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Adherence to guidelines for creatinine and potassium monitoring and discontinuation following renin-angiotensin system blockade

Running title: Creatinine and potassium monitoring after ACEI/ARB initiation

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ABSTRACT

Objectives To examine adherence to serum creatinine and potassium monitoring and discontinuation guidelines following initiation of treatment with angiotensin converting-enzyme inhibitors (ACEI) or angiotensin-receptor blockers (ARB); and whether high-risk patients are monitored.

Design Population-based cohort study using electronic health records from the UK Clinical Practice Research Datalink and Hospital Episode Statistics.

Setting UK primary care, 2004–2014.

Subjects 223,814 new ACEI/ARB users.

Main outcome measures Proportion of patients with renal function monitoring before and after ACEI/ARB initiation; adverse renal outcomes (creatinine increase $\geq 30\%$ or potassium levels >6 mmol/L) at first follow-up monitoring; and treatment discontinuation after such changes. Using logistic regression models, we also examined patient characteristics associated with such adverse outcomes, and with follow-up monitoring within the guideline-recommend two weeks of treatment initiation.

Results Ten percent of patients had neither baseline nor follow-up monitoring of creatinine within 12 months before and 2 months after initiation of an ACEI/ARB, 28% had monitoring only at baseline, 15% only at follow-up, and 47% both at baseline and follow-up. The median period between the most recent baseline monitoring and drug initiation was 40 days (interquartile range: 12-125 days). 34% of patients had baseline creatinine monitoring within one month before initiating therapy, but less than 10% also had the guideline-recommended follow-up test recorded within two weeks. Among patients experiencing a creatinine increase $\geq 30\%$ (n=567, 1.2%) or potassium level >6 mmol/L (n=191, 0.4%), 80% continued treatment. Although patients with prior myocardial infarction, hypertension, or baseline potassium >5 mmol/L were at high risk of adverse renal outcomes after ACEI/ARB initiation, there was no evidence that they were more frequently monitored.

Conclusions Only one tenth of patients initiating ACEI/ARB therapy receive the guideline-recommended creatinine monitoring. Moreover, the vast majority of the patients fulfilling post-initiation discontinuation criteria for creatinine and potassium increases continue on treatment.

Strengths and limitations of this study:

- This is the largest monitoring study to date, examining both adherence to creatinine and potassium monitoring and discontinuation guidelines following initiation of angiotensin converting-enzyme inhibitors or angiotensin-receptor blockers in UK primary care, and whether patients are monitored in accordance with their individual risk profile
- Use of the UK Clinical Practice Research Datalink and Hospital Episode Statistics ensured that the study was population-based and not restricted to specific demographic, hospital, or insurance groups.
- Blood tests performed in hospital systems were not recorded in the Clinical Practice Research Datalink, but the results were consistent for patients with no recent hospital admissions
- If the recording of creatinine levels was not missing completely at random, the associations between patient characteristics and creatinine increase may have been underestimated

INTRODUCTION

Renin angiotensin system blockade using angiotensin-converting enzyme inhibitors (ACEI) and angiotensin-receptor blockers (ARB) is a mainstay in treatment of hypertension,¹ heart failure,² diabetic microalbuminuria or proteinuric renal diseases,³ and after myocardial infarction.⁴ However, some patients experience a sudden decline in kidney function when initiating these drugs, presumably due to antagonism of the angiotensin II-mediated efferent arteriolar constriction or impaired kidney excretion of potassium.^{5 6}

The potential impact on kidney function should be evaluated by comparing pre- and post-initiation levels of serum creatinine and potassium.⁷ Discontinuation is recommended if the rise in creatinine exceeds 30% above baseline or if hyperkalaemia develops.⁸ It is unclear whether these recommendations are routinely followed in clinical practice.⁹

A few studies have compared baseline and follow-up monitoring results,⁹ but large studies using contemporary data with reference to current guidelines are lacking, and it is unknown whether patients' individual risk of renal impairment influence their likelihood of being monitored.⁹ We therefore examined adherence to creatinine and potassium monitoring and treatment discontinuation guidelines following ACEI/ARB initiation in UK primary care, and whether patients are monitored in accordance with their individual risk profile.

METHODS

Data sources

We used the UK's Clinical Practice Research Datalink (CPRD) linked to hospital record data from the Hospital Episode Statistics (HES) database. The CPRD database contains primary care electronic health record data from 7% of the UK population (~15 million patient lives, with ~8 million currently under follow-up).¹⁰ Patients included in the CPRD are largely representative of the UK population in terms of age, sex and ethnicity.^{10 11} Information recorded in the database includes demographics such as sex and year of birth, the location of the general practice, medical diagnoses (based on 'Read' codes), drug prescriptions, and a

range of routine laboratory test results. HES records cover all hospital admissions for patients covered by the NHS who receive treatment either from English NHS trusts or independent providers.^{10 11} Fifty-eight percent of general practices included in the CPRD have agreed to HES linkage.¹⁰ We obtained linked data on socioeconomic status (index of multiple deprivation) based on area of residence.

Monitoring guidelines

Consistent with other international guidelines, the National Institute for Health and Care Excellence (NICE) recommend baseline testing of creatinine when initiating ACEI/ARB therapy in patients with hypertension,¹ heart failure,² myocardial infarction,⁴ or chronic kidney disease (CKD).³ The time interval for baseline testing is not further specified.¹⁻⁴ Among patients with heart failure, myocardial infarction, and chronic kidney disease, NICE recommends follow-up monitoring within 2 weeks of treatment initiation,²⁻⁴ and for myocardial infarction patients at least annually thereafter.⁴ A baseline assessment and follow-up test within 2 weeks are also recommended by the UK Renal Association,¹² as well as the frequently used online web resource General Practice (GP) Notebook.¹³ GP Notebook additionally recommends monitoring 1, 3, 6, and 12 months after the first follow-up test.¹³ NICE recommends not to initiate ACEI/ARBs in patients with a baseline potassium level >5 mmol/L and to discontinue therapy if potassium rises above 6 mmol/L.

ACEI/ARB initiators

We identified a cohort of all HES linked CPRD patients aged ≥ 18 years, who initiated ACEI/ARB treatment between January 1, 2004 and March 31, 2014. We did not include earlier calendar periods, as laboratory data before 2004 were incomplete due to interface problems between laboratory reporting software and GP practice software.¹⁴ Also, creatinine testing was incentivised in 2004 with the introduction of the diabetes Quality and Outcomes

Framework (QOF) and further in 2006 with the CKD QOF.¹⁴ To rule out any potential influence of incomplete data around 2004, we also examined the most recent 5-year calendar period separately in a sensitivity analysis. New users were defined as persons with at least one year of continuous registration in the CPRD before their first recorded ACEI/ARB prescription.

Laboratory data

All creatinine test results were extracted from the general practice records of the study population, using creatinine-specific codes in CPRD. Cross-reference was then made to creatinine test results identified from a broad Read code search. Any irrelevant codes were excluded. The same procedure was used to identify potassium test results.

Patient characteristics

We obtained information for all patients on age, sex, calendar period of ACEI/ARB initiation (2004–2008 and 2010–2014), socioeconomic status (quintiles of the 2004 index of multiple deprivation scores), lifestyle factors (smoking, alcohol intake, and body mass index), baseline potassium level (≤ 5 or > 5 mmol/L), CKD, cardiovascular comorbidities (heart failure, myocardial infarction, hypertension, peripheral arterial disease, and arrhythmia), and diabetes.¹⁵ We used algorithms for smoking status, alcohol intake, and body mass index based on the most recent records in the CPRD before ACEI/ARB initiation.^{16 17} As measures of baseline creatinine and potassium levels, we used the single most recent measurement within 12 months before the first ACEI/ARB prescription. We calculated estimated glomerular filtration rate (eGFR) level from the most recent creatinine measurement and CKD stage from the CKD-EPI equation.¹⁸ Cardiovascular comorbidities and diabetes were identified from both the CPRD and HES based on diagnoses recorded prior to ACEI/ARB initiation. The code lists for all variables are provided in the online repository at

<https://clinicalcodes.rss.mhs.man.ac.uk/>.¹⁹

Patient involvement

The study included no patient involvement

Statistical analysis

We described ACEI/ARB users according to patient characteristics, both overall and according to creatinine monitoring status (no baseline or follow-up monitoring, baseline only, follow-up only, and both baseline and follow-up monitoring). Baseline monitoring was defined as a test performed on the date of drug initiation or within either 12 months before (generous interval) or one month before initiation (more ideal interval assumed to be driven by planned ACEI/ARB initiation). To accord with the post-initiation monitoring interval recommended from previous trial data, we considered only follow-up monitoring within the first 2 months after drug initiation.⁸

We calculated the proportion of persons in the total cohort of new users who had baseline and follow-up monitoring (within 1, 3, and 12 months before drug initiation and within 2 weeks, 1 month, and 2 months after initiation). We then computed the proportion of persons with both baseline and initial follow-up monitoring within the guideline-recommended interval of 2 weeks following drug commencement.

We repeated the analyses for continuing users, in order to examine adherence to the stricter guideline recommendations for ongoing monitoring (*i.e.*, monitoring within 1, 3, 6, and 12 months after the first retest).¹³ Continuation was defined as ACEI/ARB use beyond 30 days following the monitoring date, *i.e.*, when the end date of the first continuous course of therapy was after the date of the first monitoring date plus 30 days (to allow for stockpiling). The end date of each prescription was calculated by adding the prescription duration (total number of tablets prescribed divided by the specified number of tablets per day) to the

prescription date. In identifying continuous courses of therapy, we allowed for a 30-day gap between the end date of one prescription and the start of the next consecutive prescription.

In sensitivity analyses, we repeated the analyses (1) extending the follow-up window for the first follow-up monitoring from two to three weeks to account for minor delays; (2) including only the most recent calendar period (2009-2014) to account for temporal changes in data completeness and quality of care; (3) excluding patients with a hospital admission or discharge date within 1 month before or after their first ACEI/ARB prescription, in order to account for drug initiation and any subsequent renal function tests occurring in the hospital and therefore not captured in the CPRD; (4) focusing on specific patient subgroups (heart failure, myocardial infarction, hypertension, CKD (eGFR<60mls/min/1.73m²), peripheral arterial disease, and diabetes); and (5) defining drug use continuation as ACEI/ARB use beyond 90 days (instead of 30 days) after the first retest date.

We used the subcohort of patients with both baseline and follow-up monitoring to calculate the proportion of patients with creatinine increases $\geq 30\%$ or potassium levels >6 mmol/L at the first follow-up monitoring within 2 months after initiation, as well as the proportion of patients continuing treatment despite these contraindications for use.

Finally, we fitted a logistic regression model to identify patient characteristics associated with a severe decline in renal function (creatinine increase $\geq 30\%$ or potassium level >6 mmol/L) and compared these characteristics with those associated with receiving post-initiation follow-up monitoring within 2 weeks. The model included age, sex, CKD stage, cardiovascular comorbidities, diabetes, and baseline potassium level (>5 vs. ≤ 5 mmol/L). In three additional model-based sensitivity analyses, we repeated the analyses (1) excluding patients with a recent hospitalization (as defined above); (2) omitting baseline potassium from the model to examine the extent of potential overfitting when both baseline potassium and CKD stage were kept in the model; and (3) also adjusting additional for ethnicity.

The study protocol was made available to the journal reviewers and approved by the London School of Hygiene and Tropical Medicine Ethics Committee (No. 6536) and the Independent Scientific Advisory Committee (ISAC) for Medicines and Healthcare Products Regulatory Agency (No. 16_025). All analyses were performed using STATA 14 statistical software package.

RESULTS

Serum creatinine monitoring before and after ACEI/ARB initiation

We identified 223,814 new users of ACEI/ARBs (Table 1). Among these patients, 21,411 (10%) had no baseline or follow-up creatinine tests within 12 months before and 2 months after treatment initiation, 63,359 (28%) had only a baseline test, 33,185 (15%) had only follow-up tests, and 105,859 (47%) had both baseline and follow-up tests. Median age varied only slightly between the groups (60, 62, 59, and 63 years, respectively) and there were no substantial differences in socioeconomic status, lifestyle factors, or peripheral arterial disease. Compared with patients with neither pre- nor post-initiation monitoring, patients with both were more likely to have diagnosed hypertension (76% vs. 61%) and diabetes (20% vs. 7%), but less likely to have diagnosed heart failure (4% vs. 7%), myocardial infarction (4% vs. 18%), and arrhythmia (7% vs. 10%). Among patients with baseline monitoring, 83% did not have CKD, 13% stage 3a, 3% stage 3b, 0.5% stage 4 CKD. In the same population, 7% commenced ACEI/ARB therapy despite baseline potassium above 5 mmol/L. The median number of days between baseline monitoring and first prescription date was 40 days (interquartile range: 12-125 days).

The proportion of patients receiving creatinine testing before initiating ACEI/ARB therapy was 76% within 12 months before treatment initiation, declining to 34% within one month before initiation (Table 2). The proportion with follow-up testing after treatment initiation was 29% within 2 weeks, increasing to 62% within 2 months. 21% of ACEI/ARB initiators had both a baseline test within 12 months before and a follow-up test within 2

weeks after starting treatment (Table 3). However, among patients undergoing testing within one month prior to treatment initiation, only 9% had also the recommended follow-up test within 2 weeks of treatment start. When we extended the follow-up window to three weeks, this proportion increased only to 14% (Table 3). Among patients continuing treatment, only 1% had follow-up measurements at 1, 3, 6, and 12 months after the first retest, in compliance with the strictest recommendation (eTable 1). These results were unchanged when the analysis was restricted to the most recent calendar period (eTable 1-2) and to patients with heart failure, myocardial infarction, hypertension, peripheral arterial disease, diabetes or no recent hospitalization (eTable 3). Only patients with CKD received a slightly higher degree of monitoring (13%) within two weeks following treatment initiation (eTable 3).

Serum creatinine and potassium changes after ACEI/ARB initiation

Among patients receiving the recommended renal function monitoring, 567 (1.2%) experienced a creatinine increase $\geq 30\%$ and 191 (0.4%) a potassium level >6 mmol/L at their first follow-up test within two months of treatment initiation (1.4% received in total either) (Table 4). Among these patients, 80% continued treatment beyond 30 days following the monitoring date (Table 4). The sensitivity analysis showed that 65% of patients with a creatinine increase $\geq 30\%$ and 60% of those with a potassium level >6 mmol/L continued also treatment beyond 90 days after the monitoring date (eTable 4). The results remained consistent for longer baseline monitoring intervals (eTable 4).

Patients at high risk for creatinine increases $\geq 30\%$

When we examined patient characteristics associated with a creatinine increase $\geq 30\%$ and adjusted for the other characteristics in a multivariable analysis (Table 5), we found an increased odds ratio (OR) for women (1.6-fold increased), for age above 70 years (at least 1.3-fold increased), for CKD stage 4 (1.6-fold increased), heart failure (2.9-fold increased),

peripheral arterial disease (1.9-fold increased), myocardial infarction (1.6-fold increased), and hypertension (1.6-fold increased).

Patients at high risk for potassium >6 mmol/L

Baseline potassium level and CKD stage, but not age and sex, were associated with potassium levels >6 mmol/L after ACEI/ARB initiation. Thus, the OR was 7-fold increased for baseline potassium >5 mmol/L, two-fold increased for CKD stage 3a, 5-fold increased for stage 3b, and 11-fold increased for stage 4 (Table 5). Among cardiovascular comorbidities, heart failure was associated with the strongest OR of a potassium level >6 mmol/L (2.22, 95% CI: 1.38-3.58).

Monitoring high-risk patients

The odds of having a follow-up test within 2 weeks following drug initiation were higher for persons aged 70 years or above compared with ≤ 50 years (1.18, 95% CI: 1.13-1.23 for 70-79 years and 1.17, 95% CI: 1.11-1.23 for 80+ years). It was also increased for patients with CKD stage (1.41, 95% CI: 1.20-1.66), heart failure (1.16, 95% CI: 1.08-1.23), and peripheral arterial disease (1.11, 95% CI: 1.02-1.20). However, there was no substantially increased OR (>10%) associated with female sex (1.07, 95% CI: 1.04-1.09), prior history of myocardial infarction (0.77, 95% CI: 0.72-0.82), hypertension (1.05, 95% CI: 1.00-1.11), or baseline potassium >5 mmol/L (1.04, 95% CI: 0.99-1.09). When we excluded patients with a recent hospital admission, the reduced OR for myocardial infarction was no longer observed (0.93, 95% CI: 0.80-1.08) (eTable 5). Finally, the results remained consistent when we omitted adjustment for baseline potassium (data not shown) and when we adjusted additionally for ethnicity (eTable 5).

DISCUSSION

Only one tenth of patients initiating ACEI/ARBs in UK primary care appear to receive the guideline-recommended creatinine monitoring. One in 15 patients commenced ACEI/ARBs despite baseline potassium above the recommended level, which was also shown to be a strong predictor for severe post-initiation hyperkalaemia. Among monitored patients, a creatinine increase $\geq 30\%$ or a potassium level >6 mmol/L occurred in almost 1.5% of patients, and most did not discontinue therapy despite guideline recommendations to stop. Although patients with prior myocardial infarction, hypertension, or a high baseline potassium level were at higher risk of sudden decline in kidney function after ACEI/ARB initiation, there was no evidence that these patient groups were monitored more frequently while initiating the drugs.

Strengths and limitations

Several issues should be considered when interpreting our study results. Its large sample size increased precision. Use of the CPRD ensured that the study was population-based and not restricted to specific demographic, hospital, or insurance groups.

We did not have access to blood tests performed in hospital systems, which may have been reported to GPs, but not recorded in CPRD. However, restricting the analysis to patients with no recent hospital admissions had little effect on our findings. Although some patients may also have been seen in outpatient specialty clinics, it is common practice for specialists to ask GPs to initiate new drugs such as ACEI/ARBs, with local biochemical monitoring, limiting misclassification.

Consistent with findings from other studies,²⁰ we found that approximately 50% of all ACEI/ARB initiators were monitored both before and after treatment start. If GPs are retesting renal function in patients at higher risk of adverse outcomes, we may have overestimated the proportion of patients with high potassium levels or creatinine increases compared with the untested lower-risk general population.

GP system software is used for issuing prescriptions, ensuring the accuracy of prescription data. However, it cannot be inferred that all patients actually redeemed their prescription at the pharmacy and start medication on the same day that it was prescribed.^{18 21} Similarly, the estimated coverage of prescriptions may not be completely accurate due to such factors as stockpiling and irregular use. We also do not know whether GPs contacted patients with elevated laboratory results to advise them to stop taking the medication prior to the end of their prescriptions. However, 80% of patients who developed creatinine increase $\geq 30\%$ after ACEI/ARB initiation were still issued a subsequent ACEI/ARB prescriptions.

We aimed to detect discontinuation related closely in time to first follow-up monitoring and hence likely resulting from an elevated creatinine or potassium result. We therefore defined continuation as ACEI/ARB use beyond 30 days (the median prescription duration) after the monitoring date. Extending the definition of continuous use beyond 90 days reduced the risk of misclassifying patients as continuing treatment when they had in fact stopped. However, extending the duration also increased the risk of identifying discontinuation due to other reasons than creatinine/potassium increase, *e.g.*, death or cough. Diagnoses recorded in the CPRD generally have been found to have adequate validity for research purposes,^{22 23} particularly in the domains assessed by the QOF.^{24 25}

In the logistic regression analysis to estimate factors associated with creatinine increase $\geq 30\%$, we excluded patients without pre and post measurements (complete case analysis). If the recording of creatinine levels was not missing completely at random, the associations between patient characteristics and creatinine increase may have been underestimated.²⁶ While this assumption could not be tested directly, examination of baseline characteristics revealed no major differences in age, sex, socioeconomic status, and lifestyle between patients with and without pre- and post-monitoring. Furthermore, the results were consistent for each individual patient group examined. Patients with no testing before or after treatment

initiation (including those with potentially haemolysed samples) only accounted for 10% of all ACEI/ARB initiators.

Comparison with other studies

To our knowledge, this is the largest study conducted to date on adherence to monitoring and discontinuation guidelines after ACEI/ARB initiation. Only one previous study²⁰ examined monitoring according to guideline-recommended intervals (<14 days). All others have used longer intervals (*e.g.*, 30 days²⁷ or 6 months^{28 29}), which make interpretations and implications for clinical practice less clear. Poor adherence to monitoring guidelines after ACEI/ARB initiation is not restricted to the UK,^{20 29 30} but has also been reported in the US³¹⁻³³, Canada,³⁴ and the Netherlands.^{27 35} Owing to our sample size, we were able to show that the lack of monitoring occurred in all patient groups with an indication for ACEI/ARB therapy.

A recent Dutch study, including 3,353 patients initiating ACEI/ARBs between 2005-2011, found that 19% had creatinine measured within 30 days and 66% within one year.²⁷ Creatinine increases above 30% occurred in 1.6% of patients, and among these 70% did not discontinue treatment.²⁷ A Scottish study of 4,056 patients with type 2 diabetes, prescribed an ACEI/ARB between 2005-2009, found that 19% had both a baseline (within 90 days) and follow-up measurement (within 2 weeks) of initiation. Within this cohort, 1.7% had both a creatinine increase of $\geq 30\%$ and potassium level ≥ 5.6 mmol/L.

The magnitude of the risk of severe renal impairment, as measured by creatinine increase in these observational studies, was consistent with our findings, but substantially higher than reported in clinical trials (*e.g.*, 0.2% in the ONTARGET trial).³⁶ It is not clear from the literature how often harm occurs around the time of initiation, when the risk of nephrotoxicity is thought to be greatest.⁸ If physicians are to understand why follow-up monitoring within 2 weeks of treatment start matters, the short-term risks need to be clarified.

