impact of a modification of the treatment trajectories of patients with end-stage renal disease |
|                                     | Couillerot-Peyrondet et al. [20] | Eur J Health Econ   | 2017 | France         | Dialysis, transplantation | A comprehensive approach to assess the costs of renal replacement therapy for end-stage renal disease in France: the importance of age, diabetes status, and clinical events |
|                                     | Helmuth et al. [21]     | Clin J Am Soc Nephrol | 2019 | USA            | Transplantation | Secular trends in the cost of immunosuppressants after solid organ transplantation in the United States |
|                                     | Honeycutt et al. [22]   | J Am Soc Nephrol      | 2013 | USA            | CKD        | Medical costs of CKD in the Medicare population |
|                                     | Kao et al. [23]         | Perit Dial Int        | 2013 | Taiwan         | Dialysis   | Lifetime costs for peritoneal dialysis and haemodialysis in patients in Taiwan |
|                                     | Kitzawa et al. [24]     | Transplant Proc       | 2017 | Japan          | Transplantation | Cost analysis of transplantation in Japan, performed with the use of the National Database |
|                                     | Mohnen et al. [25]      | PLoS One              | 2019 | Netherlands    | Dialysis, transplantation | Healthcare costs of patients on different renal replacement modalities – analysis of Dutch health insurance claims data |
|                                     | van Oosten et al. [15]  | Nephrol Dial Transplant | 2019 | Netherlands    | CKD, dialysis, transplantation | Age-related difference in healthcare use and costs of patients with chronic kidney disease and matched controls: analysis of Dutch health claims data |
|                                     | Shaikh et al. [26]      | Kidney Int            | 2018 | India          | Dialysis   | Utilization, costs, and outcomes for patients receiving publicly funded haemodialysis in India |
| Descriptive and outcome studies     | Chettiar et al. [27]    | Clin J Am Soc Nephrol | 2018 | USA            | Dialysis, transplantation | Association of inpatient palliative care with health care utilization and postdischarge outcomes among Medicare beneficiaries with end stage kidney disease |
|                                     | Choi et al. [28]        | Am J Nephrol          | 2017 | South Korea    | Dialysis, transplantation | Disparities in kidney transplantation access among Korean patients initiating dialysis: a population-based cohort study using national health insurance data (2003-2013) |
|                                     | Dobbels et al. [29]     | Am J Kidney Dis       | 2008 | USA            | Transplantation | Depressive disorder in renal transplantation: an analysis of Medicare claims |
|                                     | Farrugia et al. [30]    | Kidney Int            | 2014 | UK             | Transplantation | Malignancy-related mortality following kidney transplantation is common |
|                                     | Ferreira et al. [31]    | Nephrol Dial Transplant | 2019 | France         | Dialysis   | Angiotensin-converting enzyme inhibitors/angiotensin receptor blockers, β-blockers or both in incident end-stage renal disease patients without cardiovascular disease: a propensity-matched longitudinal cohort study. |
|                                     | Ferro et al. [32]       | Clinical Transplant   | 2015 | UK             | Transplantation | Fracture risk and mortality post-kidney transplantation |
|                                     | Han et al. [33]         | Clin J Am Soc Nephrol | 2015 | South Korea    | Dialysis   | Dialysis modality and mortality in the elderly: a meta-analysis |
|                                     | Hayer et al. [34]       | Diabetologia          | 2014 | UK             | Transplantation | Infection-related mortality is higher for kidney allograft recipients with pretransplant diabetes mellitus |
|                                     | Hung et al. [35]        | Lancet Diabetes Endocrinol | 2015 | Taiwan         | CKD        | Metformin use and mortality in patients with advanced chronic kidney disease: national, retrospective, observational, cohort study |