Until now, most studies have reported only on cumulative risk over entire courses of treatment, such as the 1.1% two-year risk for potassium of >6 mmol/L in the SOLVD trials of heart failure patients.³⁷ In contrast to clinical trial reviews, reporting a 0.2% (3/1818) risk of potassium >6 mmol/L, we found a 0.4% risk of hyperkalaemia already at time of first retesting after ACEI initiation.

Extending the previous literature, our results support that advanced age, advanced CKD, and heart failure, but not sex, increase the likelihood of being monitored.^{20 27 31} Consistent with some,^{27 31} but not all, previous studies,²⁹ we found no association for diabetes. However, these previous studies reporting an association for diabetes focused on monitoring within broader intervals (*e.g.*, 6 months),²⁹ where diabetes patients, irrespective of ACEI/ARB initiation, were likely to receive blood testing owing to the diabetes QOF programme.

Determinants of increases in creatinine levels after ACEI/ARB initiation are less well understood than for hyperkalemia, but increasing age is a consistently reported factor.²⁰ Advanced CKD and a range of cardiovascular comorbidities (mostly associated with atherosclerosis) were also important determinants in our patient cohort. Consistent with previous studies, we found that the risk of hyperkalemia risk associated with CKD (likely due to impaired ability of the cortical collecting tubule to secrete potassium), heart failure (likely due to decreased delivery of sodium to the distal nephron), and high pre-treatment potassium levels.^{6 8 20 38} We did not observe an association with diabetes or increasing age, as could have been expected due to diabetic nephropathy or age-dependent hyporeninemic hypoaldosteronism.⁶

Clinical relevance

Several possible explanations exist for the divergence between the clinical guideline recommendations and the observed monitoring and response patterns in clinical practice. The

first is *clinician nonadherence* to ordering tests. This may be due to inconsistent recommendations for timing and frequency of monitoring over time,⁶ consensus-based (rather than evidence-based) monitoring guidelines, and a lack of guidelines tailored to particular high-risk patients, such as those with CKD and heart failure. Although we found that follow-up monitoring correlated well with the risk of renal impairment after ACEI/ARB initiation for most patient groups, it was not observed for patients with myocardial infarction or pre-initiation high potassium. The second explanation may be *patient nonadherence* to ordered tests. This is particularly salient in UK primary care where blood samples may be taken in phlebotomy clinics that the patient has to visit rather than the GP practice. Patients may find it burdensome to have blood tests, and GPs have no direct economic incentives to ensure that they are done. A third barrier is *lack of evidence of the clinical importance* of monitoring and its cost-effectiveness. ACEI/ARB-induced renal impairment is rare in clinical trials, even among patients with multiple risk factors for atherosclerotic renal artery stenosis.^{8 39} Trial results may therefore have led to a general perception that the rarity of renal impairment obviates the need for close monitoring. However, as observed in our data, the risks in real world practice may be somewhat higher and non-negligible. In addition, previous research has shown that potassium monitoring in high-risk patients with CKD and diabetes may reduce serious hyperkalaemia-associated adverse events.⁴⁰ Still, the extent to which an initial creatinine increase $\geq 30\%$ translates into adverse long-term outcomes in real-world patients remains to be clarified in future studies.

Generalisability, implications, and conclusions

The majority of patients initiating treatment with ACEI/ARBs experience only minor changes in renal function. However, substantial increases in creatinine levels after ACEI/ARB initiation may not be as rare as previously suggested, reinforcing the need for adherence to clinical guidelines for both pre- and post-initiating monitoring. Moreover, the post-initiation

creatinine increase and potassium levels used in this study are widely recognised cut-off levels, making the results internationally applicable. The comparison with the previous literature also confirms that the lack of systematic monitoring is not exclusive to the UK. Of particular concern was that even when appropriate monitoring was performed, severe renal impairment only rarely led to treatment discontinuation. Individual patient counselling may also be helpful to ensure that those at highest risk are closely monitored. More work is needed to determine the prognostic importance of the changes in renal function that we have observed.

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Transparency declaration: The lead author affirms that the manuscript is an honest, accurate, and transparent account of the study being reported; that no important aspects of the study have been omitted; and that any discrepancies from the study as planned have been explained.

Contributorship

LT conceived the study idea and acquired data permissions. MS, KM, and LT designed the study. MS and KM performed data management and established the cohort. MS, KM, and LT reviewed the literature. The analyses were carried out by MS. All authors participated in the discussion and interpretation of the results. MS organised the writing and wrote the initial drafts. All authors critically revised the manuscript for intellectual content and approved the final version. MS is the guarantor.

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Disclosures: None

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Table 1. Characteristics of patients initiating angiotensin converting-enzyme inhibitors or angiotensin-receptor blockers in the UK primary care during 2004-2014, by monitoring groups

	Serum creatinine monitoring*				Total
	No baseline or follow-up tests	Baseline test only	Follow-up test only	Baseline and follow-up test	
Total number	21,411 (100)	63,359 (100)	33,185 (100)	105,859 (100)	223,814 (100)
Female sex	8,882 (41)	27,722 (44)	14,570 (44)	49,109 (46)	100,283 (45)
Age (years)					
<50 years	5,019 (23)	13,697 (22)	8,732 (26)	19,910 (19)	47,358 (21)
50-59 years	5,485 (26)	15,135 (24)	9,115 (27)	24,866 (23)	54,601 (24)
60-69 years	4,863 (23)	15,586 (25)	7,776 (23)	27,790 (26)	56,015 (25)
70-79 years	3,579 (17)	12,193 (19)	5,066 (15)	22,152 (21)	42,990 (19)
80+ years	2,465 (12)	6,748 (11)	2,496 (8)	11,141 (11)	22,850 (10)
Calendar period					
2004-2008	14,814 (69)	40,667 (64)	19,808 (60)	60,902 (58)	136,191 (61)
2009-2014	6,597 (31)	22,692 (36)	13,377 (40)	44,957 (42)	87,623 (39)
SES quintiles					
1 (low)	5,153 (24)	15,290 (24)	8,533 (26)	25,577 (24)	54,553 (24)
2	4,725 (22)	14,331 (23)	7,887 (24)	24,851 (23)	51,794 (23)
3	4,341 (20)	13,028 (21)	6,890 (21)	22,629 (21)	46,888 (21)
4	4,254 (20)	12,140 (19)	5,931 (18)	19,318 (18)	41,643 (19)
5 (high)	2,925 (14)	8,508 (13)	3,898 (12)	13,359 (13)	28,690 (13)
Missing	13 (0)	62 (0)	46 (0)	125 (0)	246 (0)
Smoking status					
Never	7,860 (37)	22,496 (36)	12,229 (37)	36,895 (35)	79,480 (36)
Ever	13,433 (63)	40,797 (64)	20,915 (63)	68,939 (65)	144,084 (64)
Missing	118 (1)	66 (0)	41 (0)	25 (0)	250 (0)
Alcohol intake					
No use	2,556 (12)	7,819 (12)	3,409 (10)	11,088 (10)	24,872 (11)
Current	15,495 (72)	47,322 (75)	25,656 (77)	82,870 (78)	171,343 (77)
Former	1,328 (6)	4,499 (7)	1,933 (6)	7,490 (7)	15,250 (7)
Missing	2,032 (9)	3,719 (6)	2,187 (7)	4,411 (4)	12,349 (6)
BMI groups					
Underweight	282 (1)	700 (1)	304 (1)	1,008 (1)	2,294 (1)
Healthy weight	5,666 (26)	15,406 (24)	8,089 (24)	24,972 (24)	54,133 (24)
Overweight	7,677 (36)	23,755 (37)	12,484 (38)	40,556 (38)	84,472 (38)
Obesity	6,009 (28)	20,660 (33)	10,527 (32)	35,887 (34)	73,083 (33)
Missing	1,777 (8)	2,838 (4)	1,781 (5)	3,436 (3)	9,832 (4)
CKD (eGFR)†					
Stage ≤2 (≥60)	10,326 (48)	53,773 (85)	19,470 (59)	87,484 (83)	171,053 (76)
Stage 3a (45–59)	1,137 (5)	7,382 (12)	1,766 (5)	13,913 (13)	24,198 (11)
Stage 3b (30–44)	217 (1)	1,885 (3)	265 (1)	3,854 (4)	6,221 (3)
Stage 4 (15–29)	24 (0)	319 (1)	29 (0)	608 (1)	980 (0)
Not measured	9,707 (45)	0 (0)	11,655 (35)	0 (0)	21,362 (10)
CV comorbidities‡					
Heart failure	1,568 (7)	3,270 (5)	1,386 (4)	4,583 (4)	10,807 (5)
Myocardial infarction	3,881 (18)	4,653 (7)	3,203 (10)	4,620 (4)	16,357 (7)
Hypertension	13,023 (61)	44,273 (70)	24,195 (73)	80,946 (76)	162,437 (73)
Peripheral arterial disease	471 (2)	1,590 (3)	523 (2)	2,547 (2)	5,131 (2)
Arrhythmia	2,057 (10)	4,973 (8)	2,000 (6)	7,123 (7)	16,153 (7)
Diabetes mellitus	1,399 (7)	13,586 (21)	1,992 (6)	21,548 (20)	38,525 (17)

Abbreviations: CV, cardiovascular; CKD, chronic kidney disease; y, year
*Monitoring groups based on baseline (within 12 months before) and follow-up (within 2 months after) serum creatinine monitoring.
†Calculated from most recent creatinine measurement within 12 months before first prescription date.
‡Diagnosis ever registered before ACE/ARB initiation in CRPD or HES.

Table 2. Prevalence of baseline and follow-up serum creatinine monitoring among patients initiating angiotensin converting-enzyme inhibitors or angiotensin-receptor blockers, 2004-2014

	S-Creatinine, ≥ 1 test
Total number	n=223,814 (100%)
Baseline testing	
≤12 months before	169,218 (76%)
≤3 months before	115,348 (52%)
≤1 months before	75,476 (34%)
Follow-up testing	
≤2 weeks after	65,090 (29%)
≤1 month after	114,244 (51%)
≤2 months after	139,044 (62%)

Table 3. Prevalence of baseline and follow-up serum creatinine monitoring among patients initiating renin-angiotensin system blockade according to clinical guideline recommendations

	Clinical guidelines					All initiators n=223,814 (100%)	
	NICE Heart Failure	NICE MI	NICE/UKRA Hypertension	NICE CKD	GP Notebook	Wide baseline interval (≤12 month)	Ideal baseline interval (≤1 month)
Baseline testing	x	x	x	x	x	169,218 (76%)	75,476 (34%)
+ Follow-up test ≤2 weeks*	x	n/a	x	x	x	46,486 (21%)	19,679 (9%)
+ Follow-up test ≤3 weeks†						70,792 (32%)	30,451 (14%)

Abbreviations: CKD, chronic kidney disease; GP, General Practice; MI, myocardial infarction; n/a, not specified; UKRA, UK Renal Association

*Follow-up test among those with baseline measurements

† Sensitivity analysis illustrating the importance of 2 vs. 3-week cut-off interval in follow-up test intervals.

Table 4. Proportion of new users of angiotensin converting-enzyme inhibitors or angiotensin-receptor blockers who continue or discontinue treatment according to guideline recommended cut-off levels of serum creatinine and potassium at follow-up testing*

	Continuation†	Discontinuation†	Total
Total number, %	42,942 (93.1)	3,178 (6.9)	46,120 (100)
Serum creatinine increase $\geq 30\%$, n (%)	462 (81.5)	105 (18.5)	567 (100)
Serum potassium >6 mmol/L, n (%)	150 (78.5)	41 (21.5)	191 (100)

*Calculated from the most recent measurements within 1 month before and 2 months after drug initiation.

† A patient was considered continuous users when the end date of the first continuous course of therapy was larger than the date of the first follow-up monitoring + 30 days (to allow for stock piling and irregular use)

Table 5. Association between patient characteristics and serum creatinine increase $\geq 30\%$ and follow-up monitoring within 2 weeks following initiation of renin-angiotensin system blockade

Characteristics	Odds ratio (95% confidence intervals)					
	S-creatinine monitoring within 2 weeks		S-creatinine increase $\geq 30\%^*$		S-potassium increase $\geq 30\%^*$	
	Age- and sex adjusted	Fully adjusted†	Age- and sex adjusted	Fully adjusted†	Age- and sex adjusted	Fully adjusted†
Female sex	1.07 (1.04-1.10)	1.07 (1.04-1.09)	1.39 (1.26-1.53)	1.63 (1.47-1.80)	0.87 (0.66-1.16)	0.94 (0.70-1.26)
Age (years)						
< 50 years	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
50-59 years	0.98 (0.94-1.01)	0.98 (0.95-1.02)	0.88 (0.74-1.05)	0.86 (0.72-1.03)	1.29 (0.79-2.11)	1.10 (0.67-1.81)
60-69 years	1.05 (1.02-1.09)	1.05 (1.01-1.09)	1.03 (0.88-1.21)	1.00 (0.85-1.19)	1.35 (0.84-2.17)	0.97 (0.60-1.58)
70-79 years	1.18 (1.14-1.23)	1.18 (1.13-1.23)	1.49 (1.27-1.74)	1.36 (1.15-1.61)	1.65 (1.02-2.66)	0.74 (0.43-1.26)
80+ years	1.20 (1.14-1.25)	1.17 (1.11-1.23)	2.72 (2.32-3.20)	2.02 (1.68-2.44)	2.75 (1.67-4.53)	0.73 (0.41-1.32)
CKD stage						
No CKD (≥ 60)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Stage 3a (45-59)	1.00 (0.96-1.04)	1.00 (0.96-1.04)	0.62 (0.53-0.73)	0.60 (0.51-0.70)	2.48 (1.66-3.71)	2.06 (1.36-3.11)
Stage 3b (30-44)	0.99 (0.93-1.06)	1.01 (0.94-1.08)	1.01 (0.82-1.24)	0.88 (0.71-1.09)	7.51 (4.75-11.9)	5.10 (3.16-8.22)
Stage 4 (15-29)	1.42 (1.21-1.67)	1.41 (1.20-1.66)	2.16 (1.52-3.05)	1.72 (1.18-2.51)	24.0 (13.5-42.6)	11.4 (6.07-21.4)
Comorbidities*						
Heart failure	1.15 (1.09-1.23)	1.16 (1.08-1.23)	4.00 (3.49-4.58)	2.93 (2.51-3.42)	2.90 (1.90-4.42)	2.22 (1.38-3.58)
MI	0.80 (0.75-0.85)	0.77 (0.72-0.82)	2.33 (1.98-2.74)	1.57 (1.32-1.87)	2.12 (1.33-3.39)	1.35 (0.80-2.25)
Hypertension	1.00 (0.97-1.02)	1.05 (1.00-1.11)	0.62 (0.56-0.68)	1.58 (1.36-1.84)	0.60 (0.45-0.80)	1.02 (0.63-1.65)
PAD	1.09 (1.01-1.18)	1.11 (1.02-1.20)	2.10 (1.70-2.60)	1.87 (1.50-2.33)	2.14 (1.18-3.86)	1.53 (0.82-2.88)
Arrhythmia	1.09 (1.03-1.14)	0.98 (0.95-1.01)	2.37 (2.07-2.71)	0.77 (0.69-0.86)	1.41 (0.90-2.21)	0.77 (0.56-1.05)
Diabetes mellitus	0.93 (0.90-0.96)	0.93 (0.90-0.96)	1.09 (0.97-1.22)	1.04 (0.92-1.18)	0.97 (0.69-1.36)	0.90 (0.63-1.29)
Baseline K>5 mmol/L	1.04 (1.00-1.10)	1.04 (0.99-1.09)	1.04 (0.86-1.25)	0.97 (0.80-1.17)	8.22 (6.14-11.0)	6.68 (4.94-9.02)

Abbreviations: CKD, chronic kidney disease; MI, myocardial infarction; PAD, peripheral arterial disease
* The increase was based on the difference between the most recent baseline measurements within 12 months before and first follow-up measurement within 2 months after drug initiation. All analyses were restricted to those with both baseline and follow-up measurements (n=105,859).
† Adjusted for sex, age, CKD, heart failure, MI, hypertension, PAD, arrhythmia, diabetes, and calendar period of prescription start

ONLINE SUPPLEMENTAL MATERIAL

eTable 1. Prevalence of baseline and follow-up serum creatinine monitoring among patients initiating renin-angiotensin system blockade according to clinical guideline recommendations, overall and in most recent calendar period (extended version of Table 3)

	Clinical guidelines						All initiators		Continuing users*	
	NICE Heart Failure	NICE CKD	NICE Hypertension	NICE MI	UK Renal Association	GP Notebook	Generous baseline interpretation (≤12 month)	Strict baseline interpretation (≤1 month)	Generous baseline interpretation (≤12 month)	Strict baseline interpretation (≤1 month)
2004-2014							n=223,814 (100%)		n=173,244 (100%)	
Baseline testing	x	x	x	x	x	x	169,218 (76%)	75,476 (34%)	132,156 (76%)	58,668 (34%)
Follow-up test (≤2 weeks)†	x	x	n/a	x	x	x	46,486 (21%)	19,679 (9%)	43,420 (25%)	18,457 (11%)
+ At least within one year after				x			36,424 (16%)	15,642 (7%)	34,336 (20%)	14,800 (9%)
+ 1 month thereafter‡						x	11,866 (5%)	5,643 (3%)	11,298 (7%)	5,388 (3%)
+ 1 and 3 months thereafter ‡						x	5,787 (3%)	2,933 (1%)	5,516 (3%)	2,791 (2%)
+ 1, 3, and 6 months thereafter ‡						x	3,390 (2%)	1,807 (1%)	3,201 (2%)	1,703 (1%)
+ 1, 3, 6, and 12 months thereafter‡						x	2,618 (1%)	1,439 (1%)	2,464 (1%)	1,353 (1%)
Follow-up test (≤3 weeks)§							70,792 (32%)	30,451 (14%)	66,340 (38%)	28,644 (17%)
2009-2014							n=87,623 (100%)		n=68,042 (100%)	
Baseline testing	x	x	x	x	x	x	67,649 (77%)	30,576 (35%)	52,760 (78%)	23,716 (35%)
Follow-up test (≤2 weeks)†	x	x	n/a	x	x	x	20,172 (23%)	8,707 (10%)	18,862 (28%)	8,169 (12%)
+ At least within one year after				x			15,416 (18%)	6,726 (8%)	14,557 (21%)	6,377 (9%)
+ 1 month thereafter‡						x	5,373 (6%)	2,607 (3%)	5,123 (8%)	2,500 (4%)
+ 1 and 3 months thereafter ‡						x	2,614 (3%)	1,365 (2%)	2,492 (4%)	1,302 (2%)
+ 1, 3, and 6 months thereafter ‡						x	1,525 (2%)	841 (1%)	1,437 (2%)	792 (1%)
+ 1, 3, 6, and 12 months thereafter‡						x	1,152 (1%)	650 (1%)	1,079 (2%)	606 (1%)
Follow-up test (≤3 weeks)§							31,007 (35%)	13,511 (15%)	29,081 (43%)	12,726 (19%)

Abbreviations: CKD, chronic kidney disease; GP, General Practice; MI, myocardial infarction; n/a, not specified

* A patient was considered continuous users when the end date of the first continuous course of therapy was larger than the date of the first follow-up monitoring + 30 days (to allow for stock piling and irregular use)

† Follow-up test among those with baseline measurements

‡ Within additional 1, 3, 6, and 12 months after correspond to within 1.5, 3.5, 6.5, and 12.5 months after ACE/ARB initiation, respectively.

§ Sensitivity analysis illustrating the importance of 2 vs. 3 week cut-off interval in follow-up test intervals.

eTable 2. Prevalence of baseline and follow-up serum creatinine monitoring among patients initiating angiotensin converting-enzyme inhibitors or angiotensin-receptor blockers, overall and most recent calendar period (extended version of Table 1)

	S-Creatinine, ≥1 test, n (%)	
	Total cohort (n=223,814)	2009-2014 cohort (n=87,623)
Baseline testing		
≤12 months before	169,218 (75.6)	67,649 (77.2)
≤3 months before	115,348 (51.5)	46,925 (53.6)
≤1 months before	75,476 (33.7)	30,576 (34.9)
Follow-up testing		
≤2 weeks after	65,090 (29.1)	27,933 (31.9)
≤1 month after	114,244 (51.0)	48,877 (55.8)
≤2 months after	139,044 (62.1)	58,334 (66.6)

eTable 3. Prevalence of baseline and follow-up serum creatinine monitoring among patients initiating renin-angiotensin system blockade according to clinical guideline recommendations

	All initiators n=223,814 (100%)	
	Generous baseline interpretation (≤12 month)	Strict baseline interpretation (≤1 month)
Overall (n=223,814)		
Baseline testing	169,218 (76%)	75,476 (34%)
+ Follow-up test ≤2 weeks*	46,486 (21%)	19,679 (9%)
+ Follow-up test ≤3 weeks†	70,792 (32%)	30,451 (14%)
Overall, no recent hospitalization (n=197,291)		
Baseline testing	153,353 (78%)	68,883 (35%)
+ Follow-up test ≤2 weeks*	42,967 (22%)	18,222 (9%)
+ Follow-up test ≤3 weeks†	65,881 (33%)	28,375 (14%)
Heart failure patients (n=10,807)		
Baseline testing	7,853 (73%)	3,277 (30%)
+ Follow-up test ≤2 weeks*	2,229 (21%)	951 (9%)
+ Follow-up test ≤3 weeks†	3,085 (29%)	1,340 (12%)
Myocardial infarction patients (n=16,357)		
Baseline testing	9,273 (57%)	3,766 (23%)
+ Follow-up test ≤2 weeks*	1,816 (11%)	726 (4%)
+ Follow-up test ≤3 weeks†	2,645 (16%)	1,078 (7%)
Hypertension patients (n=162,437)		
Baseline testing	125,219 (77%)	54,174 (33%)
+ Follow-up test ≤2 weeks*	35,502 (22%)	14,559 (9%)
+ Follow-up test ≤3 weeks†	54,585 (34%)	22,710 (14%)
CKD patients (n=31,399)		
Baseline testing	27,961 (89%)	13,913 (44%)
+ Follow-up test ≤2 weeks*	8,457 (27%)	4,198 (13%)
+ Follow-up test ≤3 weeks†	12,482 (40%)	6,324 (20%)
PAD patients (n=5,131)		
Baseline testing	4,137 (81%)	1,671 (33%)
+ Follow-up test ≤2 weeks*	1,198 (23%)	458 (9%)
+ Follow-up test ≤3 weeks†	1,751 (34%)	698 (14%)
Diabetes patients (n=38,525)		
Baseline testing	35,134 (91%)	15,907 (41%)
+ Follow-up test ≤2 weeks*	9,161 (24%)	3,794 (10%)
+ Follow-up test ≤3 weeks†	14,093 (37%)	5,994 (16%)

Abbreviations: CKD, chronic kidney disease; GP, General Practice; MI, myocardial infarction; n/a, not specified; UKRA, UK Renal Association

*Follow-up test among those with baseline measurements

† Sensitivity analysis illustrating the importance of 2 vs. 3-week cut-off interval in follow-up test intervals.

eTable 4. Proportion of new users of ACEI/ARB who continue or discontinue treatment according to guideline recommended cut-off levels of serum creatinine and potassium at follow-up testing

	Wide monitoring interval (12 month before to the first monitoring within 2 month after)			Narrow monitoring interval (1 month before to the first monitoring within 2 month after)		
	Continuation*	Discontinuation*	Total	Continuation*	Discontinuation*	Total
+ 30 day window						
Total number, %	113,210 (92.5)	9,170 (7.5)	122,380 (100)	48,772 (93.0)	3,676 (7.0)	52,448 (100)
S-creatinine increase $\geq 30\%$, n (%)	1,679 (80.7)	401 (19.3)	2,080 (100)	541 (81.5)	123 (18.5)	664 (100)
S-potassium >6 mmol/L, n (%)	472 (81.9)	104 (18.1)	576 (100)	204 (80.6)	49 (19.4)	253 (100)
+ 90 day window						
Total number, %	86,620 (81.8)	19,239 (18.2)	105,859 (100)	37,920 (82.2)	8,200 (17.8)	46,120 (100)
S-creatinine increase $\geq 30\%$, n (%)	1,111 (64.7)	605 (35.3)	1,716 (100)	372 (65.6)	195 (34.4)	567 (100)
S-potassium >6 mmol/L, n (%)	261 (61.0)	167 (39.0)	428 (100)	110 (57.6)	81 (42.4)	191 (100)

Calculated from the measurements closest to the first prescription date in the corresponding intervals, i.e., the most recent baseline measurements and the first follow-up measurement.

* A patient was considered continuous users when the end date of the first continuous course of therapy was larger than the date of the first follow-up monitoring + 30 days or + 90days (to allow for stock piling and irregular use)

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eTable 5. Association between patient characteristics and serum creatinine increase $\geq 30\%$ and follow-up monitoring within 2 weeks following initiation of renin-angiotensin system blockade

Characteristics	Adjusted odds ratio (95% confidence intervals)					
	Additional adjustment for ethnicity†			Excluding patient with hospitalisation within 30 days†		
	S-creatinine monitoring within 2 weeks	S-creatinine increase $\geq 30\%*$	S-potassium >6 mmol/L*	S-creatinine monitoring within 2 weeks	S-creatinine increase $\geq 30\%*$	S-potassium >6 mmol/L*
Female sex	1.04 (1.01-1.08)	1.91 (1.64-2.23)	1.09 (0.70-1.71)	1.06 (1.04-1.09)	1.66 (1.48-1.86)	0.94 (0.69-1.27)
Age (years)						
< 50 years	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
50-59 years	0.97 (0.92-1.02)	0.91 (0.71-1.16)	1.65 (0.77-3.52)	0.99 (0.95-1.03)	0.86 (0.71-1.05)	1.17 (0.70-1.98)
60-69 years	1.04 (0.99-1.10)	0.86 (0.68-1.10)	1.23 (0.57-2.65)	1.07 (1.02-1.11)	1.05 (0.87-1.26)	0.98 (0.58-1.64)
70-79 years	1.22 (1.15-1.30)	1.22 (0.96-1.55)	0.83 (0.36-1.95)	1.19 (1.14-1.24)	1.40 (1.16-1.68)	0.72 (0.41-1.27)
80+ years	1.15 (1.06-1.24)	1.61 (1.21-2.14)	0.58 (0.21-1.58)	1.17 (1.11-1.24)	2.16 (1.76-2.66)	0.71 (0.38-1.33)
CKD stage						
No CKD (≥ 60)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Stage 3a (45–59)	1.00 (0.94-1.06)	0.51 (0.39-0.66)	1.82 (0.93-3.55)	1.00 (0.96-1.04)	0.58 (0.49-0.70)	2.40 (1.57-3.68)
Stage 3b (30–44)	1.06 (0.95-1.17)	0.99 (0.71-1.39)	5.66 (2.70-11.9)	0.98 (0.92-1.06)	0.92 (0.73-1.17)	5.15 (3.09-8.58)
Stage 4 (15–29)	1.51 (1.17-1.94)	1.65 (0.87-3.12)	15.0 (5.84-38.8)	1.42 (1.20-1.70)	1.86 (1.23-2.81)	12.2 (6.29-23.7)
Comorbidities*						
Heart failure	1.10 (0.99-1.21)	3.07 (2.41-3.91)	2.16 (0.96-4.85)	1.15 (1.06-1.24)	2.57 (2.11-3.13)	2.68 (1.59-4.51)
MI	0.78 (0.72-0.86)	1.76 (1.35-2.30)	1.10 (0.46-2.63)	0.89 (0.82-0.97)	1.59 (1.23-2.06)	1.89 (1.05-3.41)
Hypertension	0.97 (0.90-1.05)	1.51 (1.19-1.92)	1.01 (0.44-2.30)	1.04 (0.99-1.11)	1.40 (1.16-1.68)	0.81 (0.46-1.42)
PAD	1.09 (0.96-1.23)	1.63 (1.13-2.35)	1.52 (0.54-4.29)	1.16 (1.06-1.26)	1.74 (1.34-2.27)	1.75 (0.92-3.32)
Arrhythmia	0.96 (0.92-1.01)	0.84 (0.71-0.99)	0.72 (0.44-1.17)	0.98 (0.94-1.01)	0.80 (0.71-0.91)	0.74 (0.53-1.03)
Diabetes mellitus	0.91 (0.87-0.96)	1.15 (0.96-1.37)	0.58 (0.31-1.09)	0.93 (0.90-0.96)	1.06 (0.93-1.22)	0.93 (0.64-1.35)
Baseline K >6 mmol/L	1.05 (0.98-1.13)	0.86 (0.63-1.15)	7.98 (5.06-12.6)	1.03 (0.98-1.08)	1.01 (0.82-1.24)	6.44 (4.69-8.84)

Abbreviations: CKD, chronic kidney disease; MI, myocardial infarction; PAD, peripheral arterial disease
* The increase was based on the difference between the most recent baseline measurements within 12 months before and first follow-up measurement within 2 months after drug initiation. All analyses were restricted to those with both baseline and follow-up measurements (n=105,859).
† The main model was adjusted for sex, age, CKD, heart failure, MI, hypertension, PAD, arrhythmia, diabetes, and calendar period of prescription start

The RECORD statement – checklist of items, extended from the STROBE statement, that should be reported in observational studies using routinely collected health data.

	Item No.	STROBE items	Location in manuscript where items are reported	RECORD items	Location in manuscript where items are reported
Title and abstract					
	1	(a) Indicate the study's design with a commonly used term in the title or the abstract (b) Provide in the abstract an informative and balanced summary of what was done and what was found	a) 3, 6 b) 3	RECORD 1.1: The type of data used should be specified in the title or abstract. When possible, the name of the databases used should be included. RECORD 1.2: If applicable, the geographic region and timeframe within which the study took place should be reported in the title or abstract. RECORD 1.3: If linkage between databases was conducted for the study, this should be clearly stated in the title or abstract.	1.1 3, 5 1.2 3 1.3 3
Introduction					
Background rationale	2	Explain the scientific background and rationale for the investigation being reported	5		
Objectives	3	State specific objectives, including any prespecified hypotheses	5		
Methods					
Study Design	4	Present key elements of study design early in the paper	5		
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	5-6		
Participants	6	(a) Cohort study - Give the eligibility criteria, and the	a) 5-6	RECORD 6.1: The methods of study population selection (such as codes or	6.1) 5-6 6.2) 5-6, 13-14

		sources and methods of selection of participants. Describe methods of follow-up <i>Case-control study</i> - Give the eligibility criteria, and the sources and methods of case ascertainment and control selection. Give the rationale for the choice of cases and controls <i>Cross-sectional study</i> - Give the eligibility criteria, and the sources and methods of selection of participants <i>(b) Cohort study</i> - For matched studies, give matching criteria and number of exposed and unexposed <i>Case-control study</i> - For matched studies, give matching criteria and the number of controls per case		algorithms used to identify subjects) should be listed in detail. If this is not possible, an explanation should be provided. RECORD 6.2: Any validation studies of the codes or algorithms used to select the population should be referenced. If validation was conducted for this study and not published elsewhere, detailed methods and results should be provided. RECORD 6.3: If the study involved linkage of databases, consider use of a flow diagram or other graphical display to demonstrate the data linkage process, including the number of individuals with linked data at each stage.	6.3) Not needed
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable.	5-7	RECORD 7.1: A complete list of codes and algorithms used to classify exposures, outcomes, confounders, and effect modifiers should be provided. If these cannot be reported, an explanation should be provided.	7
Data sources/ measurement	8	For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe comparability of assessment methods if there is more than one group	5-7		
Bias	9	Describe any efforts to address potential sources of bias	8-9		
Study size	10	Explain how the study size was	6-7		

		arrived at			
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen, and why	5-9		
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding (b) Describe any methods used to examine subgroups and interactions (c) Explain how missing data were addressed (d) <i>Cohort study</i> - If applicable, explain how loss to follow-up was addressed <i>Case-control study</i> - If applicable, explain how matching of cases and controls was addressed <i>Cross-sectional study</i> - If applicable, describe analytical methods taking account of sampling strategy (e) Describe any sensitivity analyses	8-9		
Data access and cleaning methods		..		RECORD 12.1: Authors should describe the extent to which the investigators had access to the database population used to create the study population. RECORD 12.2: Authors should provide information on the data cleaning methods used in the study.	12.1) 5-6 12.2) 5-6
Linkage		..		RECORD 12.3: State whether the study included person-level, institutional-	12.3) 5-6

				level, or other data linkage across two or more databases. The methods of linkage and methods of linkage quality evaluation should be provided.	
Results					
Participants	13	(a) Report the numbers of individuals at each stage of the study (e.g., numbers potentially eligible, examined for eligibility, confirmed eligible, included in the study, completing follow-up, and analysed) (b) Give reasons for non-participation at each stage. (c) Consider use of a flow diagram	5-6, 9-12	RECORD 13.1: Describe in detail the selection of the persons included in the study (i.e., study population selection) including filtering based on data quality, data availability and linkage. The selection of included persons can be described in the text and/or by means of the study flow diagram.	13.1) 5-6, 9-12
Descriptive data	14	(a) Give characteristics of study participants (e.g., demographic, clinical, social) and information on exposures and potential confounders (b) Indicate the number of participants with missing data for each variable of interest (c) Cohort study - summarise follow-up time (e.g., average and total amount)	9-12, Table 1		
Outcome data	15	Cohort study - Report numbers of outcome events or summary measures over time Case-control study - Report numbers in each exposure category, or summary measures of exposure Cross-sectional study - Report numbers of outcome events or summary measures	9-12		
Main results	16	(a) Give unadjusted estimates	9-12		

		and, if applicable, confounder-adjusted estimates and their precision (e.g., 95% confidence interval). Make clear which confounders were adjusted for and why they were included (b) Report category boundaries when continuous variables were categorized (c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period			
Other analyses	17	Report other analyses done—e.g., analyses of subgroups and interactions, and sensitivity analyses	9-12		
Discussion					
Key results	18	Summarise key results with reference to study objectives	12		
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias	12-14	RECORD 19.1: Discuss the implications of using data that were not created or collected to answer the specific research question(s). Include discussion of misclassification bias, unmeasured confounding, missing data, and changing eligibility over time, as they pertain to the study being reported.	12-14
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence	17		
Generalisability	21	Discuss the generalisability (external validity) of the study results	17		

Other Information					
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	2		
Accessibility of protocol, raw data, and programming code		..	7, protocol made available for reviewers	RECORD 22.1: Authors should provide information on how to access any supplemental information such as the study protocol, raw data, or programming code.	7, protocol made available for reviewers

*Reference: Benchimol EI, Smeeth L, Guttman A, Harron K, Moher D, Petersen I, Sørensen HT, von Elm E, Langan SM, the RECORD Working Committee. The REporting of studies Conducted using Observational Routinely-collected health Data (RECORD) Statement. *PLoS Medicine* 2015; in press.

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**Adherence to guidelines for creatinine and potassium monitoring and
discontinuation following renin-angiotensin system blockade:
a UK general practice-based cohort study**

Running title: Creatinine and potassium monitoring after ACEI/ARB initiation

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Competing interests: All authors have completed the Unified Competing Interest form at www.icmje.org/coi_disclosure.pdf (available on request from the corresponding author) and declare that (1) no authors have support from any company for the submitted work. The Department of Clinical Epidemiology is involved, however, in studies with funding from various companies as research grants to (and administered by) Aarhus University. None of these studies have any relation to the present study; (2) no authors have relationships with companies that might have an interest in the submitted work in the previous 3 years; (3) their spouses, partners, or children have no financial relationships that may be relevant to the submitted work; and (4) no authors have non-financial interests that may be relevant to the submitted work.

ABSTRACT

Objectives To examine adherence to serum creatinine and potassium monitoring and discontinuation guidelines following initiation of treatment with angiotensin converting-enzyme inhibitors (ACEI) or angiotensin-receptor blockers (ARB); and whether high-risk patients are monitored.

Design General practice-based cohort study using electronic health records from the UK Clinical Practice Research Datalink and Hospital Episode Statistics.

Setting UK primary care, 2004–2014.

Subjects 223,814 new ACEI/ARB users.

Main outcome measures Proportion of patients with renal function monitoring before and after ACEI/ARB initiation; creatinine increase $\geq 30\%$ or potassium levels >6 mmol/L at first follow-up monitoring; and treatment discontinuation after such changes. Using logistic regression models, we also examined patient characteristics associated with these biochemical changes, and with follow-up monitoring within the guideline-recommendation of two weeks after treatment initiation.

Results Ten percent of patients had neither baseline nor follow-up monitoring of creatinine within 12 months before and 2 months after initiation of an ACEI/ARB, 28% had monitoring only at baseline, 15% only at follow-up, and 47% both at baseline and follow-up. The median period between the most recent baseline monitoring and drug initiation was 40 days (interquartile range: 12-125 days). 34% of patients had baseline creatinine monitoring within one month before initiating therapy, but less than 10% also had the guideline-recommended follow-up test recorded within two weeks. Among patients experiencing a creatinine increase $\geq 30\%$ (n=567, 1.2%) or potassium level >6 mmol/L (n=191, 0.4%), 80% continued treatment. Although patients with prior myocardial infarction, hypertension, or baseline potassium >5 mmol/L were at high risk of $\geq 30\%$ increase in creatinine after ACEI/ARB initiation, there was no evidence that they were more frequently monitored.

Conclusions Only one tenth of patients initiating ACEI/ARB therapy receive the guideline-recommended creatinine monitoring. Moreover, the vast majority of the patients fulfilling post-initiation discontinuation criteria for creatinine and potassium increases continue on treatment.

Article Summary:

- This is the largest monitoring study to date, examining both adherence to creatinine and potassium monitoring and discontinuation guidelines following initiation of angiotensin converting-enzyme inhibitors or angiotensin-receptor blockers in UK primary care, and whether patients are monitored in accordance with their individual risk profile
- Use of the UK Clinical Practice Research Datalink and Hospital Episode Statistics ensured that the study was population-based and not restricted to specific demographic, hospital, or insurance groups.
- Blood tests performed in hospital systems were not recorded in the Clinical Practice Research Datalink, but the results were consistent for patients with no recent hospital admissions
- If the recording of creatinine levels was not missing completely at random, the associations between patient characteristics and creatinine increase may have been underestimated

INTRODUCTION

Renin angiotensin system blockade using angiotensin-converting enzyme inhibitors (ACEI) and angiotensin-receptor blockers (ARB) is a mainstay in treatment of hypertension,¹ heart failure,² diabetic microalbuminuria or proteinuric renal diseases,³ and after myocardial infarction.⁴ However, some patients experience a sudden decline in kidney function when initiating these drugs, presumably due to antagonism of the angiotensin II-mediated efferent arteriolar constriction or impaired kidney excretion of potassium.^{5 6}

The potential impact on kidney function should be evaluated by comparing pre- and post-initiation levels of serum creatinine and potassium.⁷ Discontinuation is recommended if the rise in creatinine exceeds 30% above baseline or if hyperkalaemia develops.⁸ It is unclear whether these recommendations are routinely followed in clinical practice.⁹

A few studies have compared baseline and follow-up monitoring results,⁹ but large studies using contemporary data with reference to current guidelines are lacking, and it is unknown whether patients' individual risk of renal impairment influence their likelihood of being monitored.⁹ We therefore examined adherence to creatinine and potassium monitoring and treatment discontinuation guidelines following ACEI/ARB initiation in UK primary care, and whether patients are monitored in accordance with their individual risk profile.

METHODS

Data sources

We used the UK's Clinical Practice Research Datalink (CPRD) linked to hospital record data from the Hospital Episode Statistics (HES) database. The CPRD database contains primary care electronic health record data from 7% of the UK population (~15 million patient lives, with ~8 million currently under follow-up).¹⁰ Patients included in the CPRD are largely representative of the UK population in terms of age, sex and ethnicity.^{10 11} Information recorded in the database includes demographics such as sex and year of birth, the location of the general practice, medical diagnoses (based on 'Read' codes), drug prescriptions, and a

range of routine laboratory test results. HES records cover all hospital admissions for patients covered by the NHS who receive treatment either from English NHS trusts or independent providers.^{10 11} Fifty-eight percent of general practices included in the CPRD have agreed to HES linkage.¹⁰ We obtained linked data on socioeconomic status (index of multiple deprivation) based on area of residence.

Monitoring guidelines

Consistent with other international guidelines, the National Institute for Health and Care Excellence (NICE) recommend baseline testing of creatinine when initiating ACEI/ARB therapy in patients with hypertension,¹ heart failure,² myocardial infarction,⁴ or chronic kidney disease (CKD).³ The time interval for baseline testing is not further specified.¹⁻⁴ Among patients with heart failure, myocardial infarction, and chronic kidney disease, NICE recommends follow-up monitoring within 2 weeks of treatment initiation,²⁻⁴ and for myocardial infarction patients at least annually thereafter.⁴ A baseline assessment and follow-up test within 2 weeks are also recommended by the UK Renal Association,¹² as well as the frequently used online web resource General Practice (GP) Notebook.¹³ GP Notebook additionally recommends monitoring 1, 3, 6, and 12 months after the first follow-up test.¹³ NICE recommends not to initiate ACEI/ARBs in patients with a baseline potassium level >5 mmol/L and to discontinue therapy if potassium rises above 6 mmol/L.

ACEI/ARB initiators

We identified a cohort of all HES linked CPRD patients aged ≥ 18 years, who initiated ACEI/ARB treatment between January 1, 2004 and March 31, 2014. We did not include earlier calendar periods, as laboratory data before 2004 were incomplete due to interface problems between laboratory reporting software and GP practice software.¹⁴ Also, creatinine testing was incentivised in 2004 with the introduction of the diabetes Quality and Outcomes

Framework (QOF) and further in 2006 with the CKD QOF.¹⁴ To rule out any potential influence of incomplete data around 2004, we also examined the most recent 5-year calendar period separately in a sensitivity analysis. New users were defined as persons with at least one year of continuous registration in the CPRD before their first recorded ACEI/ARB prescription.

Laboratory data

All creatinine test results were extracted from the general practice records of the study population, using creatinine-specific codes in CPRD. Cross-reference was then made to creatinine test results identified from a broad Read code search. Any irrelevant codes were excluded. Renal function testing in the UK includes creatinine and potassium so it can be inferred that testing frequency is similar to creatinine for potassium. When we conducted analyses related to potassium levels, we repeated the procedure used to identify creatinine levels for potassium test results.

Patient characteristics

We obtained information for all patients on age, sex, calendar period of ACEI/ARB initiation (2004–2008 and 2010–2014), socioeconomic status (quintiles of the 2004 index of multiple deprivation scores), lifestyle factors (smoking, alcohol intake, and body mass index), baseline potassium level (≤ 5 or > 5 mmol/L), CKD, cardiovascular comorbidities (heart failure, myocardial infarction, hypertension, peripheral arterial disease, and arrhythmia), and diabetes.¹⁵ We used algorithms for smoking status, alcohol intake, and body mass index based on the most recent records in the CPRD before ACEI/ARB initiation.^{16 17} As measures of baseline creatinine and potassium levels, we used the single most recent measurement within 12 months before the first ACEI/ARB prescription. We calculated estimated glomerular filtration rate (eGFR) level from the most recent creatinine measurement and

CKD stage from the CKD-EPI equation.¹⁸ Cardiovascular comorbidities and diabetes were identified from both the CPRD and HES based on diagnoses recorded prior to ACEI/ARB initiation. The code lists for all variables are provided in the Appendix.