(continued)
| Study types | Author | Journal | Year | Country/region | Population | Title |
|------------|--------|---------|------|----------------|------------|-------|
|            | Kim et al. [36] | Int J Cardiol | 2015 | South Korea | Dialysis | Risk of major cardiovascular events among incident dialysis patients: a Korean national population-based study |
|            | Kitchlu et al. [37] | Nephrol Dial Transplant | 2012 | Canada, Ontario | Dialysis | Beta-blockers and cardiovascular outcomes in dialysis patients: a cohort study in Ontario, Canada |
|            | Komenda et al. [38] | CMAJ Open | 2015 | Canada, Manitoba | Dialysis | Secular trends in end-stage renal disease requiring dialysis in Manitoba, Canada: a population-based study |
|            | Kuo et al. [39] | Am J Kidney Dis | 2007 | Taiwan | CKD | Epidemiological features of CKD in Taiwan |
|            | Lam et al. [40] | Transplantation | 2017 | Canada, Ontario | Transplantation | The risk of cardiovascular disease is not increasing over time despite aging and higher comorbidity burden of kidney transplant recipients |
|            | Lenihan et al. [41] | Transplantation | 2019 | USA | Transplantation | Trends in the medical complexity and outcomes of Medicare-insured patients undergoing kidney transplant in the years 1998–2014 |
|            | Li et al. [42] | Nephrol Dial Transplant | 2012 | Taiwan | Transplantation | Malignancies after renal transplantation in Taiwan: a nationwide population-based study |
|            | Liao et al. [43] | Kidney Int | 2015 | Taiwan | Dialysis | Incidence and risk factors for new-onset atrial fibrillation among patients with end-stage renal disease undergoing renal replacement therapy |
|            | René et al. [44] | Nephrol Dial Transplant | 2017 | Canada, Quebec | Dialysis | Association of erythropoiesis-stimulating agents and the incidence of cancer diagnosis among chronic dialysis patients: a nested case–control study |
|            | Tonelli et al. [45] | Kidney Int | 2018 | Canada, Alberta | CKD | A population based cohort study defines prognoses in severe chronic kidney disease |
|            | Wang et al. [46] | Kidney Int | 2019 | China | CKD, dialysis, transplantation | Executive summary for the 2015 Annual Data Report of the China Kidney Disease Network (CK-NET) |
|            | Wang et al. [47] | Am J Kidney Dis | 2014 | Taiwan | Dialysis | Risk of stroke in long-term dialysis patients compared with the general population |
|            | Wang et al. [48] | Clin J Am Soc Nephrol | 2015 | Taiwan | Dialysis | Comparison of subdural haematoma risk between haemodialysis and peritoneal dialysis patients with ESRD |
|            | Wang et al. [49] | Nephrol Dial Transplant | 2018 | Taiwan | Dialysis | Risk of new-onset diabetes in end-stage renal disease patients undergoing dialysis: analysis from registry data of Taiwan |
|            | Weinhandl et al. [50] | Am J Nephrol | 2019 | USA | Dialysis | Contemporary trends in clinical outcomes among dialysis patients with Medicare coverage |
|            | Wu et al. [51] | Gut | 2011 | Taiwan | Dialysis | Long-term peptic ulcer rebleeding risk estimation in patients undergoing haemodialysis: a 10-year nationwide cohort study |
|            | Wu et al. [52] | Am J Obstet Gynecol | 2014 | Taiwan | Dialysis | End-stage renal disease after hypertensive disorders in pregnancy |
|            | Yoon et al. [53] | Stroke | 2017 | South Korea | Dialysis | Warfarin use in patients with atrial fibrillation undergoing haemodialysis |
|            | Yu et al. [54] | Lancet Oncol | 2016 | Taiwan | Polycystic kidney disease | Risk of cancer in patients with polycystic kidney disease: a propensity-score matched analysis of a nationwide, population-based cohort study |

Validation studies | For more details see Table 3
IDENTIFICATION OF PERSONS WITH KIDNEY DISEASE

Administrative databases are the result of administrative processes, such as reimbursement in the case of health claims databases, and are therefore not designed for clinical research purposes [16]. Because clinical data and information on renal function are lacking in health claims databases, the identification of patients with kidney disease depends on diagnosis codes only, which in turn depends on the proper registration of the codes by the involved healthcare professional or organization. Investigating the accuracy of this identification is therefore crucial to assess whether data derived from health claims data are truly representative and this information on the validity of the diagnosis codes should be provided in research articles using health claims data [58, 69]. Some studies assessed the validity of health claims data to identify patients with CKD, dialysis or kidney transplantation (Table 3). Apart from one study using Dutch health claims data, all studies were performed in Canada or the USA. Of all available studies, five studied the validity of health claims data in the identification of CKD patients, three of dialysis patients and one of kidney transplant patients. However, the use of different case definitions and reference populations does impede the direct comparison between studies. In the following paragraphs we will discuss the studies that have assessed the validity of health claims data to identify patients with CKD, dialysis or kidney transplantation in more detail.