Patient involvement

The study included no patient involvement

Statistical analysis

We described ACEI/ARB users according to patient characteristics, both overall and according to creatinine monitoring status (no baseline or follow-up monitoring, baseline only, follow-up only, and both baseline and follow-up monitoring). Baseline monitoring was defined as a test performed on the date of drug initiation or within either 12 months before (generous interval) or one month before initiation (more ideal interval assumed to be driven by planned ACEI/ARB initiation). To accord with the post-initiation monitoring interval recommended from previous trial data, we considered only follow-up monitoring within the first 2 months after drug initiation.⁸

We calculated the proportion of persons in the total cohort of new users who had baseline and follow-up monitoring (within 1, 3, and 12 months before drug initiation and within 2 weeks, 1 month, and 2 months after initiation). We then computed the proportion of persons with both baseline and initial follow-up monitoring within the guideline-recommended interval of 2 weeks following drug commencement.

We repeated the analyses for continuing users, in order to examine adherence to the stricter guideline recommendations for ongoing monitoring (*i.e.*, monitoring within 1, 3, 6, and 12 months after the first retest).¹³ Continuation was defined as ACEI/ARB use beyond 30 days following the monitoring date, *i.e.*, when the end date of the first continuous course of therapy was after the date of the first monitoring date plus 30 days (to allow for stockpiling).

The end date of each prescription was calculated by adding the prescription duration (total number of tablets prescribed divided by the specified number of tablets per day) to the prescription date. In identifying continuous courses of therapy, we allowed for a 30-day gap between the end date of one prescription and the start of the next consecutive prescription.

In sensitivity analyses, we repeated the analyses (1) extending the follow-up window for the first follow-up monitoring from two to three weeks to account for minor delays; (2) including only the most recent calendar period (2009-2014) to account for temporal changes in data completeness and quality of care; (3) excluding patients with a hospital admission or discharge date within 1 month before or after their first ACEI/ARB prescription, in order to account for drug initiation and any subsequent renal function tests occurring in the hospital and therefore not captured in the CPRD; (4) focusing on specific patient subgroups (heart failure, myocardial infarction, hypertension, CKD (eGFR<60mls/min/1.73m²), peripheral arterial disease, and diabetes); and (5) defining drug use continuation as ACEI/ARB use beyond 90 days (instead of 30 days) after the first retest date.

We used the subcohort of patients with both baseline and follow-up monitoring to calculate the proportion of patients with creatinine increases $\geq 30\%$ or potassium levels >6 mmol/L at the first follow-up monitoring within 2 months after initiation, as well as the proportion of patients continuing treatment despite these contraindications for use.

Finally, we fitted a logistic regression model to identify patient characteristics associated with a severe decline in renal function (creatinine increase $\geq 30\%$ or potassium level >6 mmol/L) and compared these characteristics with those associated with receiving post-initiation follow-up monitoring within 2 weeks. The model included age, sex, CKD stage, cardiovascular comorbidities, diabetes, and baseline potassium level (>5 vs. ≤ 5 mmol/L). In three additional model-based sensitivity analyses, we repeated the analyses (1) excluding patients with a recent hospitalization (as defined above); (2) omitting baseline potassium from the model to examine the extent of potential overfitting when both baseline

potassium and CKD stage were kept in the model; and (3) also adjusting additional for ethnicity.

The study protocol was made available to the journal reviewers and approved by the London School of Hygiene and Tropical Medicine Ethics Committee (No. 6536) and the Independent Scientific Advisory Committee (ISAC) for Medicines and Healthcare Products Regulatory Agency (No. 16_025). All analyses were performed using STATA 14 statistical software package.

RESULTS

Serum creatinine monitoring before and after ACEI/ARB initiation

We identified 223,814 new users of ACEI/ARB. We compared these patients in four groups: 21,411 (10%) had no baseline or follow-up creatinine tests within 12 months before and 2 months after treatment initiation, 63,359 (28%) had only a baseline test, 33,185 (15%) had only follow-up tests, and 105,859 (47%) had both baseline and follow-up tests. (Table 1). Median age varied only slightly between the groups (60, 62, 59, and 63 years, respectively) and there were no substantial differences in socioeconomic status, lifestyle factors, or peripheral arterial disease. Compared with patients with neither pre- nor post-initiation monitoring, patients with both were more likely to have diagnosed hypertension (76% vs. 61%) and diabetes (20% vs. 7%), but less likely to have diagnosed heart failure (4% vs. 7%), myocardial infarction (4% vs. 18%), and arrhythmia (7% vs. 10%). Among patients with baseline monitoring, 83% did not have CKD, 13% stage 3a, 3% stage 3b, 0.5% stage 4 CKD. In the same population, 7% commenced ACEI/ARB therapy despite baseline potassium above 5 mmol/L. The median number of days between baseline monitoring and first prescription date was 40 days (interquartile range: 12-125 days).

Among all patients initiating ACEI/ARB therapy, the proportion of patients receiving creatinine testing before was 76% within 12 months before treatment initiation, declining to 34% within one month before initiation (Table 2). The proportion with follow-up testing after

treatment initiation was 29% within 2 weeks, increasing to 62% within 2 months. Among ACEI/ARB initiators who had a baseline test within 12 months, 21% also had a follow-up test within 2 weeks after starting treatment (Table 3). However, among patients undergoing testing within one month prior to treatment initiation, only 9% had also the recommended follow-up test within 2 weeks of treatment start. When we extended the follow-up window to three weeks, this proportion increased only to 14% (Table 3). Among patients continuing treatment, only 1% had follow-up measurements at 1, 3, 6, and 12 months after the first retest, in compliance with the strictest recommendation (eTable 1). These results were unchanged when the analysis was restricted to the most recent calendar period (eTable 1-2) and to patients with heart failure, myocardial infarction, hypertension, peripheral arterial disease, diabetes or no recent hospitalization (eTable 3). Only patients with CKD received a slightly higher degree of monitoring (13%) within two weeks following treatment initiation (eTable 3). The proportion with follow-up testing after treatment initiation was also unchanged when results were stratified by date of ACEI/ARB initiation in two-year intervals (eTable 4).

Serum creatinine and potassium changes after ACEI/ARB initiation

Among patients receiving the recommended renal function monitoring, 567 (1.2%) experienced a creatinine increase $\geq 30\%$ and 191 (0.4%) a potassium level >6 mmol/L at their first follow-up test within two months of treatment initiation (1.4% received in total either) (Table 4). Among these patients, 80% continued treatment beyond 30 days following the monitoring date (Table 4). The sensitivity analysis showed that 65% of patients with a creatinine increase $\geq 30\%$ and 60% of those with a potassium level >6 mmol/L continued also treatment beyond 90 days after the monitoring date (eTable 5). The results remained consistent for longer baseline monitoring intervals (eTable 5).

Patients at high risk for creatinine increases $\geq 30\%$

When we examined patient characteristics associated with a creatinine increase $\geq 30\%$ and adjusted for the other characteristics in a multivariable analysis (Table 5), we found an increased odds ratio (OR) for women (1.6-fold increased), for age above 70 years (at least 1.3-fold increased), for CKD stage 4 (1.6-fold increased), heart failure (2.9-fold increased), peripheral arterial disease (1.9-fold increased), myocardial infarction (1.6-fold increased), and hypertension (1.6-fold increased).

Patients at high risk for potassium >6 mmol/L

Baseline potassium level and CKD stage, but not age and sex, were associated with potassium levels >6 mmol/L after ACEI/ARB initiation. Thus, the OR was 7-fold increased for baseline potassium >5 mmol/L, two-fold increased for CKD stage 3a, 5-fold increased for stage 3b, and 11-fold increased for stage 4 (Table 5). Among cardiovascular comorbidities, heart failure was associated with the strongest OR of a potassium level >6 mmol/L (2.22, 95% CI: 1.38-3.58).

Monitoring high-risk patients

Some characteristics associated with an increased odds of having $\geq 30\%$ rise in creatinine were also associated with a greater likelihood of having a follow-up test within 2 weeks following drug initiation. These included older age: persons aged 70 years or above compared with ≤ 50 years (1.18, 95% CI: 1.13-1.23 for 70-79 years and 1.17, 95% CI: 1.11-1.23 for 80+ years), CKD stage 4 compared to no CKD (1.41, 95% CI: 1.20-1.66), heart failure (1.16, 95% CI: 1.08-1.23), and peripheral arterial disease (1.11, 95% CI: 1.02-1.20). However, other characteristics associated with an increased odds of having $\geq 30\%$ rise in creatinine were not associated with a greater likelihood of having a follow-up test within 2 weeks following drug initiation: there was no substantially increased OR ($>10\%$) associated with female sex (1.07,

95% CI: 1.04-1.09), prior history of myocardial infarction (0.77, 95% CI: 0.72-0.82), hypertension (1.05, 95% CI: 1.00-1.11), or baseline potassium >5 mmol/L (1.04, 95% CI: 0.99-1.09). When we excluded patients with a recent hospital admission, the reduced OR for myocardial infarction was no longer observed (0.93, 95% CI: 0.80-1.08) (eTable 6). Finally, the results remained consistent when we omitted adjustment for baseline potassium (data not shown) and when we adjusted additionally for ethnicity (eTable 6).

DISCUSSION

Only one tenth of patients initiating ACEI/ARBs in UK primary care appear to receive the guideline-recommended creatinine monitoring. One in 15 patients commenced ACEI/ARBs despite baseline potassium above the recommended level, which was also shown to be a strong predictor for severe post-initiation hyperkalaemia. Among monitored patients, a creatinine increase $\geq 30\%$ or a potassium level >6 mmol/L occurred in almost 1.5% of patients, and most did not discontinue therapy despite guideline recommendations to stop. Although patients with prior myocardial infarction, hypertension, or a high baseline potassium level were at higher risk of sudden decline in kidney function after ACEI/ARB initiation, there was no evidence that these patient groups were monitored more frequently while initiating the drugs.

Strengths and limitations

Several issues should be considered when interpreting our study results. Its large sample size increased precision. Use of the CPRD ensured that the study was general practice-based and not restricted to specific demographic, hospital, or insurance groups.

Over the time course of this study multiple factors have impacted on prescribing of ACEI/ARB and measurement of renal function in primary care, for example the introduction of the relevant NICE guidelines, and QOF reimbursement for testing in certain sub-groups. We also did not have information about clinical initiatives such as heart failure nurses and

ACEI/ARB stopping rules ('sick-day rules'). While our main results provide summary measures over a ten-year period, sensitivity analyses confirm that despite these changes, the proportion receiving the guideline suggested biochemical monitoring does not vary during the study period. We did not have access to blood tests performed in hospital systems, which may have been reported to GPs, but not recorded in CPRD. However, restricting the analysis to patients with no recent hospital admissions who were most likely to have had renal function measured and acted upon in secondary care had little effect on our findings. We did not examine testing during initiation of dual blockade with ACEI and ARB as this combination is now used very infrequently for patients with severe comorbidities who are likely to be monitored in secondary care. Although some patients may also have been seen in outpatient specialty clinics, it is common practice for specialists to ask GPs to initiate new drugs such as ACEI/ARBs, with local biochemical monitoring, limiting misclassification.

Consistent with findings from other studies,¹⁹ we found that approximately 50% of all ACEI/ARB initiators were monitored both before and after treatment start. If GPs are retesting renal function in patients at higher risk of substantial biochemical changes, we may have overestimated the proportion of patients with high potassium levels or creatinine increases compared with the untested lower-risk general population.

GP system software is used for issuing prescriptions, ensuring the accuracy of prescription data. However, it cannot be inferred that all patients actually redeemed their prescription at the pharmacy and start medication on the same day that it was prescribed.^{18 20} Similarly, the estimated coverage of prescriptions may not be completely accurate due to such factors as stockpiling and irregular use. We also do not know whether GPs contacted patients with elevated laboratory results to advise them to stop taking the medication prior to the end of their prescriptions. However, 80% of patients who developed creatinine increase $\geq 30\%$ after ACEI/ARB initiation were still issued a subsequent ACEI/ARB prescription.

We aimed to detect discontinuation related closely in time to first follow-up monitoring

and hence likely resulting from an elevated creatinine or potassium result. We therefore defined continuation as ACEI/ARB use beyond 30 days (the median prescription duration) after the monitoring date. Extending the definition of continuous use beyond 90 days reduced the risk of misclassifying patients as continuing treatment when they had in fact stopped. However, extending the duration also increased the risk of identifying discontinuation due to other reasons than creatinine/potassium increase, *e.g.*, death or cough. Diagnoses recorded in the CPRD generally have been found to have adequate validity for research purposes,^{21 22} particularly in the domains assessed by the QOF.^{23 24}

In the logistic regression analysis to estimate factors associated with creatinine increase $\geq 30\%$, we excluded patients without pre and post measurements (complete case analysis). If the recording of creatinine levels was not missing completely at random, the associations between patient characteristics and creatinine increase may have been underestimated.²⁵ While this assumption could not be tested directly, examination of baseline characteristics revealed no major differences in age, sex, socioeconomic status, and lifestyle between patients with and without pre- and post-monitoring. Furthermore, the results were consistent for each individual patient group examined. Patients with no testing before or after treatment initiation (including those with potentially haemolysed samples) only accounted for 10% of all ACEI/ARB initiators.

Comparison with other studies

To our knowledge, this is the largest study conducted to date on adherence to monitoring and discontinuation guidelines after ACEI/ARB initiation. Only one previous study¹⁹ examined monitoring according to guideline-recommended intervals (<14 days). All others have used longer intervals (*e.g.*, 30 days²⁶ or 6 months^{27 28}), which make interpretations and implications for clinical practice less clear. Poor adherence to monitoring guidelines after ACEI/ARB initiation is not restricted to the UK,^{19 28 29} but has also been reported in the US³⁰⁻

³², Canada,³³ and the Netherlands.^{26 34} Owing to our sample size, we were able to show that the lack of monitoring occurred in all patient groups with an indication for ACEI/ARB therapy.

A recent Dutch study, including 3,353 patients initiating ACEI/ARBs between 2005-2011, found that 19% had creatinine measured within 30 days and 66% within one year.²⁶ Creatinine increases above 30% occurred in 1.6% of patients, and among these 70% did not discontinue treatment.²⁶ A Scottish study of 4,056 patients with type 2 diabetes, prescribed an ACEI/ARB between 2005-2009, found that 19% had both a baseline (within 90 days) and follow-up measurement (within 2 weeks) of initiation. Within this cohort, 1.7% had both a creatinine increase of $\geq 30\%$ and potassium level ≥ 5.6 mmol/L.

The magnitude of the risk of severe renal impairment, as measured by creatinine increase in these observational studies, was consistent with our findings, but substantially higher than reported in clinical trials (*e.g.*, 0.2% in the ONTARGET trial).³⁵ It is not clear from the literature how often harm occurs around the time of initiation, when the risk of nephrotoxicity is thought to be greatest.⁸ If physicians are to understand why follow-up monitoring within 2 weeks of treatment start matters, the short-term risks need to be clarified. Until now, most studies have reported only on cumulative risk over entire courses of treatment, such as the 1.1% two-year risk for potassium of >6 mmol/L in the SOLVD trials of heart failure patients.³⁶ In contrast to clinical trial reviews, reporting a 0.2% (3/1818) risk of potassium >6 mmol/L, we found a 0.4% risk of hyperkalaemia already at time of first retesting after ACEI initiation.

Extending the previous literature, our results support that advanced age, advanced CKD, and heart failure, but not sex, increase the likelihood of being monitored.^{19 26 30} Consistent with some,^{26 30} but not all, previous studies,²⁸ we found no association for diabetes. However, these previous studies reporting an association for diabetes focused on monitoring within broader intervals (*e.g.*, 6 months),²⁸ where diabetes patients, irrespective of

ACEI/ARB initiation, were likely to receive blood testing owing to the diabetes QOF programme.

Determinants of increases in creatinine levels after ACEI/ARB initiation are less well understood than for hyperkalaemia, but increasing age is a consistently reported factor.¹⁹ Advanced CKD and a range of cardiovascular comorbidities (mostly associated with atherosclerosis) were also important determinants in our patient cohort. Consistent with previous studies, we found that the risk of hyperkalaemia was associated with CKD (likely due to impaired ability of the cortical collecting tubule to secrete potassium), heart failure (likely due to decreased delivery of sodium to the distal nephron), and high pre-treatment potassium levels.^{6 8 19 37} We did not observe an association with diabetes or increasing age, as could have been expected due to diabetic nephropathy or age-dependent hyporeninemic hypoaldosteronism.⁶

Clinical relevance

Several possible explanations exist for the divergence between the clinical guideline recommendations and the observed monitoring and response patterns in clinical practice. The first is *clinician nonadherence* to ordering tests. This may be due to inconsistent recommendations for timing and frequency of monitoring over time,⁶ consensus-based (rather than evidence-based) monitoring guidelines, and a lack of guidelines tailored to particular high-risk patients, such as those with CKD and heart failure. Although we found that follow-up monitoring correlated well with the risk of renal impairment after ACEI/ARB initiation for most patient groups, it was not observed for patients with myocardial infarction or pre-initiation high potassium. The second explanation may be *patient nonadherence* to ordered tests. This is particularly salient in UK primary care where blood samples may be taken in phlebotomy clinics that the patient has to visit rather than the GP practice. Patients may find it burdensome to have blood tests, and GPs have no direct economic incentives to ensure that

they are done. A third barrier is *lack of evidence of the clinical importance* of monitoring and its cost-effectiveness. ACEI/ARB-induced renal impairment is rare in clinical trials, even among patients with multiple risk factors for atherosclerotic renal artery stenosis.^{8 38} Trial results may therefore have led to a general perception that the rarity of renal impairment obviates the need for close monitoring. However, as observed in our data, the risks in real world practice may be somewhat higher and non-negligible. In addition, previous research has shown that potassium monitoring in high-risk patients with CKD and diabetes may reduce serious hyperkalaemia-associated adverse events.³⁹ Still, the extent to which an initial creatinine increase $\geq 30\%$ translates into adverse long-term outcomes in real-world patients remains to be clarified in future studies.

Generalisability, implications, and conclusions

The majority of patients initiating treatment with ACEI/ARBs experience only minor changes in renal function. However, substantial increases in creatinine levels after ACEI/ARB initiation may not be as rare as previously suggested, reinforcing the need for adherence to clinical guidelines for both pre- and post-initiating monitoring. Moreover, the post-initiation creatinine increase and potassium levels used in this study are widely recognised cut-off levels, making the results internationally applicable. The comparison with the previous literature also confirms that the lack of systematic monitoring is not exclusive to the UK. Of particular concern was that even when appropriate monitoring was performed, severe renal impairment only rarely led to treatment discontinuation. Individual patient counselling may also be helpful to ensure that those at highest risk are closely monitored. More work is needed to determine the prognostic importance of the changes in renal function that we have observed.

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Contributions

LT conceived the study idea and acquired data permissions. MS, KM, and LT designed the study. MS and KM performed data management and established the cohort. MS, KM, and LT reviewed the literature. The analyses were carried out by MS. All authors participated in the discussion and interpretation of the results. MS organised the writing and wrote the initial drafts. All authors critically revised the manuscript for intellectual content and approved the final version. MS is the guarantor.

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Table 1. Characteristics of patients initiating angiotensin converting-enzyme inhibitors or angiotensin-receptor blockers in the UK primary care during 2004-2014, by monitoring groups

	Serum creatinine monitoring*				Total
	No baseline or follow-up tests	Baseline test only	Follow-up test only	Baseline and follow-up test	
Total number	21,411 (100)	63,359 (100)	33,185 (100)	105,859 (100)	223,814 (100)
Female sex	8,882 (41)	27,722 (44)	14,570 (44)	49,109 (46)	100,283 (45)
Age (years)					
<50 years	5,019 (23)	13,697 (22)	8,732 (26)	19,910 (19)	47,358 (21)
50-59 years	5,485 (26)	15,135 (24)	9,115 (27)	24,866 (23)	54,601 (24)
60-69 years	4,863 (23)	15,586 (25)	7,776 (23)	27,790 (26)	56,015 (25)
70-79 years	3,579 (17)	12,193 (19)	5,066 (15)	22,152 (21)	42,990 (19)
80+ years	2,465 (12)	6,748 (11)	2,496 (8)	11,141 (11)	22,850 (10)
Calendar period					
2004-2008	14,814 (69)	40,667 (64)	19,808 (60)	60,902 (58)	136,191 (61)
2009-2014	6,597 (31)	22,692 (36)	13,377 (40)	44,957 (42)	87,623 (39)
SES quintiles					
1 (low)	5,153 (24)	15,290 (24)	8,533 (26)	25,577 (24)	54,553 (24)
2	4,725 (22)	14,331 (23)	7,887 (24)	24,851 (23)	51,794 (23)
3	4,341 (20)	13,028 (21)	6,890 (21)	22,629 (21)	46,888 (21)
4	4,254 (20)	12,140 (19)	5,931 (18)	19,318 (18)	41,643 (19)
5 (high)	2,925 (14)	8,508 (13)	3,898 (12)	13,359 (13)	28,690 (13)
Missing	13 (0)	62 (0)	46 (0)	125 (0)	246 (0)
Smoking status					
Never	7,860 (37)	22,496 (36)	12,229 (37)	36,895 (35)	79,480 (36)
Ever	13,433 (63)	40,797 (64)	20,915 (63)	68,939 (65)	144,084 (64)
Missing	118 (1)	66 (0)	41 (0)	25 (0)	250 (0)
Alcohol intake					
No use	2,556 (12)	7,819 (12)	3,409 (10)	11,088 (10)	24,872 (11)
Current	15,495 (72)	47,322 (75)	25,656 (77)	82,870 (78)	171,343 (77)
Former	1,328 (6)	4,499 (7)	1,933 (6)	7,490 (7)	15,250 (7)
Missing	2,032 (9)	3,719 (6)	2,187 (7)	4,411 (4)	12,349 (6)
BMI groups					
Underweight	282 (1)	700 (1)	304 (1)	1,008 (1)	2,294 (1)
Healthy weight	5,666 (26)	15,406 (24)	8,089 (24)	24,972 (24)	54,133 (24)
Overweight	7,677 (36)	23,755 (37)	12,484 (38)	40,556 (38)	84,472 (38)
Obesity	6,009 (28)	20,660 (33)	10,527 (32)	35,887 (34)	73,083 (33)
Missing	1,777 (8)	2,838 (4)	1,781 (5)	3,436 (3)	9,832 (4)
CKD (eGFR)†					
Stage ≤2 (≥60)	10,326 (48)	53,773 (85)	19,470 (59)	87,484 (83)	171,053 (76)
Stage 3a (45-59)	1,137 (5)	7,382 (12)	1,766 (5)	13,913 (13)	24,198 (11)
Stage 3b (30-44)	217 (1)	1,885 (3)	265 (1)	3,854 (4)	6,221 (3)
Stage 4 (15-29)	24 (0)	319 (1)	29 (0)	608 (1)	980 (0)
Not measured	9,707 (45)	0 (0)	11,655 (35)	0 (0)	21,362 (10)
CV comorbidities‡					
Heart failure	1,568 (7)	3,270 (5)	1,386 (4)	4,583 (4)	10,807 (5)
Myocardial infarction	3,881 (18)	4,653 (7)	3,203 (10)	4,620 (4)	16,357 (7)
Hypertension	13,023 (61)	44,273 (70)	24,195 (73)	80,946 (76)	162,437 (73)
Peripheral arterial disease	471 (2)	1,590 (3)	523 (2)	2,547 (2)	5,131 (2)
Arrhythmia	2,057 (10)	4,973 (8)	2,000 (6)	7,123 (7)	16,153 (7)
Diabetes mellitus	1,399 (7)	13,586 (21)	1,992 (6)	21,548 (20)	38,525 (17)

Abbreviations: CV, cardiovascular; CKD, chronic kidney disease; y, year

*Monitoring groups based on baseline (within 12 months before) and follow-up (within 2 months after) serum creatinine monitoring.