CKD

A study in Alberta, Canada, tested health claims data for the identification of CKD (defined as an eGFR <60 mL/min/1.73 m²) against a gold standard derived from outpatient serum creatinine measurements. In this study, 19% of CKD patients were correctly identified as such with health claims data (sensitivity or true-positive rate) and 60% of the patients with CKD-related claims data did have an eGFR <60 mL/min/1.73 m² [positive predictive value (PPV) 60%] [57]. Similar results were found in a study from Ontario, Canada, using patients with a serum creatinine laboratory test following a prescription of medication as the gold standard (sensitivity 18%, PPV 65%) [55]. In both studies, sensitivity was markedly higher for CKD patients defined as an eGFR <30 mL/min/1.73 m² (65% and 59%, respectively). Two US studies tested the validity of Medicare data. One study used patients with hospitalization for myocardial infarction as the gold standard (sensitivity 27%, PPV 89%) [58], while the second study used research study measurements as a reference (sensitivity 16%, PPV 76%) [56]. Recently we tested the validity of Dutch health claims data using a laboratory database as the gold standard. Sensitivity was markedly higher in patients with advanced CKD (eGFR <30 mL/min/1.73 m²) than in patients with CKD (eGFR <60 mL/min/1.73 m²), being 51% and 27%, respectively [59]. All studies had high specificity for CKD. The negative predictive value (NPV) varied in all studies and with a wide range (36–98%).
Table 3. Overview of studies of the validity of health claims data in the identification of CKD, dialysis and kidney transplant patients

| Author       | Health claims database                                      | Study Population                                                                 | Reference population                                                                 | Age          | Case definition                                                                 | Sensitivity (95% CI) | Specificity (95% CI) | PPV (95% CI)   | NPV (95% CI)   |
|--------------|------------------------------------------------------------|-----------------------------------------------------------------------------------|--------------------------------------------------------------------------------------|--------------|----------------------------------------------------------------------------------|----------------------|----------------------|----------------|----------------|----------------|
| Fleet et al. | Ontario Health Insurance Plan database (OHIP), Ontario (Canada) | Patients with ICD-10 and physician claims diagnostic codes for CKD, between 1 July 2007 and 31 December 2010 | Patients with an outpatient prescription medication and a serum creatinine test the year prior to the prescription date from a laboratory in Southwestern Ontario | ≥65 years    | eGFR < 60 mL/min/1.73 m² or ACR > 30 mg/g | 18.0 (17.7–18.4) | 98.2 (98.1–98.3) | 85.2 (84.5–85.9) | 67.7 (67.4–68.0) |
|              |                                                            |                                                                                   |                                                                                      |              | eGFR < 45 mL/min/1.73 m²                                                        | 32.7 (32.0–33.3) | 96.9 (96.7–97.0) | 65.4 (64.4–66.3) | 88.8 (88.6–89.0) |
|              |                                                            |                                                                                   |                                                                                      |              | eGFR < 30 mL/min/1.73 m²                                                        | 58.8 (57.4–60.1) | 94.6 (94.5–94.7) | 32.5 (31.6–33.5) | 98.1 (98.0–98.2) |
| Muntner et al. | Medicare claims data, USA | Patients with ICD-9 discharge codes associated with hospitalization or physician evaluation and claims associated with outpatient physician visits for CKD, between January 2003 and October 2007 | Participants enrolled in the Reasons for Geographic and Racial Differences in Stroke (REGARDS) study with data available on eGFR and albumin–creatinine ratio | ≥65 years    | eGFR < 60 mL/min/1.73 m² or ACR > 30 mg/g | 15.5 (14.0–17.1) | 97.7 (97.2–98.1) | 75.6 (71.4–79.5) | 71.5 (70.4–72.6) |
|              |                                                            |                                                                                   |                                                                                      |              | eGFR < 60 mL/min/1.73 m² or ACR > 30 mg/g | 20.6 (18.5–22.8) | 97.1 (96.6–97.5) | 63.9 (59.2–68.3) | 83.0 (82.1–83.9) |
|              |                                                            |                                                                                   |                                                                                      |              | eGFR < 45 mL/min/1.73 m²                                                        | 37.1 (32.7–41.6) | 95.8 (95.3–96.2) | 39.0 (34.5–43.7) | 95.4 (94.9–95.9) |
|              |                                                            |                                                                                   |                                                                                      |              | eGFR < 30 mL/min/1.73 m²                                                        | 56.4 (45.8–66.6) | 94.2 (93.6–94.8) | 11.8 (8.9–15.1) | 99.4 (99.1–99.5) |
| Ronksley et al. | Alberta Provincial Physician Claims database, Alberta (Canada) | Patients with ICD-9 and ICD-10 codes for CKD, between 1 January 2004 and 31 December 2004 | Patients with at least two outpatient serum creatinine measurements within a 1-year time period | ≥18 years    | eGFR < 60 mL/min/1.73 m²                                                        | 18.9 (–)        | 97.29 (–)         | 60.59 (–)     | 83.99 (–)    |
|              |                                                            |                                                                                   |                                                                                      |              | One claim or one hospitalization in one year                                      | 14.29 (–)       | 98.19 (–)         | 63.69 (–)     | 83.39 (–)    |
|              |                                                            |                                                                                   |                                                                                      |              | Two claims or one hospitalization in 1 year                                      | 11.89 (–)       | 98.59 (–)         | 64.09 (–)     | 82.99 (–)    |
|              |                                                            |                                                                                   |                                                                                      |              | eGFR < 30 mL/min/1.73 m²                                                        | 64.79 (–)       | 96.59 (–)         | 24.09 (–)     | 99.49 (–)    |
|              |                                                            |                                                                                   |                                                                                      |              | One claim or one hospitalization in one year                                      | 56.59 (–)       | 97.79 (–)         | 29.39 (–)     | 99.29 (–)    |
|              |                                                            |                                                                                   |                                                                                      |              | Two claims or one hospitalization in 1 year                                      | 49.99 (–)       | 98.19 (–)         | 31.49 (–)     | 99.19 (–)    |

(continued)
| Author                  | Health claims database                      | Study Population | Reference population | Age          | Case definition                                                                 | Sensitivity (95% CI) | Specificity (95% CI) | PPV (95% CI) | NPV (95% CI) |
|-------------------------|---------------------------------------------|------------------|----------------------|--------------|----------------------------------------------------------------------------------|----------------------|----------------------|--------------|--------------|
| Winkelmayer et al. [58] | Medicare claims data, USA                    | Patients with ICD-9 diagnosis codes for CKD, during 1999 and/or 2000 | Patients with hospitalization for myocardial infarction with a serum creatinine measurement | ≥65 years    | eGFR < 60 mL/min/1.73 m² for 6-months period                                      | 20.7 (18.5–22.9)     | 96.0 (94.4–97.5)     | 91.6 (88.5–94.8) | 36.3 (34.0–38.7) |
|                         |                                             |                   |                      |              | eGFR < 60 mL/min/1.73 m² for 12-month period                                      | 26.6 (24.2–28.9)     | 93.3 (91.3–95.2)     | 89.3 (86.3–92.4) | 37.4 (35.0–39.8) |
| van Oosten et al. [59]  | Vektis database, The Netherlands              | Patients with DBC codes for CKD, between 1 January 2014 and 31 December 2014 | Patients with an outpatient serum creatinine measurement in 2014 | ≥18 years    | One eGFR < 60 mL/min/1.73 m²                                                      | 20 (19–21)           | 100 (100–100)       | 96 (95–97)  | 84 (83–84)   |
|                         |                                             |                   |                      |              | At least two eGFR < 60 mL/min/1.73 m² at least 90 days apart                    | 27 (25–28)           | 100 (100–100)       | 90 (88–91)  | 52 (51–52)   |
|                         |                                             |                   |                      |              | One eGFR < 30 mL/min/1.73 m²                                                     | 42 (38–46)           | 100 (100–100)       | 83 (79–87)  | 98 (98–99)   |
|                         |                                             |                   |                      |              | At least two eGFR < 30 mL/min/1.73 m² at least 90 days apart                    | 51 (47–56)           | 100 (99–100)       | 80 (76–84)  | 98 (98–99)   |
| Dialysis                |                                             |                   |                      |              |                                                                                  |                      |                      |              |              |
| Clement et al. [60]     | Alberta Provincial Physician Claims database, Alberta (Canada) | Patients with physician claims for outpatient dialysis, between 1 January 2008 and 31 December 2008 | ESRD registry [Northern Alberta (NARP) and Southern Alberta (SARP) registries] | ≥18 years    | 1. At least one claim                                                            | 81.1 (–)             | NA       | 77.7 (–)  | NA           |
|                         |                                             |                   |                      |              | 2. At least two claims                                                            | 78.6 (–)             | NA       | 80.7 (–)  | NA           |
|                         |                                             |                   |                      |              | 3. At least two claims at least 90 days apart                                     | 63.1 (–)             | NA       | 84.8 (–)  | NA           |
|                         |                                             |                   |                      |              | 4. Continuous claims at least 90 days apart with no gaps in claims >21 days      | 58.2 (–)             | NA       | 85.9 (–)  | NA           |
| Komenda et al. [61]     | Manitoba Health Physician Claims database, Manitoba (Canada) | Patients with physician claims for outpatient dialysis, between 1 January 2004 to 31 December 2010 | Manitoba Renal Program Dialysis Registry | ≥18 years    | 1-year period (2010)                                                            | 77.0 (74.7–79.2)      | 93.8 (92.9–94.7)     | 85.2 (83.2–87.2) | 89.8 (88.7–90.9) |
|                         |                                             |                   |                      |              | 2. Any two claims                                                                | 74.6 (72.3–76.9)      | 94.4 (93.6–95.2)     | 86.0 (84.0–88.0) | 88.9 (87.8–90.0) |
|                         |                                             |                   |                      |              | 3. Any two claims at least 90 days apart                                         | 64.8 (62.2–67.3)      | 97.1 (96.5–97.7)     | 91.2 (89.5–93.0) | 85.6 (84.4–86.8) |
|                         |                                             |                   |                      |              | 4. Any two claims at least 90 days apart with no gaps in treatment >21 days       | 52.7 (50.1–55.4)      | 97.5 (96.9–98.1)     | 90.7 (88.7–92.7) | 81.7 (80.4–83.0) |
|                         |                                             |                   |                      |              | 5-year period (2004–2008)                                                        |                      |                      |              |              |
|                         |                                             |                   |                      |              | 1. At least one claim                                                            | 87.6 (86.3–89.0)      | 91.3 (90.7–91.9)     | 96.2 (95.8–96.7) | 95.9 (95.4–96.3) |
|                         |                                             |                   |                      |              | 2. Any two claims                                                                | 86.0 (84.7–87.4)      | 93.4 (92.9–93.9)     | 78.9 (77.4–80.4) | 95.9 (95.4–96.3) |