†Calculated from most recent creatinine measurement within 12 months before first prescription date.

‡Diagnosis ever registered before ACE/ARB initiation in CRPD or HES.

Table 2. Prevalence of baseline and follow-up serum creatinine monitoring among patients initiating angiotensin converting-enzyme inhibitors or angiotensin-receptor blockers, 2004-2014

	S-Creatinine, ≥1 test
Total number	n=223,814 (100%)
Baseline testing	
≤12 months before	169,218 (76%)
≤3 months before	115,348 (52%)
≤1 months before	75,476 (34%)
Follow-up testing	
≤2 weeks after	65,090 (29%)
≤1 month after	114,244 (51%)
≤2 months after	139,044 (62%)

Table 3. Prevalence of baseline and follow-up serum creatinine monitoring among patients initiating renin-angiotensin system blockade according to clinical guideline recommendations

	Clinical guidelines					All initiators n=223,814 (100%)	
	NICE Heart Failure	NICE MI	NICE/UKRA Hypertension	NICE CKD	GP Notebook	Wide baseline interval (≤12 month)	Ideal baseline interval (≤1 month)
Baseline testing	x	x	x	x	x	169,218 (76%)	75,476 (34%)
+ Follow-up test ≤2 weeks*	x	n/a	x	x	x	46,486 (21%)	19,679 (9%)
+ Follow-up test ≤3 weeks†						70,792 (32%)	30,451 (14%)

Abbreviations: CKD, chronic kidney disease; GP, General Practice; MI, myocardial infarction

*Follow-up test among those with baseline measurements

† Sensitivity analysis illustrating the importance of 2 vs. 3-week cut-off interval in follow-up test intervals.

Table 4. Proportion of new users of angiotensin converting-enzyme inhibitors or angiotensin-receptor blockers who continue or discontinue treatment according to guideline recommended cut-off levels of serum creatinine and potassium at follow-up testing*

	Continuation†	Discontinuation†	Total
Total number, %	42,942 (93.1)	3,178 (6.9)	46,120 (100)
Serum creatinine increase ≥30%, n (%)	462 (81.5)	105 (18.5)	567 (100)
Serum potassium >6 mmol/L, n (%)	150 (78.5)	41 (21.5)	191 (100)

*Calculated from the most recent measurements within 1 month before and 2 months after drug initiation.

† A patient was considered continuous users when the end date of the first continuous course of therapy was larger than the date of the first follow-up monitoring + 30 days (to allow for stock piling and irregular use)

Table 5. Association between patient characteristics and serum creatinine increase $\geq 30\%$ and follow-up monitoring within 2 weeks following initiation of renin-angiotensin system blockade

Characteristics	Odds ratio (95% confidence intervals)					
	S-creatinine monitoring within 2 weeks		S-creatinine increase $\geq 30\%*$		S-potassium increase $\geq 30\%*$	
	Age- and sex adjusted	Fully adjusted†	Age- and sex adjusted	Fully adjusted†	Age- and sex adjusted	Fully adjusted†
Female sex	1.07 (1.04-1.10)	1.07 (1.04-1.09)	1.39 (1.26-1.53)	1.63 (1.47-1.80)	0.87 (0.66-1.16)	0.94 (0.70-1.26)
Age (years)						
< 50 years	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
50-59 years	0.98 (0.94-1.01)	0.98 (0.95-1.02)	0.88 (0.74-1.05)	0.86 (0.72-1.03)	1.29 (0.79-2.11)	1.10 (0.67-1.81)
60-69 years	1.05 (1.02-1.09)	1.05 (1.01-1.09)	1.03 (0.88-1.21)	1.00 (0.85-1.19)	1.35 (0.84-2.17)	0.97 (0.60-1.58)
70-79 years	1.18 (1.14-1.23)	1.18 (1.13-1.23)	1.49 (1.27-1.74)	1.36 (1.15-1.61)	1.65 (1.02-2.66)	0.74 (0.43-1.26)
80+ years	1.20 (1.14-1.25)	1.17 (1.11-1.23)	2.72 (2.32-3.20)	2.02 (1.68-2.44)	2.75 (1.67-4.53)	0.73 (0.41-1.32)
CKD stage						
No CKD (≥ 60)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Stage 3a (45-59)	1.00 (0.96-1.04)	1.00 (0.96-1.04)	0.62 (0.53-0.73)	0.60 (0.51-0.70)	2.48 (1.66-3.71)	2.06 (1.36-3.11)
Stage 3b (30-44)	0.99 (0.93-1.06)	1.01 (0.94-1.08)	1.01 (0.82-1.24)	0.88 (0.71-1.09)	7.51 (4.75-11.9)	5.10 (3.16-8.22)
Stage 4 (15-29)	1.42 (1.21-1.67)	1.41 (1.20-1.66)	2.16 (1.52-3.05)	1.72 (1.18-2.51)	24.0 (13.5-42.6)	11.4 (6.07-21.4)
Comorbidities*						
Heart failure	1.15 (1.09-1.23)	1.16 (1.08-1.23)	4.00 (3.49-4.58)	2.93 (2.51-3.42)	2.90 (1.90-4.42)	2.22 (1.38-3.58)
MI	0.80 (0.75-0.85)	0.77 (0.72-0.82)	2.33 (1.98-2.74)	1.57 (1.32-1.87)	2.12 (1.33-3.39)	1.35 (0.80-2.25)
Hypertension	1.00 (0.97-1.02)	1.05 (1.00-1.11)	0.62 (0.56-0.68)	1.58 (1.36-1.84)	0.60 (0.45-0.80)	1.02 (0.63-1.65)
PAD	1.09 (1.01-1.18)	1.11 (1.02-1.20)	2.10 (1.70-2.60)	1.87 (1.50-2.33)	2.14 (1.18-3.86)	1.53 (0.82-2.88)
Arrhythmia	1.09 (1.03-1.14)	0.98 (0.95-1.01)	2.37 (2.07-2.71)	0.77 (0.69-0.86)	1.41 (0.90-2.21)	0.77 (0.56-1.05)
Diabetes mellitus	0.93 (0.90-0.96)	0.93 (0.90-0.96)	1.09 (0.97-1.22)	1.04 (0.92-1.18)	0.97 (0.69-1.36)	0.90 (0.63-1.29)
Baseline K>5 mmol/L	1.04 (1.00-1.10)	1.04 (0.99-1.09)	1.04 (0.86-1.25)	0.97 (0.80-1.17)	8.22 (6.14-11.0)	6.68 (4.94-9.02)

Abbreviations: CKD, chronic kidney disease; MI, myocardial infarction; PAD, peripheral arterial disease

* The increase was based on the difference between the most recent baseline measurements within 12 months before and first follow-up measurement within 2 months after drug initiation. All analyses were restricted to those with both baseline and follow-up measurements (n=105,859).

† Adjusted for sex, age, CKD, heart failure, MI, hypertension, PAD, arrhythmia, diabetes, and calendar period of prescription start

ONLINE SUPPLEMENTAL MATERIAL

eTable 1. Prevalence of baseline and follow-up serum creatinine monitoring among patients initiating renin-angiotensin system blockade according to clinical guideline recommendations, overall and in most recent calendar period (extended version of Table 3)

	Clinical guidelines						All initiators		Continuing users*	
	NICE Heart Failure	NICE CKD	NICE Hypertension	NICE MI	UK Renal Association	GP Notebook	Generous baseline interpretation (≤12 month)	Strict baseline interpretation (≤1 month)	Generous baseline interpretation (≤12 month)	Strict baseline interpretation (≤1 month)
2004-2014							n=223,814 (100%)		n=173,244 (100%)	
Baseline testing	x	x	x	x	x	x	169,218 (76%)	75,476 (34%)	169,218 (76%)	58,668 (34%)
Follow-up test (≤2 weeks)†	x	x	n/a	x	x	x	46,486 (21%)	19,679 (9%)	46,486 (21%)	18,457 (11%)
+ At least within one year after				x			36,424 (16%)	15,642 (7%)	36,424 (16%)	14,800 (9%)
+ 1 month thereafter‡						x	11,866 (5%)	5,643 (3%)	11,866 (5%)	5,388 (3%)
+ 1 and 3 months thereafter ‡						x	5,787 (3%)	2,933 (1%)	5,787 (3%)	2,791 (2%)
+ 1, 3, and 6 months thereafter ‡						x	3,390 (2%)	1,807 (1%)	3,390 (2%)	1,703 (1%)
+ 1, 3, 6, and 12 months thereafter‡						x	2,618 (1%)	1,439 (1%)	2,618 (1%)	1,353 (1%)
Follow-up test (≤3 weeks)§							70,792 (32%)	30,451 (14%)	70,792 (32%)	28,644 (17%)
2009-2014							n=87,623 (100%)		n=68,042 (100%)	
Baseline testing	x	x	x	x	x	x	67,649 (77%)	30,576 (35%)	67,649 (77%)	23,716 (35%)
Follow-up test (≤2 weeks)†	x	x	n/a	x	x	x	20,172 (23%)	8,707 (10%)	20,172 (23%)	8,169 (12%)
+ At least within one year after				x			15,416 (18%)	6,726 (8%)	15,416 (18%)	6,377 (9%)
+ 1 month thereafter‡						x	5,373 (6%)	2,607 (3%)	5,373 (6%)	2,500 (4%)
+ 1 and 3 months thereafter ‡						x	2,614 (3%)	1,365 (2%)	2,614 (3%)	1,302 (2%)
+ 1, 3, and 6 months thereafter ‡						x	1,525 (2%)	841 (1%)	1,525 (2%)	792 (1%)
+ 1, 3, 6, and 12 months thereafter‡						x	1,152 (1%)	650 (1%)	1,152 (1%)	606 (1%)
Follow-up test (≤3 weeks)§							31,007 (35%)	13,511 (15%)	31,007 (35%)	12,726 (19%)

Abbreviations: CKD, chronic kidney disease; GP, General Practice; MI, myocardial infarction; n/a, not specified

* A patient was considered continuous users when the end date of the first continuous course of therapy was larger than the date of the first follow-up monitoring + 30 days (to allow for stock piling and irregular use)

† Follow-up test among those with baseline measurements

‡ Within additional 1, 3, 6, and 12 months after correspond to within 1.5, 3.5, 6.5, and 12.5 months after ACE/ARB initiation, respectively.

§ Sensitivity analysis illustrating the importance of 2 vs. 3-week cut-off interval in follow-up test intervals.

eTable 2. Prevalence of baseline and follow-up serum creatinine monitoring among patients initiating angiotensin converting-enzyme inhibitors or angiotensin-receptor blockers, overall and most recent calendar period (extended version of Table 1)

	S-Creatinine, ≥ 1 test, n (%)	
	Total cohort (n=223,814)	2009-2014 cohort (n=87,623)
Baseline testing		
≤ 12 months before	169,218 (75.6)	67,649 (77.2)
≤ 3 months before	115,348 (51.5)	46,925 (53.6)
≤ 1 months before	75,476 (33.7)	30,576 (34.9)
Follow-up testing		
≤ 2 weeks after	65,090 (29.1)	27,933 (31.9)
≤ 1 month after	114,244 (51.0)	48,877 (55.8)
≤ 2 months after	139,044 (62.1)	58,334 (66.6)

eTable 3. Prevalence of baseline and follow-up serum creatinine monitoring among patients initiating renin-angiotensin system blockade according to clinical guideline recommendations

	All initiators n=223,814 (100%)	
	Generous baseline interpretation (≤ 12 month)	Strict baseline interpretation (≤ 1 month)
Overall (n=223,814)		
Baseline testing	169,218 (76%)	75,476 (34%)
+ Follow-up test ≤ 2 weeks*	46,486 (21%)	19,679 (9%)
+ Follow-up test ≤ 3 weeks†	70,792 (32%)	30,451 (14%)
Overall, no recent hospitalization (n=197,291)		
Baseline testing	153,353 (78%)	68,883 (35%)
+ Follow-up test ≤ 2 weeks*	42,967 (22%)	18,222 (9%)
+ Follow-up test ≤ 3 weeks†	65,881 (33%)	28,375 (14%)
Heart failure patients (n=10,807)		
Baseline testing	7,853 (73%)	3,277 (30%)
+ Follow-up test ≤ 2 weeks*	2,229 (21%)	951 (9%)
+ Follow-up test ≤ 3 weeks†	3,085 (29%)	1,340 (12%)
Myocardial infarction patients (n=16,357)		
Baseline testing	9,273 (57%)	3,766 (23%)
+ Follow-up test ≤ 2 weeks*	1,816 (11%)	726 (4%)
+ Follow-up test ≤ 3 weeks†	2,645 (16%)	1,078 (7%)
Hypertension patients (n=162,437)		
Baseline testing	125,219 (77%)	54,174 (33%)
+ Follow-up test ≤ 2 weeks*	35,502 (22%)	14,559 (9%)
+ Follow-up test ≤ 3 weeks†	54,585 (34%)	22,710 (14%)
CKD patients (n=31,399)		
Baseline testing	27,961 (89%)	13,913 (44%)
+ Follow-up test ≤ 2 weeks*	8,457 (27%)	4,198 (13%)
+ Follow-up test ≤ 3 weeks†	12,482 (40%)	6,324 (20%)
PAD patients (n=5,131)		
Baseline testing	4,137 (81%)	1,671 (33%)
+ Follow-up test ≤ 2 weeks*	1,198 (23%)	458 (9%)
+ Follow-up test ≤ 3 weeks†	1,751 (34%)	698 (14%)
Diabetes patients (n=38,525)		
Baseline testing	35,134 (91%)	15,907 (41%)
+ Follow-up test ≤ 2 weeks*	9,161 (24%)	3,794 (10%)
+ Follow-up test ≤ 3 weeks†	14,093 (37%)	5,994 (16%)

Abbreviations: CKD, chronic kidney disease

*Follow-up test among those with baseline measurements

† Sensitivity analysis illustrating the importance of 2 vs. 3-week cut-off interval in follow-up test intervals.

eTable 4: Prevalence of baseline and follow-up serum creatinine monitoring among patients initiating renin-angiotensin system blockade according to clinical guideline recommendations, stratified by year of ACEI/ARB initiation, 2004-2014

	All initiators n=223,814		2004/2005 n=51,529		2006/2007 n=59,881		2008/2009 n=47,472		2010/2011 n=37,183		2012/2014 n=27,749	
	Wide baseline interval (≤12 month)	Ideal baseline interval (≤1 month)	Wide baseline interval (≤12 month)	Ideal baseline interval (≤1 month)	Wide baseline interval (≤12 month)	Ideal baseline interval (≤1 month)	Wide baseline interval (≤12 month)	Ideal baseline interval (≤1 month)	Wide baseline interval (≤12 month)	Ideal baseline interval (≤1 month)	Wide baseline interval (≤12 month)	Ideal baseline interval (≤1 month)
Baseline testing	169,218 (76%)	75,476 (34%)	36,649 (71%)	14,710 (29%)	45,801 (76%)	21,079 (35%)	36,585 (77%)	17,265 (36%)	13,045 (35%)	21,443 (77%)	9,377 (34%)	
+ Follow-up test ≤2 weeks*	46,486 (21%)	19,679 (9%)	8,139 (16%)	2,929 (6%)	12,587 (21%)	5,509 (9%)	10,595 (22%)	4,769 (10%)	3,720 (10%)	6,556 (24%)	2,752 (10%)	
+ Follow-up test ≤3 weeks†	70,792 (32%)	30,451 (14%)	12,222 (24%)	4,550 (9%)	19,102 (32%)	8,504 (14%)	16,229 (34%)	7,392 (16%)	5,768 (16%)	10,054 (36%)	4,237 (15%)	

*Follow-up test among those with baseline measurements
† Sensitivity analysis illustrating the importance of 2 vs. 3-week cut-off interval in follow-up test intervals.

eTable 5. Proportion of new users of ACEI/ARB who continue or discontinue treatment according to guideline recommended cut-off levels of serum creatinine and potassium at follow-up testing

	Wide monitoring interval (12 month before to the first monitoring within 2 month after)			Narrow monitoring interval (1 month before to the first monitoring within 2 month after)		
	Continuation*	Discontinuation*	Total	Continuation*	Discontinuation*	Total
+ 30 day window						
Total number, %	113,210 (92.5)	9,170 (7.5)	122,380 (100)	48,772 (93.0)	3,676 (7.0)	52,448 (100)
S-creatinine increase $\geq 30\%$, n (%)	1,679 (80.7)	401 (19.3)	2,080 (100)	541 (81.5)	123 (18.5)	664 (100)
S-potassium >6 mmol/L, n (%)	472 (81.9)	104 (18.1)	576 (100)	204 (80.6)	49 (19.4)	253 (100)
+ 90 day window						
Total number, %	86,620 (81.8)	19,239 (18.2)	105,859 (100)	37,920 (82.2)	8,200 (17.8)	46,120 (100)
S-creatinine increase $\geq 30\%$, n (%)	1,111 (64.7)	605 (35.3)	1,716 (100)	372 (65.6)	195 (34.4)	567 (100)
S-potassium >6 mmol/L, n (%)	261 (61.0)	167 (39.0)	428 (100)	110 (57.6)	81 (42.4)	191 (100)

Calculated from the measurements closest to the first prescription date in the corresponding intervals, i.e., the most recent baseline measurements and the first follow-up measurement.

* A patient was considered continuous users when the end date of the first continuous course of therapy was larger than the date of the first follow-up monitoring + 30 days or + 90days (to allow for stock piling and irregular use)

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eTable 6. Association between patient characteristics and serum creatinine increase $\geq 30\%$ and follow-up monitoring within 2 weeks following initiation of renin-angiotensin system blockade

Characteristics	Adjusted odds ratio (95% confidence intervals)					
	Additional adjustment for ethnicity†			Excluding patient with hospitalisation within 30 days†		
	S-creatinine monitoring within 2 weeks	S-creatinine increase $\geq 30\%*$	S-potassium >6 mmol/L*	S-creatinine monitoring within 2 weeks	S-creatinine increase $\geq 30%*$	S-potassium >6 mmol/L*
Female sex	1.04 (1.01-1.08)	1.91 (1.64-2.23)	1.09 (0.70-1.71)	1.06 (1.04-1.09)	1.66 (1.44-1.86)	0.94 (0.69-1.27)
Age (years)						
< 50 years	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
50-59 years	0.97 (0.92-1.02)	0.91 (0.71-1.16)	1.65 (0.77-3.52)	0.99 (0.95-1.03)	0.86 (0.70-1.05)	1.17 (0.70-1.98)
60-69 years	1.04 (0.99-1.10)	0.86 (0.68-1.10)	1.23 (0.57-2.65)	1.07 (1.02-1.11)	1.05 (0.84-1.26)	0.98 (0.58-1.64)
70-79 years	1.22 (1.15-1.30)	1.22 (0.96-1.55)	0.83 (0.36-1.95)	1.19 (1.14-1.24)	1.40 (1.11-1.73)	0.72 (0.41-1.27)
80+ years	1.15 (1.06-1.24)	1.61 (1.21-2.14)	0.58 (0.21-1.58)	1.17 (1.11-1.24)	2.16 (1.70-2.66)	0.71 (0.38-1.33)
CKD stage						
No CKD (≥ 60)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Stage 3a (45-59)	1.00 (0.94-1.06)	0.51 (0.39-0.66)	1.82 (0.93-3.55)	1.00 (0.96-1.04)	0.58 (0.44-0.76)	2.40 (1.57-3.68)
Stage 3b (30-44)	1.06 (0.95-1.17)	0.99 (0.71-1.39)	5.66 (2.70-11.9)	0.98 (0.92-1.06)	0.92 (0.77-1.17)	5.15 (3.09-8.58)
Stage 4 (15-29)	1.51 (1.17-1.94)	1.65 (0.87-3.12)	15.0 (5.84-38.8)	1.42 (1.20-1.70)	1.86 (1.27-2.61)	12.2 (6.29-23.7)
Comorbidities*						
Heart failure	1.10 (0.99-1.21)	3.07 (2.41-3.91)	2.16 (0.96-4.85)	1.15 (1.06-1.24)	2.57 (2.11-3.13)	2.68 (1.59-4.51)
MI	0.78 (0.72-0.86)	1.76 (1.35-2.30)	1.10 (0.46-2.63)	0.89 (0.82-0.97)	1.59 (1.27-2.06)	1.89 (1.05-3.41)
Hypertension	0.97 (0.90-1.05)	1.51 (1.19-1.92)	1.01 (0.44-2.30)	1.04 (0.99-1.11)	1.40 (1.11-1.78)	0.81 (0.46-1.42)
PAD	1.09 (0.96-1.23)	1.63 (1.13-2.35)	1.52 (0.54-4.29)	1.16 (1.06-1.26)	1.74 (1.33-2.27)	1.75 (0.92-3.32)
Arrhythmia	0.96 (0.92-1.01)	0.84 (0.71-0.99)	0.72 (0.44-1.17)	0.98 (0.94-1.01)	0.80 (0.70-0.91)	0.74 (0.53-1.03)
Diabetes mellitus	0.91 (0.87-0.96)	1.15 (0.96-1.37)	0.58 (0.31-1.09)	0.93 (0.90-0.96)	1.06 (0.91-1.22)	0.93 (0.64-1.35)
Baseline K >6 mmol/L	1.05 (0.98-1.13)	0.86 (0.63-1.15)	7.98 (5.06-12.6)	1.03 (0.98-1.08)	1.01 (0.88-1.14)	6.44 (4.69-8.84)

Abbreviations: CKD, chronic kidney disease; MI, myocardial infarction; PAD, peripheral arterial disease

* The increase was based on the difference between the most recent baseline measurements within 12 months before and first follow-up measurement within 2 months after drug initiation. All analyses were restricted to those with both baseline and follow-up measurements (n=105,859).