(continued)
| Author | Health claims database | Study Population | Reference population | Age | Case definition | Sensitivity (95% CI) | Specificity (95% CI) | PPV (95% CI) | NPV (95% CI) |
|--------|------------------------|------------------|---------------------|-----|----------------|---------------------|---------------------|--------------|--------------|
| Taneja et al. [62] | Health Alliance Plan (HAP) database, USA | Patients with ESRD and dialysis-related billing codes for peritoneal dialysis or haemodialysis, between 1 January 2005 and 31 December 2008. | Patient medical record | 18–63 years | Any PD-related claim—in a 30-day window | 72.0 (70.2–73.8) | 99.6 (99.5–99.8) | 98.3 (97.7–98.9) | 92.5 (92.0–93.1) |
| | | | | | Any PD-related claim—in a 90-day window | 47.6 (45.6–49.6) | 99.8 (99.7–99.9) | 98.3 (97.6–99.0) | 86.9 (86.2–87.5) |
| Kidney transplantation | Lam et al. [63] | Ontario Health Insurance Plan (OHIP) database (OHIP), Ontario (Canada) | Patients with kidney transplantation related claims, between 1 January 2008 and 31 December 2011 | | Any PD-related claim—in a 180-day window | NA | NA | 34.9 (–) | NA |
| | | | | | Any HD-related claim—in a 30-day window | NA | NA | 67.4 (–) | NA |
| | | | | | Any HD-related claim—in a 90-day window | NA | NA | 67.4 (–) | NA |
| | | | | | Any HD-related claim—in a 180-day window | NA | NA | 86.7 (–) | NA |
| | | | | | A claim for a kidney-only transplant | 98 (97–99) | NA | 96 (95–97) | NA |

CI, confidence interval; ICD, International Classification of Diseases.
Health claims data have low sensitivity for the estimation of the overall CKD prevalence in the general population, since health claims data only detect CKD patients treated in a hospital and registered for the specific Diagnose Behandelcombinaties (DBCs) and not the ones treated by a general practitioner or those who are not treated at all. However, they do, to a large extent, reflect the population of CKD with an actual referral to a nephrologist.

Dialysis
All available studies indicate that identifying patients undergoing dialysis with health claims data is accurate. A study in Ontario, Canada, showed a reasonably good identification of dialysis patients with a sensitivity of 81% and a PPV of 78% when compared to a gold standard of registry data [60]. These results were confirmed in patients in Manitoba, Canada (sensitivity 77%, PPV 85%), also using registry data as the gold standard [61]. A US study accurately identified haemodialysis using health claims data compared with medical records data as the gold standard (PPV 91%), but was less precise for patients treated with peritoneal dialysis (PPV 67%) [62]. In The Netherlands, the number of dialysis (haemodialysis and peritoneal dialysis) patients identified with Dutch health claims data was compared with the number of patients in the Dutch registry for end-stage renal disease [RENINE (Registratie Nierfunctievervanging Nederland)]. Since the analysis was only possible on aggregated data, sensitivity could not be calculated, but the correspondence between the databases was very high (99%) [25].

Kidney transplantation
Identification of performed kidney transplantations was shown to be very accurate using Canadian health claims data compared with data from three Canadian transplantation centres serving as the gold standard (sensitivity 98%, PPV 96%) [63]. Also, Dutch health claims data were shown to very accurately identify the number of performed kidney transplantations per year when compared with the Dutch registry for end-stage renal disease, with a correspondence of 99% between the databases [25].

In RRT patients, who are generally all treated within the standard healthcare system, both the PPV and NPV are high. This makes it possible to compare outcomes between patients with and without RRT. In contrast, not all affected CKD patients (not treated with RRT) are known or treated within the (hospital) care system. Therefore, in CKD patients, the PPV is high, but the NPV generally is low. In CKD patients, it may therefore be more difficult to identify those without CKD using health claims data [58]. Please note that both the PPV and NPV depend on the prevalence of the disease in the population [70].

OPPORTUNITIES AND CHALLENGES OF HEALTH CLAIMS DATA
Health claims databases have strengths and limitations, depending on the research question that investigators would like to address [71].

Opportunities
Claims databases have several specific advantages over other types of research data, regarding scope, flexibility, costs and statistical power [71]. Regarding the scope of health claims databases, they are generally very comprehensive and complete and often cover the inhabitants of an entire country or region. These databases usually contain data on demographics (e.g. age, sex and postal code), prescribed medication, diagnosis, hospital care and other delivered healthcare, including a complete care pathway of a patient from the first contact with a nephrologist until the last treatment. Since health claims data are collected for payment purposes, they contain data on healthcare costs. Moreover, with health claims data, it is possible to study CKD patients treated with and without RRT in the same data set. Health claims databases often contain data over longer periods of time as well as geographic data, making it possible to study regional differences. Furthermore, since health claims data are collected on a routine basis, they provide information on a non-experimental setting.

The flexibility of health claims data provides opportunities to use different study designs (e.g. cohort studies and case-control design). In addition, for investigators, the data collection is relatively inexpensive and less time-consuming compared with other data collections such as randomized controlled trials or cohort studies. Considering this, with health claims data it is relatively easy to obtain a sufficient number of cases and provide adequate statistical power at relatively low cost [71]. These unique features of health claims databases make it possible to monitor trends in disease prevalence, treatment or healthcare costs over time, providing insight into the effect of changes in policy or guidelines. Therefore health claims data could play a valuable role in guiding health policy and improving quality of care, with the Alberta Kidney Disease Network as an example of a unique collaboration between researchers and policymakers [4]. In addition, health claims data could potentially be used as a quality indicator without providing extra administrative burden for the caregiver, as outcomes can be traced back to individual healthcare providers. For instance, the number of cardiovascular complications in CKD patients, identified using health claims data, can be used to comment on the quality of cardiovascular care on the condition that one can adjust for patient case mix.