† The main model was adjusted for sex, age, CKD, heart failure, MI, hypertension, PAD, arrhythmia, diabetes, and calendar period of prescription start

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1
2 **Appendix: Code list**
3
4

5 **Serum creatinine**

6 Code	Classification	Description
7 4....00	Read	laboratory procedures
8 41...00	Read	laboratory procedures -general
9 4....11	Read	investigation-laboratory
10 4....12	Read	test - laboratory
11 412..00	Read	laboratory procedure performed
12 413..00	Read	laboratory test requested
13 41B..00	Read	laboratory test due
14 41BZ.00	Read	laboratory test nos due
15 41G..00	Read	laboratory administration procedures
16 41G3.00	Read	specimen received in laboratory
17 41H..00	Read	review of patient laboratory test report
18 44J..00	Read	blood urea/renal function
19 44J3.00	Read	serum creatinine
20 44J3000	Read	serum creatinine abnormal
21 44J3100	Read	serum creatinine low
22 44J3200	Read	serum creatinine normal
23 44J3300	Read	serum creatinine raised
24 44J3z00	Read	serum creatinine nos
25 44JC.00	Read	corrected plasma creatinine level
26 44JD.00	Read	corrected serum creatinine level
27 44JF.00	Read	plasma creatinine level
28 44JZ.00	Read	blood urea/renal function nos
29 451..00	Read	renal function tests
30 4511	Read	renal function tests normal
31 4512	Read	renal function tests abnormal
32 4515	Read	differential renal function
33 4516	Read	renal function tests borderline
34 4519	Read	deteriorating renal function
35 451H.00	Read	recovery of renal function
36 451Z.00	Read	renal function test nos
37 4Q40.00	Read	creatinine level
38 4Z...00	Read	laboratory procedure nos
39 8A6..00	Read	renal function monitoring
40 8H7P.11	Read	refer to pathology laboratory
41 8HP..00	Read	referral for laboratory tests
42 98F..00	Read	reason for repeat laboratory test
43 9b0P.00	Read	laboratory request
44 9b0Q.00	Read	laboratory result
45 9mG..00	Read	renal function monitoring invitation
46 9mG0.00	Read	renal function monitoring invitation first letter
47 9N1Q.00	Read	seen in diabetic clinic

C345.00	Read	gout due to impairment of renal function
K060.11	Read	impaired renal function
K08..00	Read	impaired renal function disorder
K08y.00	Read	other impaired renal function disorder
K08y300	Read	renal function impairment with growth failure
K08yz00	Read	other impaired renal function disorder nos
K08z.00	Read	impaired renal function disorder nos
R144.00	Read	[d]renal function test abnormal
R1yz.11	Read	[d]unexplained laboratory result
ZV72600	Read	[v]laboratory examination

Serum potassium

Code	Classification	Description
44I4.00	Read	serum potassium
4....12	Read	test - laboratory
4Z...00	Read	laboratory procedure nos
451..00	Read	renal function tests
C368.00	Read	hypokalaemia
412..00	Read	laboratory procedure performed
4512.00	Read	renal function tests abnormal
4511.00	Read	renal function tests normal
8A6..00	Read	renal function monitoring
C367.00	Read	hyperkalaemia
K060.11	Read	impaired renal function
4....11	Read	investigation-laboratory
K08..00	Read	impaired renal function disorder
R144.00	Read	[d]renal function test abnormal
8HP..00	Read	referral for laboratory tests
4....00	Read	laboratory procedures
44J..00	Read	blood urea/renal function
44h8.00	Read	plasma potassium level
44h0.00	Read	blood potassium level
4I46.00	Read	sweat potassium level
413..00	Read	laboratory test requested
C345.00	Read	gout due to impairment of renal function
41...00	Read	laboratory procedures -general
41B..00	Read	laboratory test due
411..00	Read	laboratory test not necessary
4516.00	Read	renal function tests borderline
K08z.00	Read	impaired renal function disorder nos
4519.00	Read	deteriorating renal function
44I4200	Read	low serum potassium level
44I4000	Read	normal serum potassium level
44I4100	Read	raised serum potassium level
44JZ.00	Read	blood urea/renal function nos

1			
2	1F34.00	Read	dietary potassium - low
3	1F3..11	Read	dietary potassium intake
4	41BZ.00	Read	laboratory test nos due
5	ZC61c00	Read	potassium supplementation
6	ZV72600	Read	[v]laboratory examination
7	451Z.00	Read	renal function test nos
8	K08y.00	Read	other impaired renal function disorder
9	4Q42.00	Read	potassium level
10	R1yz.11	Read	[d]unexplained laboratory result
11	ZC28.00	Read	advice to change potassium intake
12	4I34.11	Read	potassium in sample
13	9b0Q.00	Read	laboratory result
14	K08yz00	Read	other impaired renal function disorder nos
15	4515.00	Read	differential renal function
16	1F36.00	Read	dietary potassium - high
17	4I34.00	Read	fluid sample potassium
18	1F35.00	Read	dietary potassium - average
19	5714.00	Read	potassium 40 whole body count
20	41G..00	Read	laboratory administration procedures
21	41H..00	Read	review of patient laboratory test report
22	9b0P.00	Read	laboratory request
23	8B77.00	Read	potassium supplementation
24	451H.00	Read	recovery of renal function
25	9mG..00	Read	renal function monitoring invitation
26	98F..00	Read	reason for repeat laboratory test
27	13BD.00	Read	low potassium diet
28	9mG0.00	Read	renal function monitoring invitation first letter
29	41G3.00	Read	specimen received in laboratory
30	4IA5.00	Read	unsatisfactory laboratory analysis

End stage renal disease

31	Code	Classification	Description
32	14S2.00	Read	h/o: kidney recipient
33	14V2.00	Read	h/o: renal dialysis
34	14V2.11	Read	h/o: kidney dialysis
35	1Z14.00	Read	chronic kidney disease stage 5
36	1Z1K.00	Read	chronic kidney disease stage 5 with proteinuria
37	1Z1K.11	Read	ckd stage 5 with proteinuria
38	1Z1L.00	Read	chronic kidney disease stage 5 without proteinuria
39	1Z1L.11	Read	ckd stage 5 without proteinuria
40	4I29.00	Read	peritoneal dialysis sample
41	4N0..00	Read	dialysis fluid urea level
42	4N2..00	Read	dialysis fluid glucose level
43	67P4100	Read	discussion about kidney transplantation
44	7A60.00	Read	arteriovenous shunt

1			
2	7A60000	Read	insertion of arteriovenous prosthesis
3	7A60100	Read	creation of arteriovenous fistula nec
4	7A60111	Read	creation of radial-cephalic fistula
5	7A60112	Read	creation of brachial-cephalic fistula
6	7A60200	Read	attention to arteriovenous shunt
7	7A60300	Read	removal of infected arteriovenous shunt
8	7A60400	Read	banding of arteriovenous fistula
9	7A60500	Read	thrombectomy of arteriovenous fistula
10	7A60600	Read	creation of graft fistula for dialysis
11	7A60y00	Read	other specified arteriovenous shunt
12	7A60z00	Read	arteriovenous shunt nos
13	7A61100	Read	repair of acquired arteriovenous fistula
14	7A61111	Read	ligation of acquired arteriovenous fistula
15	7A61400	Read	ligation of acquired arteriovenous fistula
16	7A61900	Read	ligation of arteriovenous dialysis fistula
17	7A61A00	Read	ligation of arteriovenous dialysis graft
18	7B00.00	Read	transplantation of kidney
19	7B00100	Read	transplantation of kidney from live donor
20	7B00111	Read	allotransplantation of kidney from live donor
21	7B00200	Read	transplantation of kidney from cadaver
22	7B00211	Read	allotransplantation of kidney from cadaver
23	7B00212	Read	cadaveric renal transplant
24	7B00300	Read	allotransplantation of kidney from cadaver heart-beating
25	7B00400	Read	allotransplantation kidney from cadaver heart non-beating
26	7B00600	Read	xenograft renal transplant
27	7B00y00	Read	other specified transplantation of kidney
28	7B00z00	Read	transplantation of kidney nos
29	7B01500	Read	transplant nephrectomy
30	7B01511	Read	excision of rejected transplanted kidney
31	7B06300	Read	exploration of renal transplant
32	7B0F.00	Read	interventions associated with transplantation of kidney
33	7B0F100	Read	pre-transplantation of kidney work-up
34	7B0F300	Read	post-transplantation of kidney examination
35	7B0Fy00	Read	os interventions associated with transplantation of kidney
36	7B0Fz00	Read	interventions associated with transplantation of kidney nos
37	7L1A.00	Read	compensation for renal failure
38	7L1A000	Read	renal dialysis
39	7L1A011	Read	thomas intravascular shunt for dialysis
40	7L1A100	Read	peritoneal dialysis
41	7L1A.11	Read	dialysis for renal failure
42	7L1A200	Read	haemodialysis nec
43	7L1A300	Read	haemofiltration
44	7L1A400	Read	automated peritoneal dialysis
45	7L1A500	Read	continuous ambulatory peritoneal dialysis
46	7L1A600	Read	peritoneal dialysis nec

1			
2	7L1Ay00	Read	other specified compensation for renal failure
3	7L1Az00	Read	compensation for renal failure nos
4			
5	7L1B.00	Read	placement ambulatory apparatus compensation renal failure
6	7L1B000	Read	insertion of ambulatory peritoneal dialysis catheter
7			
8	7L1B100	Read	removal of ambulatory peritoneal dialysis catheter
9	7L1B.11	Read	placement ambulatory dialysis apparatus - compens renal fail
10	7L1B200	Read	flushing of peritoneal dialysis catheter
11			
12	7L1By00	Read	placement ambulatory apparatus- compensate renal failure os
13	7L1C.00	Read	placement other apparatus for compensation for renal failure
14	7L1C000	Read	insertion of temporary peritoneal dialysis catheter
15			
16	7L1Cy00	Read	placement other apparatus- compensate for renal failure os
17	7L1Cz00	Read	placement other apparatus- compensate for renal failure nos
18			
19	8L50.00	Read	renal transplant planned
20	9b8K.00	Read	transplantation surgery
21	9Ot5.00	Read	predicted stage chronic kidney disease
22			
23	G72C.00	Read	ruptured aneurysm of dialysis vascular access
24	G72D.00	Read	aneurysm of dialysis arteriovenous fistula
25	G72D100	Read	aneurysm of needle site of dialysis arteriovenous fistula
26			
27	G72D200	Read	aneurysm of anastomotic site of dialysis av fistula
28	G760.00	Read	acquired arteriovenous fistula
29			
30	Gy1..00	Read	stenosis of dialysis vascular access
31	Gy21.00	Read	thrombosis of dialysis arteriovenous fistula
32	Gy3..00	Read	occlusion of dialysis vascular access
33			
34	Gy31.00	Read	occlusion of dialysis arteriovenous fistula
35	Gy40.00	Read	infection of dialysis arteriovenous graft
36	Gy41.00	Read	infection of dialysis arteriovenous fistula
37			
38	Gy51.00	Read	haemorrhage of dialysis arteriovenous fistula
39	Gy60.00	Read	rupture of dialysis arteriovenous graft
40			
41	K050.00	Read	end stage renal failure
42	K05..12	Read	end stage renal failure
43	K055.00	Read	chronic kidney disease stage 5
44			
45	K0B5.00	Read	renal tubulo-interstitial disorders in transplant rejectn
46	K0D..00	Read	end-stage renal disease
47			
48	Kyu1C00	Read	[x]renal tubulo-interstitial disorders/transplant rejection
49	SP01500	Read	mechanical complication of dialysis catheter
50	SP01700	Read	mechanical complication of arterio-venous surgical fistula
51			
52	SP05613	Read	[x] peritoneal dialysis associated peritonitis
53	SP06B00	Read	continuous ambulatory peritoneal dialysis associated perit
54	SP07G00	Read	stenosis of arteriovenous dialysis fistula
55			
56	SP07N00	Read	arteriovenous fistula thrombosis
57	SP08011	Read	det.ren.func.after ren.transpl
58	SP08300	Read	kidney transplant failure and rejection
59	SP08J00	Read	chronic rejection of renal transplant
60			
	SP08N00	Read	unexplained episode of renal transplant dysfunction
	SP08R00	Read	renal transplant rejection

1			
2	SP08T00	Read	urological complication of renal transplant
3	SP08W00	Read	vascular complication of renal transplant
4	SP0E.00	Read	disorders associated with peritoneal dialysis
5	SP0G.00	Read	anaphylactoid reaction due to haemodialysis
6	TA02000	Read	accid cut
7	TA22.00	Read	failure of sterile precautions during perfusion
8	TA22000	Read	failure of sterile precautions during kidney dialysis
9	TB00100	Read	kidney transplant with complication
10	TB00111	Read	renal transplant with complication
11	TB11.00	Read	kidney dialysis with complication
12	TB11.11	Read	renal dialysis with complication
13	U612200	Read	[x]failure sterile precautions dur kidney dialys/other perf
14	Z1A..00	Read	dialysis training
15	Z1A1.00	Read	peritoneal dialysis training
16	Z1A2.00	Read	haemodialysis training
17	Z919.00	Read	care of haemodialysis equipment
18	Z919100	Read	priming haemodialysis lines
19	Z919300	Read	reversing haemodialysis lines
20	Z91A.00	Read	peritoneal dialysis bag procedure
21	ZV42000	Read	[v]kidney transplanted
22	ZV45100	Read	[v]renal dialysis status
23	ZV56.00	Read	[v]aftercare involving intermittent dialysis
24	ZV56000	Read	[v]aftercare involving extracorporeal dialysis
25	ZV56011	Read	[v]aftercare involving renal dialysis nos
26	ZV56100	Read	[v]preparatory care for dialysis
27	ZV56y00	Read	[v]other specified aftercare involving intermittent dialysis
28	ZV56y11	Read	[v]aftercare involving peritoneal dialysis
29	ZVu3G00	Read	[x]other dialysis
30	I77.0	ICD-10	arteriovenous fistula
31	N16.5	ICD-10	renal tubulo-interstitial disorders in transplant rejection
32	N18.5	ICD-10	chronic kidney disease
33	T82.4	ICD-10	mechanical complication of vascular dialysis catheter
34	T86.1	ICD-10	kidney transplant failure and rejection
35	Y60.2	ICD-10	during kidney dialysis or other perfusion
36	Y61.2	ICD-10	during kidney dialysis or other perfusion
37	Y62.2	ICD-10	during kidney dialysis or other perfusion
38	Y84.1	ICD-10	kidney dialysis
39	Z49	ICD-10	care involving dialysis
40	Z49.0	ICD-10	preparatory care for dialysis
41	Z49.1	ICD-10	extracorporeal dialysis
42	Z49.2	ICD-10	other dialysis
43	Z94.0	ICD-10	kidney transplant status
44	Z99.2	ICD-10	dependence on renal dialysis

Ischemic heart disease

	Code	Classification	Description
1			
2			
3	14A3.00	Read	h/o: myocardial infarct <60
4	14A4.00	Read	h/o: myocardial infarct >60
5	14A5.00	Read	h/o: angina pectoris
6	14AA.00	Read	h/o: heart disease nos
7	14AH.00	Read	h/o: myocardial infarction in last year
8	14AJ.00	Read	h/o: angina in last year
9	14AL.00	Read	h/o: treatment for ischaemic heart disease
10	14AT.00	Read	history of myocardial infarction
11	14AW.00	Read	h/o acute coronary syndrome
12	182A.00	Read	Chest pain on exertion
13	187..00	Read	frequency of angina
14	1J61.00	Read	suspected ischaemic heart disease
15	3213111	Read	Positive exercise ECG test
16	322..00	Read	ecg: myocardial ischaemia
17	3222	Read	ecg:shows myocardial ischaemia
18	322Z.00	Read	ecg: myocardial ischaemia nos
19	323..00	Read	ecg: myocardial infarction
20	3232	Read	ecg: old myocardial infarction
21	3233	Read	ecg: antero-septal infarct.
22	3234	Read	ecg:posterior/inferior infarct
23	3235	Read	ecg: subendocardial infarct
24	3236	Read	ecg: lateral infarction
25	323Z.00	Read	ecg: myocardial infarct nos
26	32B..00	Read	ecg: q wave
27	32B2.00	Read	ecg: q wave abnormal
28	32B3.00	Read	ecg: q wave pathological
29	32BZ.00	Read	ecg: q wave nos
30	32E4.00	Read	ecg: s-t depression
31	44H3.00	Read	cardiac enzymes abnormal
32	44H3000	Read	cardiac enzymes abnormal - first set
33	44HJ.00	Read	Plasma creatinine phosphokinase MB isoenzyme level
34	44MH.00	Read	Plasma troponin T level
35	44p2.00	Read	cardiac troponin positive
36	5533	Read	angiocardiology abnormal
37	5543	Read	coronary arteriograph.abnormal
38	5C11.00	Read	radionuclide heart study abnormal
39	661M000	Read	angina self-management plan agreed
40	662..00	Read	Cardiac disease monitoring
41	662..11	Read	Heart disease monitoring
42	662K.00	Read	angina control
43	662K000	Read	angina control - good
44	662K100	Read	angina control - poor
45	662K200	Read	angina control - improving
46	662K300	Read	angina control - worsening

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1			
2	662Kz00	Read	angina control nos
3	662N.00	Read	CHD monitoring
4	662Z.00	Read	Cardiac disease monitoring NOS
5	66f.00	Read	Cardiovascular disease monitoring
6	66f1.00	Read	Cardiovascular disease interim monitoring
7	6A2..00	Read	coronary heart disease annual review
8	6A4..00	Read	coronary heart disease review
9	7920	Read	saphenous vein graft replacement of coronary artery
10	7920000	Read	saphenous vein graft replacement of one coronary artery
11	7920100	Read	saphenous vein graft replacement of two coronary arteries
12	7920.11.00	Read	saphenous vein graft bypass of coronary artery
13	7920200	Read	saphenous vein graft replacement of three coronary arteries
14	7920300	Read	saphenous vein graft replacement of four+ coronary arteries
15	7920y00	Read	saphenous vein graft replacement of coronary artery os
16	7920z00	Read	saphenous vein graft replacement coronary artery nos
17	7921	Read	other autograft replacement of coronary artery
18	7921000	Read	autograft replacement of one coronary artery nec
19	792..11	Read	coronary artery bypass graft operations
20	7921100	Read	autograft replacement of two coronary arteries nec
21	7921.11.00	Read	other autograft bypass of coronary artery
22	7921200	Read	autograft replacement of three coronary arteries nec
23	7921300	Read	autograft replacement of four of more coronary arteries nec
24	7921y00	Read	other autograft replacement of coronary artery os
25	7921z00	Read	other autograft replacement of coronary artery nos
26	7922	Read	allograft replacement of coronary artery
27	7922000	Read	allograft replacement of one coronary artery
28	7922100	Read	allograft replacement of two coronary arteries
29	7922.11.00	Read	allograft bypass of coronary artery
30	7922200	Read	allograft replacement of three coronary arteries
31	7922300	Read	allograft replacement of four or more coronary arteries
32	7922y00	Read	other specified allograft replacement of coronary artery
33	7922z00	Read	allograft replacement of coronary artery nos
34	7923	Read	prosthetic replacement of coronary artery
35	7923000	Read	prosthetic replacement of one coronary artery
36	7923100	Read	prosthetic replacement of two coronary arteries
37	7923.11.00	Read	prosthetic bypass of coronary artery
38	7923200	Read	prosthetic replacement of three coronary arteries
39	7923300	Read	prosthetic replacement of four or more coronary arteries
40	7923z00	Read	prosthetic replacement of coronary artery nos
41	7924	Read	revision of bypass for coronary artery
42	7924000	Read	revision of bypass for one coronary artery
43	7924100	Read	revision of bypass for two coronary arteries
44	7924200	Read	revision of bypass for three coronary arteries
45	7924y00	Read	other specified revision of bypass for coronary artery
46	7924z00	Read	revision of bypass for coronary artery nos

1			
2	7925	Read	connection of mammary artery to coronary artery
3	7925000	Read	double anastomosis of mammary arteries to coronary arteries
4	7925100	Read	double implant of mammary arteries into coronary arteries
5	7925.11.00	Read	creation of bypass from mammary artery to coronary artery
6	7925300	Read	single anastomosis of mammary artery to coronary artery nec
7	7925311	Read	lima single anastomosis
8	7925312	Read	rima single anastomosis
9	7925400	Read	single implantation of mammary artery into coronary artery
10	7925y00	Read	connection of mammary artery to coronary artery os
11	7925z00	Read	connection of mammary artery to coronary artery nos
12	7926	Read	connection of other thoracic artery to coronary artery
13	7926000	Read	double anastom thoracic arteries to coronary arteries nec
14	7926200	Read	single anastomosis of thoracic artery to coronary artery nec
15	7926300	Read	single implantation thoracic artery into coronary artery nec
16	7926z00	Read	connection of other thoracic artery to coronary artery nos
17	7927500	Read	open angioplasty of coronary artery
18	7928	Read	transluminal balloon angioplasty of coronary artery
19	7928000	Read	percut transluminal balloon angioplasty one coronary artery
20	7928100	Read	percut translum balloon angioplasty mult coronary arteries
21	7928.11.00	Read	percutaneous balloon coronary angioplasty
22	7928200	Read	percut translum balloon angioplasty bypass graft coronary a
23	7928300	Read	percut translum cutting balloon angioplasty coronary artery
24	7928y00	Read	transluminal balloon angioplasty of coronary artery os
25	7928z00	Read	transluminal balloon angioplasty of coronary artery nos
26	7929000	Read	percutaneous transluminal laser coronary angioplasty
27	7929100	Read	percut transluminal coronary thrombolysis with streptokinase
28	7929111	Read	percut translum coronary thrombolytic therapy- streptokinase
29	7929300	Read	rotary blade coronary angioplasty
30	7929400	Read	insertion of coronary artery stent
31	7929500	Read	insertion of drug-eluting coronary artery stent
32	7929600	Read	percutaneous transluminal atherectomy of coronary artery
33	792B000	Read	endarterectomy of coronary artery nec
34	792C.00	Read	other replacement of coronary artery
35	792C000	Read	replacement of coronary arteries using multiple methods
36	792Cy00	Read	other specified replacement of coronary artery
37	792Cz00	Read	replacement of coronary artery nos
38	792D.00	Read	other bypass of coronary artery
39	792Dy00	Read	other specified other bypass of coronary artery
40	792Dz00	Read	other bypass of coronary artery nos
41	793G.00	Read	perc translumin balloon angioplasty stenting coronary artery
42	793G000	Read	perc translum ball angio insert 1-2 drug elut stents cor art
43	793G100	Read	perc tran ball angio ins 3 or more drug elut stents cor art
44	793G200	Read	perc translum balloon angioplasty insert 1-2 stents cor art
45	793G300	Read	percutaneous cor balloon angiop 3 more stents cor art nec
46	793Gy00	Read	os perc translumina balloon angioplast stenting coronary art

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1			
2	793Gz00	Read	perc translum balloon angioplasty stenting coronary art nos
3	7A4B800	Read	percut translum thrombolysis femoral graft streptokinase
4	7A54000	Read	percutaneous transluminal angioplasty of artery nec
5	7A54500	Read	rotary blade angioplasty
6	7A54700	Read	percutaneous transluminal thrombolysis of artery
7	7A54800	Read	percutaneous transluminal atherectomy
8	7A56000	Read	percutaneous transluminal arterial thrombolysis reconstruct
9	7A56400	Read	percutaneous transluminal balloon angioplasty of artery
10	7A6G100	Read	peroperative angioplasty
11	7A6H300	Read	prosthetic graft patch angioplasty
12	7A6H400	Read	percutaneous transluminal angioplasty of vascular graft
13	7A6S300	Read	percutaneous transluminal venous thrombolysis nec
14	889A.00	Read	diab mellit insulin-glucose infus acute myocardial infarct
15	8B27.00	Read	antianginal therapy
16	8B3k.00	Read	coronary heart disease medication review
17	8BGC.00	Read	long term dual antiplatelet drug therapy indicated
18	8CMP.00	Read	coronary heart disease care plan
19	8F9..00	Read	Cardiac rehabilitation
20	8F90.00	Read	Cardiac rehabilitation - phase 1
21	8F91.00	Read	Cardiac rehabilitation - phase 2
22	8F92.00	Read	Cardiac rehabilitation - phase 3
23	8F93.00	Read	Cardiac rehabilitation - phase 4
24	8H2V.00	Read	admit ischaemic heart disease emergency
25	8H7v.00	Read	Referral to cardiac rehabilitation nurse
26	8I37.00	Read	Coronary heart disease monitoring refused
27	8I3a.00	Read	Cardiac rehabilitation declined
28	8IEY.00	Read	referral to angina plan self-management programme declined
29	8L40.00	Read	coronary artery bypass graft operation planned
30	8L41.00	Read	coronary angioplasty planned
31	8LF..00	Read	coronary angiography planned
32	8T04.00	Read	referral to angina plan self-management programme
33	9Ob..00	Read	Coronary heart disease monitoring administration
34	9Ob0.00	Read	Attends coronary heart disease monitoring
35	9Ob1.00	Read	Refuses coronary heart disease monitoring
36	9Ob2.00	Read	Coronary heart disease monitoring default
37	9Ob3.00	Read	Coronary heart disease monitoring 1st letter
38	9Ob4.00	Read	Coronary heart disease monitoring 2nd letter
39	9Ob5.00	Read	Coronary heart disease monitoring 3rd letter
40	9Ob6.00	Read	Coronary heart disease monitoring verbal invitation
41	9Ob8.00	Read	Coronary heart disease monitoring check done
42	9Ob9.00	Read	Coronary heart disease monitoring telephone invite
43	G....12	Read	Cardiac diseases
44	G....13	Read	Heart diseases
45	G3...00	Read	ischaemic heart disease
46	G30..00	Read	acute myocardial infarction

1			
2	G300.00	Read	acute anterolateral infarction
3	G301.00	Read	other specified anterior myocardial infarction
4	G301000	Read	acute anteroapical infarction
5	G30..11	Read	attack - heart
6	G301100	Read	acute anteroseptal infarction
7	G30..12	Read	coronary thrombosis
8	G30..13	Read	cardiac rupture following myocardial infarction (mi)
9	G30..14	Read	heart attack
10	G30..15	Read	mi - acute myocardial infarction
11	G30..16	Read	thrombosis - coronary
12	G30..17	Read	silent myocardial infarction
13	G301z00	Read	anterior myocardial infarction nos
14	G302.00	Read	acute inferolateral infarction
15	G303.00	Read	acute inferoposterior infarction
16	G304.00	Read	posterior myocardial infarction nos
17	G305.00	Read	lateral myocardial infarction nos
18	G306.00	Read	true posterior myocardial infarction
19	G307.00	Read	acute subendocardial infarction
20	G307000	Read	acute non-q wave infarction
21	G307100	Read	acute non-st segment elevation myocardial infarction
22	G308.00	Read	inferior myocardial infarction nos
23	G309.00	Read	acute q-wave infarct
24	G30A.00	Read	mural thrombosis
25	G30B.00	Read	acute posterolateral myocardial infarction
26	G30X.00	Read	acute transmural myocardial infarction of unspecif site
27	G30X000	Read	acute st segment elevation myocardial infarction
28	G30y.00	Read	other acute myocardial infarction
29	G30y000	Read	acute atrial infarction
30	G30y100	Read	acute papillary muscle infarction
31	G30y200	Read	acute septal infarction
32	G30yz00	Read	other acute myocardial infarction nos
33	G30z.00	Read	acute myocardial infarction nos
34	G31..00	Read	other acute and subacute ischaemic heart disease
35	G310.00	Read	postmyocardial infarction syndrome
36	G310.11	Read	dressler's syndrome
37	G3...11	Read	arteriosclerotic heart disease
38	G311.00	Read	preinfarction syndrome
39	G311000	Read	myocardial infarction aborted
40	G311011	Read	mi - myocardial infarction aborted
41	G311100	Read	unstable angina
42	G311.11	Read	crescendo angina
43	G311.12	Read	impending infarction
44	G311.13	Read	unstable angina
45	G311.14	Read	angina at rest
46	G311200	Read	angina at rest

1			
2	G311300	Read	refractory angina
3	G311400	Read	worsening angina
4	G311500	Read	acute coronary syndrome
5	G311z00	Read	preinfarction syndrome nos
6	G3...12	Read	atherosclerotic heart disease
7	G312.00	Read	coronary thrombosis not resulting in myocardial infarction
8	G3...13	Read	ihd - ischaemic heart disease
9	G31y.00	Read	other acute and subacute ischaemic heart disease
10	G31y000	Read	acute coronary insufficiency
11	G31y100	Read	microinfarction of heart
12	G31y200	Read	subendocardial ischaemia
13	G31y300	Read	transient myocardial ischaemia
14	G31yz00	Read	other acute and subacute ischaemic heart disease nos
15	G32..00	Read	old myocardial infarction
16	G32..11	Read	healed myocardial infarction
17	G32..12	Read	personal history of myocardial infarction
18	G33..00	Read	angina pectoris
19	G330.00	Read	angina decubitus
20	G330000	Read	nocturnal angina
21	G330z00	Read	angina decubitus nos
22	G331.11	Read	variant angina pectoris
23	G332.00	Read	coronary artery spasm
24	G33z.00	Read	angina pectoris nos
25	G33z000	Read	status anginosus
26	G33z100	Read	stenocardia
27	G33z200	Read	syncope anginosa
28	G33z300	Read	angina on effort
29	G33z400	Read	ischaemic chest pain
30	G33z500	Read	post infarct angina
31	G33z600	Read	new onset angina
32	G33z700	Read	stable angina
33	G33zz00	Read	angina pectoris nos
34	G34..00	Read	other chronic ischaemic heart disease
35	G340.00	Read	coronary atherosclerosis
36	G340000	Read	single coronary vessel disease
37	G340100	Read	double coronary vessel disease
38	G340.11	Read	triple vessel disease of the heart
39	G340.12	Read	coronary artery disease
40	G341.00	Read	Aneurysm of heart
41	G341000	Read	Ventricular cardiac aneurysm
42	G341100	Read	Other cardiac wall aneurysm
43	G341.11	Read	Cardiac aneurysm
44	G341z00	Read	Aneurysm of heart NOS
45	G342.00	Read	atherosclerotic cardiovascular disease
46	G343.00	Read	Ischaemic cardiomyopathy

1			
2	G344.00	Read	silent myocardial ischaemia
3	G34y.00	Read	other specified chronic ischaemic heart disease
4	G34y000	Read	chronic coronary insufficiency
5	G34y100	Read	chronic myocardial ischaemia
6	G34yz00	Read	other specified chronic ischaemic heart disease nos
7	G34z.00	Read	other chronic ischaemic heart disease nos
8	G34z000	Read	asymptomatic coronary heart disease
9	G35..00	Read	subsequent myocardial infarction
10	G350.00	Read	subsequent myocardial infarction of anterior wall
11	G351.00	Read	subsequent myocardial infarction of inferior wall
12	G353.00	Read	subsequent myocardial infarction of other sites
13	G35X.00	Read	subsequent myocardial infarction of unspecified site
14	G36..00	Read	certain current complication follow acute myocardial infarct
15	G360.00	Read	haemopericardium/current comp folow acute myocardial infarct
16	G361.00	Read	atrial septal defect/curr comp folow acute myocardial infarct
17	G362.00	Read	ventricular septal defect/curr comp folow acute myocardial infarct
18	G363.00	Read	rupture cardiac wall w/out haemopericard/curr comp folow acute myocardial infarct
19	G364.00	Read	rupture chordae tendinae/curr comp folow acute myocardial infarct
20	G365.00	Read	rupture papillary muscle/curr comp folow acute myocardial infarct
21	G366.00	Read	thrombosis atrium
22	G38..00	Read	postoperative myocardial infarction
23	G380.00	Read	postoperative transmural myocardial infarction anterior wall
24	G381.00	Read	postoperative transmural myocardial infarction inferior wall
25	G383.00	Read	postoperative transmural myocardial infarction unspecified site
26	G384.00	Read	postoperative subendocardial myocardial infarction
27	G38z.00	Read	postoperative myocardial infarction
28	G39..00	Read	coronary microvascular disease
29	G3y..00	Read	other specified ischaemic heart disease
30	G3z..00	Read	ischaemic heart disease nos
31	G5...00	Read	other forms of heart disease
32	G501.00	Read	post infarction pericarditis
33	G574000	Read	Ventricular fibrillation
34	G5y..00	Read	other specified heart disease
35	G5yyz00	Read	Other ill-defined heart disease NOS
36	G5yz.00	Read	other heart disease nos
37	G5z..00	Read	heart disease nos
38	Gyu3.00	Read	[x]ischaemic heart diseases
39	Gyu3000	Read	[x]other forms of angina pectoris
40	Gyu3200	Read	[x]other forms of acute ischaemic heart disease
41	Gyu3300	Read	[x]other forms of chronic ischaemic heart disease
42	Gyu3400	Read	[x]acute transmural myocardial infarction of unspecified site
43	Gyu3600	Read	[x]subsequent myocardial infarction of unspecified site
44	Gyu5.00	Read	[x]other forms of heart disease
45	Gyu7000	Read	[x]atherosclerosis of other arteries
46	SP00300	Read	mechanical complication of coronary bypass

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SP07600	Read	coronary artery bypass graft occlusion
Z677.00	Read	Cardiac rehabilitation class
ZL22200	Read	Under care of cardiac rehabilitation nurse
ZV45700	Read	[v]presence of aortocoronary bypass graft
ZV45800	Read	[v]presence of coronary angioplasty implant and graft
ZV45K00	Read	[v]presence of coronary artery bypass graft
ZV45K11	Read	[v]presence of coronary artery bypass graft - cabg
ZV45L00	Read	[v]status following coronary angioplasty nos
ZV57900	Read	[V]Cardiac rehabilitation
I20	ICD-10	angina pectoris
I20.0	ICD-10	unstable angina
I20.1	ICD-10	angina pectoris with documented spasm
I20.8	ICD-10	other forms of angina pectoris
I20.9	ICD-10	angina pectoris
I21	ICD-10	acute myocardial infarction
I21.0	ICD-10	acute transmural myocardial infarction of anterior wall
I21.1	ICD-10	acute transmural myocardial infarction of inferior wall
I21.2	ICD-10	acute transmural myocardial infarction of other sites
I21.3	ICD-10	acute transmural myocardial infarction of unspecified site
I21.4	ICD-10	acute subendocardial myocardial infarction
I21.9	ICD-10	acute myocardial infarction
I22	ICD-10	subsequent myocardial infarction
I22.0	ICD-10	subsequent myocardial infarction of anterior wall
I22.1	ICD-10	subsequent myocardial infarction of inferior wall
I22.8	ICD-10	subsequent myocardial infarction of other sites
I22.9	ICD-10	subsequent myocardial infarction of unspecified site
I23	ICD-10	certain current complications following acute myocardial infarction
I23.0	ICD-10	haemopericardium as current complication following acute myocardial infarction
I23.1	ICD-10	atrial septal defect as current complication following acute myocardial infarction
I23.2	ICD-10	ventricular septal defect as current complication following acute myocardial infarction
I23.3	ICD-10	rupture of cardiac wall without haemopericardium as current complication following acute myocardial
I23.4	ICD-10	rupture of chordae tendineae as current complication following acute myocardial infarction
I23.5	ICD-10	rupture of papillary muscle as current complication following acute myocardial infarction
I23.6	ICD-10	thrombosis of atrium
I23.8	ICD-10	other current complications following acute myocardial infarction
I24	ICD-10	other acute ischaemic heart diseases
I24.0	ICD-10	coronary thrombosis not resulting in myocardial infarction
I24.1	ICD-10	dressler's syndrome
I24.8	ICD-10	other forms of acute ischaemic heart disease
I24.9	ICD-10	acute ischaemic heart disease
I25	ICD-10	chronic ischaemic heart disease
I25.0	ICD-10	atherosclerotic cardiovascular disease
I25.1	ICD-10	atherosclerotic heart disease
I25.2	ICD-10	old myocardial infarction
I25.5	ICD-10	ischaemic cardiomyopathy

1			
2	I25.6	ICD-10	silent myocardial ischaemia
3	I25.8	ICD-10	other forms of chronic ischaemic heart disease
4	I25.9	ICD-10	chronic ischaemic heart disease
5			
6	T82.2	ICD-10	mechanical complication of coronary artery bypass and valve grafts
7			
8	Z95.5	ICD-10	presence of coronary angioplasty implant and graft
9			

Heart failure

Code	Classification	Description
14A6.00	Read	h/o: heart failure
14AM.00	Read	h/o: heart failure in last year
1736	Read	paroxysmal nocturnal dyspnoea
1J60.00	Read	suspected heart failure
1O1..00	Read	heart failure confirmed
388D.00	Read	new york heart assoc classification heart failure symptoms
585f.00	Read	echocardiogram shows left ventricular systolic dysfunction
585g.00	Read	echocardiogram shows left ventricular diastolic dysfunction
661M500	Read	heart failure self-management plan agreed
662f.00	Read	new york heart association classification - class i
662g.00	Read	new york heart association classification - class ii
662h.00	Read	new york heart association classification - class iii
662i.00	Read	new york heart association classification - class iv
662p.00	Read	heart failure 6 month review
662T.00	Read	congestive heart failure monitoring
662W.00	Read	heart failure annual review
679W100	Read	education about deteriorating heart failure
679X.00	Read	heart failure education
67D4.00	Read	heart failure information given to patient
8B29.00	Read	cardiac failure therapy
8CeC.00	Read	preferred place of care for next exacerbation heart failure
8CL3.00	Read	heart failure care plan discussed with patient
8CMK.00	Read	has heart failure management plan
8CMW800	Read	heart failure clinical pathway
8H2S.00	Read	admit heart failure emergency
8HBE.00	Read	heart failure follow-up
8Hg8.00	Read	discharge from practice nurse heart failure clinic
8HgD.00	Read	discharge from heart failure nurse service
8HHb.00	Read	referral to heart failure nurse
8HHz.00	Read	referral to heart failure exercise programme
8Hk0.00	Read	referred to heart failure education group
8HTL.00	Read	referral to heart failure clinic
8HTL000	Read	referral to rapid access heart failure clinic
8IE0.00	Read	referral to heart failure education group declined
8IE1.00	Read	referral to heart failure exercise programme declined
9h1..00	Read	exception reporting: lvd quality indicators

1			
2	9h11.00	Read	excepted from lvd quality indicators: patient unsuitable
3	9h12.00	Read	excepted from lvd quality indicators: informed dissent
4	9hH..00	Read	exception reporting: heart failure quality indicators
5	9hH0.00	Read	excepted heart failure quality indicators: patient unsuitable
6	9hH1.00	Read	excepted heart failure quality indicators: informed dissent
7	9N0k.00	Read	seen in heart failure clinic
8	9N2p.00	Read	seen by community heart failure nurse
9	9N4s.00	Read	did not attend practice nurse heart failure clinic
10	9N4w.00	Read	did not attend heart failure clinic
11	9N6T.00	Read	referred by heart failure nurse specialist
12	9On..00	Read	left ventricular dysfunction monitoring administration
13	9On0.00	Read	left ventricular dysfunction monitoring first letter
14	9On1.00	Read	left ventricular dysfunction monitoring second letter
15	9On2.00	Read	left ventricular dysfunction monitoring third letter
16	9On3.00	Read	left ventricular dysfunction monitoring verbal invite
17	9On4.00	Read	left ventricular dysfunction monitoring telephone invite
18	9Or..00	Read	heart failure monitoring administration
19	9Or0.00	Read	heart failure review completed
20	9Or1.00	Read	heart failure monitoring telephone invite
21	9Or2.00	Read	heart failure monitoring verbal invite
22	9Or3.00	Read	heart failure monitoring first letter
23	9Or4.00	Read	heart failure monitoring second letter
24	9Or5.00	Read	heart failure monitoring third letter
25	G1yz100	Read	rheumatic left ventricular failure
26	G210.00	Read	malignant hypertensive heart disease
27	G210100	Read	malignant hypertensive heart disease with ccf
28	G211100	Read	benign hypertensive heart disease with ccf
29	G21z100	Read	hypertensive heart disease nos with ccf
30	G230.00	Read	malignant hypertensive heart and renal disease
31	G232.00	Read	hypertensive heart&renal dis with (congestive) heart failure
32	G234.00	Read	hyperten heart&renal dis+both(congestv)heart and renal fail
33	G400.00	Read	acute cor pulmonale
34	G41z.11	Read	chronic cor pulmonale
35	G554000	Read	congestive cardiomyopathy
36	G554011	Read	congestive obstructive cardiomyopathy
37	G557100	Read	beriberi heart disease
38	G58..00	Read	heart failure
39	G580.00	Read	congestive heart failure
40	G580000	Read	acute congestive heart failure
41	G580100	Read	chronic congestive heart failure
42	G580.11	Read	congestive cardiac failure
43	G580.12	Read	right heart failure
44	G580.13	Read	right ventricular failure
45	G580.14	Read	biventricular failure
46	G580200	Read	decompensated cardiac failure

1			
2	G580300	Read	compensated cardiac failure
3	G580400	Read	congestive heart failure due to valvular disease
4	G581.00	Read	left ventricular failure
5	G581000	Read	acute left ventricular failure
6	G58..11	Read	cardiac failure
7	G581.11	Read	asthma - cardiac
8	G581.13	Read	impaired left ventricular function
9	G582.00	Read	acute heart failure
10	G583.00	Read	heart failure with normal ejection fraction
11	G583.11	Read	hfnef - heart failure with normal ejection fraction
12	G583.12	Read	heart failure with preserved ejection fraction
13	G584.00	Read	right ventricular failure
14	G58z.00	Read	heart failure nos
15	G58z.11	Read	weak heart
16	G58z.12	Read	cardiac failure nos
17	G5y4z00	Read	post cardiac operation heart failure nos
18	G5yy900	Read	left ventricular systolic dysfunction
19	G5yyA00	Read	left ventricular diastolic dysfunction
20	G5yyB00	Read	right ventricular diastolic dysfunction
21	Q48y100	Read	congenital cardiac failure
22	R2y1000	Read	[d]cardiorespiratory failure
23	SP11111	Read	heart failure as a complication of care
24	ZRad.00	Read	new york heart assoc classification heart failure symptoms
25	I11.0	ICD-10	hypertensive heart disease with (congestive) heart failure
26	I13.0	ICD-10	hypertensive heart and renal disease with (congestive) heart failure
27	I13.2	ICD-10	hypertensive heart and renal disease with both (congestive) heart failure and renal failure
28	I26.0	ICD-10	pulmonary embolism with mention of acute cor pulmonale
29	I50	ICD-10	heart failure
30	I50.0	ICD-10	congestive heart failure
31	I50.1	ICD-10	left ventricular failure
32	I50.9	ICD-10	heart failure