Challenges
Health claims data were not designed for clinical research and therefore the researcher cannot control the design, collection and processing of data [16]. Studies have shown that administrative databases, including health claims databases, have limitations in scope (availability of relevant data), data quality and the ability to adjust for patient case mix [72]. Since the majority of health claims databases lack both clinical and laboratory data, the identification of patients is based on specific diagnosis or procedure codes. As a result, the identification of patient groups with health claims data may result in undercoding or overcoding of diagnoses or outcomes [58, 73]. In a database designed for reimbursement purposes, this undercoding or overcoding can be related to coding optimization (i.e. a diagnosis or procedure with higher reimbursement fees is more likely to be coded than the one with lower reimbursement fees). In addition, as previously discussed in the CKD validation studies, usually health claims data are only able to identify CKD patients who are treated by nephrologists while patients who do not come to the attention of health services remain undetected with health claims data. This may be an important limitation in countries that do not have universal healthcare coverage.
Furthermore, the lack of clinical information like severity and progression of the disease, clinical parameters (e.g. smoking and bodyweight) or risk factors might lead, for some research questions, to an incomplete adjustment for potential confounders [71]. Next, in countries without universal health care coverage, such as the USA, elderly individuals or those with lower socio-economic status, may be overrepresented in the population [71].

There are several ways to deal with these challenges of health claims data. First, part of the lacking information with regard to the prevalence of chronic diseases or morbidity in CKD patients may be derived from specific medication use. Studies show that data on prescribed medication can be used as a proxy for the prevalence of several chronic diseases [74]. A proxy can be very valid, for example, in case of identifying patients with diabetes mellitus using anti-diabetic drugs and insulin analogues, but less valid if drugs have overlapping indications (e.g. inhalation therapy prescribed for patients with asthma and those with chronic obstructive pulmonary disease).

Second, record linkage of other unrelated administrative databases to health claims databases is a promising tool to add value to the data and it improves and broadens their usage for health research [16]. Record linkage, by means of a unique and direct identifier (i.e. social security number or NHS number), is increasingly used worldwide to combine administrative datasets (e.g. the Alberta kidney disease database) [4]. In case this unique identifier is lacking, indirect linkage tools could be a valid option, although it may introduce less precise record matching [75]. In many countries, maintainers of administrative health databases, such as governmental agencies and health insurance companies, are often reluctant to share administrative databases. The general data protection rules, among others, underlie the restrictive sharing and linking of health data. Therefore the World Health Organization advocates the development of a metadata standard to improve data-sharing policies, thereby increasing the research potential of routinely collected health datasets [76]. We stress the importance of improving the utility of claims data while protecting confidentiality.

Nevertheless, when comparing the results of different health claims databases one must bear in mind the differences in characteristics of the study populations, the regional differences in insurance coverage and the registration of diagnosis or health care use. This might limit the extrapolation of the results to other countries. Apart from important privacy issues this also limits the possibilities of merging international databases. In addition, coding and coverage of diagnoses, procedures or treatments may change over time. One should be aware of these possible changes within a healthcare system when comparing health claims data over a longer period of time.

CONCLUSION

Health claims databases offer important opportunities for studies on large populations of patients with (kidney) disease and health outcomes in a non-experimental setting. However, one should take into account the limitations of health claims data and consider the characteristics of a health claims database, especially when making international comparisons. Since research with health claims data uses codes to identify kidney disease patients and to define other key study variables, information on the validity of these codes in measuring the association of the code with the real variable is indispensable. Available studies indicate that identifying patients undergoing dialysis and the number of performed kidney transplantations using health claims data is accurate, whereas health claims data have low sensitivity for the estimation of the overall CKD prevalence in the general population.

So far, health claims data in 10 countries have been used for studies on kidney disease patients. The unique features of health claims data provide specific research opportunities, such as studying healthcare costs or studying longitudinal, epidemiological data with nationwide coverage. For the optimal utility of health claims data, it is important to keep financial and technical barriers low, while protecting confidentiality. In addition, health claims data can be used to create a nationwide atlas (e.g. the Dutch Kidney Atlas) providing national and regional information on, for instance, the numbers, healthcare costs, prescribed medications, treatments and outcomes of kidney patients.

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CONFLICT OF INTEREST STATEMENT

None declared.

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