Hypertensi
on

Code	Classification	Description
14A2.00	Read	h/o: hypertension
1JD..00	Read	suspected hypertension
2126100	Read	hypertension resolved
212K.00	Read	hypertension resolved
246M.00	Read	white coat hypertension
662..12	Read	hypertension monitoring
6624	Read	borderline hyperten:yearly obs
6627	Read	good hypertension control
6628	Read	poor hypertension control
6629	Read	hypertension:follow-up default

1			
2	662b.00	Read	moderate hypertension control
3	662c.00	Read	hypertension six month review
4	662d.00	Read	hypertension annual review
5	662F.00	Read	hypertension treatm. started
6	662G.00	Read	hypertensive treatm.changed
7	662H.00	Read	hypertension treatm.stopped
8			
9	662O.00	Read	on treatment for hypertension
10	662P.00	Read	hypertension monitoring
11	662P000	Read	hypertension 9 month review
12	662q.00	Read	trial reduction of antihypertensive therapy
13	662r.00	Read	trial withdrawal of antihypertensive therapy
14	67H8.00	Read	lifestyle advice regarding hypertension
15	7Q01.00	Read	high cost hypertension drugs
16	7Q01y00	Read	other specified high cost hypertension drugs
17	8B26.00	Read	antihypertensive therapy
18	8BL0.00	Read	patient on maximal tolerated antihypertensive therapy
19	8CR4.00	Read	hypertension clinical management plan
20	8HT5.00	Read	referral to hypertension clinic
21	8I3N.00	Read	hypertension treatment refused
22	9h3..00	Read	exception reporting: hypertension quality indicators
23	9h31.00	Read	excepted from hypertension qual indicators: patient unsuit
24	9h32.00	Read	excepted from hypertension qual indicators: informed dissent
25	9N03.00	Read	seen in hypertension clinic
26	9N1y200	Read	seen in hypertension clinic
27	9N4L.00	Read	dna - did not attend hypertension clinic
28	9OI..00	Read	hypertension monitoring admin.
29	9OI1.00	Read	attends hypertension monitor.
30	9OI..11	Read	hypertension clinic admin.
31	9OI2.00	Read	refuses hypertension monitor.
32	9OI3.00	Read	hyperten.monitor offer default
33	9OI4.00	Read	hypertens.monitor.1st letter
34	9OI5.00	Read	hypertens.monitor 2nd letter
35	9OI6.00	Read	hypertens.monitor 3rd letter
36	9OI7.00	Read	hypertens.monitor verbal inv.
37	9OI8.00	Read	hypertens.monitor phone invite
38	9OI9.00	Read	hypertens.monitor deleted
39	9OIA.00	Read	hypertension monitor.chk done
40	9OIA.11	Read	hypertension monitored
41	9OIZ.00	Read	hypertens.monitoring admin.nos
42	F404200	Read	blind hypertensive eye
43	F421300	Read	hypertensive retinopathy
44	G2...00	Read	hypertensive disease
45	G20..00	Read	essential hypertension
46	G200.00	Read	malignant essential hypertension
47	G201.00	Read	benign essential hypertension

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2	G20..11	Read	high blood pressure
3	G20..12	Read	primary hypertension
4	G202.00	Read	systolic hypertension
5	G203.00	Read	diastolic hypertension
6	G20z.00	Read	essential hypertension nos
7	G20z.11	Read	hypertension nos
8	G21..00	Read	hypertensive heart disease
9	G210.00	Read	malignant hypertensive heart disease
10	G210000	Read	malignant hypertensive heart disease without ccf
11	G210100	Read	malignant hypertensive heart disease with ccf
12	G210z00	Read	malignant hypertensive heart disease nos
13	G2...11	Read	bp - hypertensive disease
14	G211.00	Read	benign hypertensive heart disease
15	G211000	Read	benign hypertensive heart disease without ccf
16	G211100	Read	benign hypertensive heart disease with ccf
17	G211z00	Read	benign hypertensive heart disease nos
18	G21z.00	Read	hypertensive heart disease nos
19	G21z000	Read	hypertensive heart disease nos without ccf
20	G21z011	Read	cardiomegaly - hypertensive
21	G21z100	Read	hypertensive heart disease nos with ccf
22	G21zz00	Read	hypertensive heart disease nos
23	G22..00	Read	hypertensive renal disease
24	G220.00	Read	malignant hypertensive renal disease
25	G221.00	Read	benign hypertensive renal disease
26	G22..11	Read	nephrosclerosis
27	G222.00	Read	hypertensive renal disease with renal failure
28	G22z.00	Read	hypertensive renal disease nos
29	G22z.11	Read	renal hypertension
30	G23..00	Read	hypertensive heart and renal disease
31	G230.00	Read	malignant hypertensive heart and renal disease
32	G231.00	Read	benign hypertensive heart and renal disease
33	G232.00	Read	hypertensive heart&renal dis wth (congestive) heart failure
34	G233.00	Read	hypertensive heart and renal disease with renal failure
35	G234.00	Read	hyperten heart&renal dis+both(congestv)heart and renal fail
36	G23z.00	Read	hypertensive heart and renal disease nos
37	G24..00	Read	secondary hypertension
38	G240.00	Read	secondary malignant hypertension
39	G240000	Read	secondary malignant renovascular hypertension
40	G240z00	Read	secondary malignant hypertension nos
41	G241.00	Read	secondary benign hypertension
42	G241000	Read	secondary benign renovascular hypertension
43	G241z00	Read	secondary benign hypertension nos
44	G244.00	Read	hypertension secondary to endocrine disorders
45	G24z.00	Read	secondary hypertension nos
46	G24z000	Read	secondary renovascular hypertension nos

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1			
2	G24z100	Read	hypertension secondary to drug
3	G24zz00	Read	secondary hypertension nos
4	G25..00	Read	stage 1 hypertension (nice - nat ins for hth clin excl 2011)
5	G25..11	Read	stage 1 hypertension
6	G26..00	Read	severe hypertension (nat inst for health clinical ex 2011)
7	G26..11	Read	severe hypertension
8			
9	G27..00	Read	hypertension resistant to drug therapy
10	G28..00	Read	stage 2 hypertension (nice - nat ins for hth clin excl 2011)
11	G2y..00	Read	other specified hypertensive disease
12	G2z..00	Read	hypertensive disease nos
13	G672.00	Read	hypertensive encephalopathy
14	G672.11	Read	hypertensive crisis
15	Gyu2.00	Read	[x]hypertensive diseases
16	Gyu2000	Read	[x]other secondary hypertension
17	Gyu2100	Read	[x]hypertension secondary to other renal disorders
18	L122.00	Read	other pre-existing hypertension in preg/childbirth/puerp
19	L122000	Read	other pre-existing hypertension in preg/childb/puerp unsp
20	L122100	Read	other pre-existing hypertension in preg/childb/puerp - deliv
21	L122300	Read	other pre-exist hypertension in preg/childb/puerp-not deliv
22	L122z00	Read	other pre-existing hypertension in preg/childb/puerp nos
23	L127.00	Read	pre-eclampsia or eclampsia with pre-existing hypertension
24	L127z00	Read	pre-eclampsia or eclampsia + pre-existing hypertension nos
25	L128.00	Read	pre-exist hypertension compl preg childbirth and puerperium
26	L128000	Read	pre-exist hyperten heart dis compl preg childbth+puerperium
27	L128200	Read	pre-exist 2ndry hypertens comp preg childbth and puerperium
28	TJC7.00	Read	adverse reaction to other antihypertensives
29	TJC7z00	Read	adverse reaction to antihypertensives nos
30	U60C500	Read	[x]oth antihyperten drug caus advers eff in therap use
31	U60C511	Read	[x] adverse reaction to other antihypertensives
32	U60C51A	Read	[x] adverse reaction to antihypertensives nos
33	I10	ICD-10	essential (primary) hypertension
34	I11	ICD-10	hypertensive heart disease
35	I11.0	ICD-10	hypertensive heart disease with (congestive) heart failure
36	I11.9	ICD-10	hypertensive heart disease without (congestive) heart failure
37	I12	ICD-10	hypertensive renal disease
38	I12.0	ICD-10	hypertensive renal disease with renal failure
39	I12.9	ICD-10	hypertensive renal disease without renal failure
40	I13	ICD-10	hypertensive heart and renal disease
41	I13.0	ICD-10	hypertensive heart and renal disease with (congestive) heart failure
42	I13.1	ICD-10	hypertensive heart and renal disease with renal failure
43	I13.2	ICD-10	hypertensive heart and renal disease with both (congestive) heart failure and renal failure
44	I13.9	ICD-10	hypertensive heart and renal disease
45	I15	ICD-10	secondary hypertension
46	I15.0	ICD-10	renovascular hypertension
47	I15.1	ICD-10	hypertension secondary to other renal disorders

1			
2	I15.2	ICD-10	hypertension secondary to endocrine disorders
3	I15.8	ICD-10	other secondary hypertension
4	I15.9	ICD-10	secondary hypertension
5			
6	I67.4	ICD-10	hypertensive encephalopathy
7			
8			

9 **Arrhythmia**

10	Code	Classification	Description
11	14AN.00	Read	h/o: atrial fibrillation
12	14AR.00	Read	history of atrial flutter
13	212R.00	Read	atrial fibrillation resolved
14	327..00	Read	ecg: supraventricular arrhythmia
15	3272	Read	ecg: atrial fibrillation
16	3273	Read	ecg: atrial flutter
17	328..00	Read	ecg: ventricular arrhythmia
18	3282	Read	ecg: ventricular tachycardia
19	328Z.00	Read	ecg: ventricular arrhythmia nos
20	662S.00	Read	atrial fibrillation monitoring
21	6A9..00	Read	atrial fibrillation annual review
22	7936A00	Read	implant intravenous pacemaker for atrial fibrillation
23	8CMW200	Read	atrial fibrillation care pathway
24	8HTy.00	Read	referral to atrial fibrillation clinic
25	9hF..00	Read	exception reporting: atrial fibrillation quality indicators
26	9hF1.00	Read	excepted from atrial fibrillation qual indic: inform dissent
27	9Os..00	Read	atrial fibrillation monitoring administration
28	9Os0.00	Read	atrial fibrillation monitoring first letter
29	9Os1.00	Read	atrial fibrillation monitoring second letter
30	9Os2.00	Read	atrial fibrillation monitoring third letter
31	9Os3.00	Read	atrial fibrillation monitoring verbal invite
32	9Os4.00	Read	atrial fibrillation monitoring telephone invite
33	G559.00	Read	arrhythmogenic right ventricular cardiomyopathy
34	G55A.11	Read	tachycardia-induced cardiomyopathy
35	G56..00	Read	conduction disorders
36	G56..11	Read	conduction disorders of heart
37	G567400	Read	wolff-parkinson-white syndrome
38	G56y.00	Read	other conduction disorders
39	G56y000	Read	lown-ganong-levine syndrome
40	G56zz00	Read	conduction disorders nos
41	G57..00	Read	cardiac dysrhythmias
42	G570.00	Read	paroxysmal supraventricular tachycardia
43	G570000	Read	paroxysmal atrial tachycardia
44	G570100	Read	paroxysmal atrioventricular tachycardia
45	G570200	Read	paroxysmal junctional tachycardia
46	G570300	Read	paroxysmal nodal tachycardia
47	G570z00	Read	paroxysmal supraventricular tachycardia nos
48	G571.00	Read	paroxysmal ventricular tachycardia

G57..11	Read	cardiac arrhythmias
G571.11	Read	ventricular tachycardia
G572.00	Read	paroxysmal tachycardia unspecified
G572000	Read	essential paroxysmal tachycardia
G572z00	Read	paroxysmal tachycardia nos
G573.00	Read	atrial fibrillation and flutter
G573000	Read	atrial fibrillation
G573100	Read	atrial flutter
G573200	Read	paroxysmal atrial fibrillation
G573300	Read	non-rheumatic atrial fibrillation
G573400	Read	permanent atrial fibrillation
G573500	Read	persistent atrial fibrillation
G573600	Read	paroxysmal atrial flutter
G573z00	Read	atrial fibrillation and flutter nos
G574.00	Read	ventricular fibrillation and flutter
G574100	Read	ventricular flutter
G574z00	Read	ventricular fibrillation and flutter nos
G576300	Read	atrial premature depolarization
G576400	Read	junctional premature depolarization
G576500	Read	ventricular premature depolarization
G57y.00	Read	other cardiac dysrhythmias
G57y600	Read	nodal rhythm disorder
G57y900	Read	supraventricular tachycardia nos
G57yA00	Read	re-entry ventricular arrhythmia
G57yz00	Read	other cardiac dysrhythmia nos
G57z.00	Read	cardiac dysrhythmia nos
Gyu5a00	Read	[x]other specified cardiac arrhythmias
I45.6	ICD-10	pre-excitation syndrome
I47	ICD-10	paroxysmal tachycardia
I47.0	ICD-10	re-entry ventricular arrhythmia
I47.1	ICD-10	supraventricular tachycardia
I47.2	ICD-10	ventricular tachycardia
I47.9	ICD-10	paroxysmal tachycardia
I48	ICD-10	atrial fibrillation and flutter
I49	ICD-10	other cardiac arrhythmias
I49.0	ICD-10	ventricular fibrillation and flutter
I49.1	ICD-10	atrial premature depolarization
I49.2	ICD-10	junctional premature depolarization
I49.3	ICD-10	ventricular premature depolarization
I49.4	ICD-10	other and unspecified premature depolarization
I49.5	ICD-10	sick sinus syndrome
R00.0	ICD-10	tachycardia

Peripheral arterial disease

Code Classification Description

1			
2	G73z000	Read	intermittent claudication
3	G73z011	Read	claudication
4	G73..12	Read	ischaemia of legs
5	G73zz00	Read	peripheral vascular disease nos
6	G73z.00	Read	peripheral vascular disease nos
7	G73yz00	Read	other specified peripheral vascular disease nos
8	G73..11	Read	peripheral ischaemic vascular disease
9	G73..00	Read	other peripheral vascular disease
10	G73..13	Read	peripheral ischaemia
11	2G63.00	Read	ischaemic toe
12	G702.00	Read	Extremity artery atheroma
13	G742z00	Read	peripheral arterial embolism and thrombosis nos
14	G702z00	Read	Extremity artery atheroma NOS
15	G76A.00	Read	arterial insufficiency
16	G73y100	Read	Peripheral angiopathic disease EC NOS
17	R055011	Read	[d]peripheral circulatory failure
18	G73y.00	Read	other specified peripheral vascular disease
19	14NB.00	Read	h/o: peripheral vascular disease procedure
20	Gyu7400	Read	[x]other specified peripheral vascular diseases
21	7A56600	Read	percutaneous transluminal placement peripheral stent artery
22	G733.00	Read	ischaemic foot
23	G73z012	Read	vascular claudication
24	G734.00	Read	peripheral arterial disease
25	16I..00	Read	claudication distance
26	I70.2	ICD-10	atherosclerosis of arteries of extremities
27	I70.20	ICD-10	atherosclerosis of arteries of extremities
28	I70.21	ICD-10	atherosclerosis of arteries of extremities
29	I73	ICD-10	other peripheral vascular diseases
30	I73.8	ICD-10	other specified peripheral vascular diseases
31	I73.9	ICD-10	peripheral vascular disease, unspecified
32	I79.2	ICD-10	peripheral angiopathy in diseases classified elsewhere

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46
Myocardial infarction

Code	Classification	Description
47		
48		
49	14A3.00	Read H/O: myocardial infarct <60
50	14A4.00	Read H/O: myocardial infarct >60
51	14AH.00	Read H/O: Myocardial infarction in last year
52	323..00	Read ECG: myocardial infarction
53	3232.00.00	Read ECG: old myocardial infarction
54	3233.00.00	Read ECG: antero-septal infarct.
55	3234.00.00	Read ECG:posterior/inferior infarct
56	3235.00.00	Read ECG: subendocardial infarct
57	3236.00.00	Read ECG: lateral infarction
58	323Z.00	Read ECG: myocardial infarct NOS
59	32B2.00	Read ECG: Q wave abnormal
60		

32B3.00	Read	ECG: Q wave pathological
32E3.00	Read	ECG: S-T elevation
44H3.00	Read	Cardiac enzymes abnormal
44H3000	Read	Cardiac enzymes abnormal - first set
44p2.00	Read	Cardiac troponin positive
7929100	Read	Percut transluminal coronary thrombolysis with streptokinase
7929111	Read	Percut translum coronary thrombolytic therapy- streptokinase
889A.00	Read	Diab mellit insulin-glucose infus acute myocardial infarct
8B3a.00	Read	Door to needle time
8B3g.00	Read	Pain to thrombolysis time
G30..00	Read	Acute myocardial infarction
G30..11	Read	Attack - heart
G30..12	Read	Coronary thrombosis
G30..13	Read	Cardiac rupture following myocardial infarction (MI)
G30..14	Read	Heart attack
G30..15	Read	MI - acute myocardial infarction
G30..16	Read	Thrombosis - coronary
G30..17	Read	Silent myocardial infarction
G300.00	Read	Acute anterolateral infarction
G301.00	Read	Other specified anterior myocardial infarction
G301000	Read	Acute anteroapical infarction
G301100	Read	Acute anteroapical infarction
G301z00	Read	Anterior myocardial infarction NOS
G302.00	Read	Acute inferolateral infarction
G303.00	Read	Acute inferoposterior infarction
G304.00	Read	Posterior myocardial infarction NOS
G305.00	Read	Lateral myocardial infarction NOS
G306.00	Read	True posterior myocardial infarction
G307.00	Read	Acute subendocardial infarction
G307000	Read	Acute non-Q wave infarction
G307100	Read	Acute non-ST segment elevation myocardial infarction
G308.00	Read	Inferior myocardial infarction NOS
G309.00	Read	Acute Q-wave infarct
G30A.00	Read	Mural thrombosis
G30B.00	Read	Acute posterolateral myocardial infarction
G30X.00	Read	Acute transmural myocardial infarction of unspecif site
G30X000	Read	Acute ST segment elevation myocardial infarction
G30y.00	Read	Other acute myocardial infarction
G30y000	Read	Acute atrial infarction
G30y100	Read	Acute papillary muscle infarction
G30y200	Read	Acute septal infarction
G30yz00	Read	Other acute myocardial infarction NOS
G30z.00	Read	Acute myocardial infarction NOS
G310.00	Read	Postmyocardial infarction syndrome
G310.11	Read	Dressler's syndrome

1			
2	G31y100	Read	Microinfarction of heart
3	G32..00	Read	Old myocardial infarction
4	G32..11	Read	Healed myocardial infarction
5	G32..12	Read	Personal history of myocardial infarction
6	G33z500	Read	Post infarct angina
7	G35..00	Read	Subsequent myocardial infarction
8	G350.00	Read	Subsequent myocardial infarction of anterior wall
9	G351.00	Read	Subsequent myocardial infarction of inferior wall
10	G353.00	Read	Subsequent myocardial infarction of other sites
11	G35X.00	Read	Subsequent myocardial infarction of unspecified site
12	G36..00	Read	Certain current complication follow acute myocardial infarct
13	G360.00	Read	Haemopericardium/current comp folow acut myocard infarct
14	G361.00	Read	Atrial septal defect/curr comp folow acut myocardal infarct
15	G362.00	Read	Ventric septal defect/curr comp fol acut myocardal infarctn
16	G363.00	Read	Ruptur cardiac wall w/out haemopericard/cur comp fol ac MI
17	G364.00	Read	Ruptur chordae tendinae/curr comp fol acute myocard infarct
18	G365.00	Read	Rupture papillary muscle/curr comp fol acute myocard infarct
19	G366.00	Read	Thrombosis atrium,auric append&vent/curr comp foll acute MI
20	G38..00	Read	Postoperative myocardial infarction
21	G380.00	Read	Postoperative transmural myocardial infarction anterior wall
22	G381.00	Read	Postoperative transmural myocardial infarction inferior wall
23	G384.00	Read	Postoperative subendocardial myocardial infarction
24	G38z.00	Read	Postoperative myocardial infarction, unspecified
25	G501.00	Read	Post infarction pericarditis
26	Gyu3400	Read	[X]Acute transmural myocardial infarction of unspecif site
27	I21	ICD-10	Acute myocardial infarction
28	I22	ICD-10	Subsequent myocardial infarction
29	I23	ICD-10	Certain current complications following acute myocardial infarction

Diabetes mellitus

Code	Classification	Description
13AB.00	Read	diabetic lipid lowering diet
13AC.00	Read	diabetic weight reducing diet
13B1.00	Read	diabetic diet
13L4.11	Read	diabetic child
1434	Read	h/o: diabetes mellitus
14F4.00	Read	h/o: admission in last year for diabetes foot problem
14P3.00	Read	h/o: insulin therapy
2BBF.00	Read	retinal abnormality - diabetes related
2BBJ.00	Read	o/e - no right diabetic retinopathy
2BBk.00	Read	o/e - right eye stable treated prolif diabetic retinopathy
2BBK.00	Read	o/e - no left diabetic retinopathy
2BBI.00	Read	o/e - left eye stable treated prolif diabetic retinopathy
2BBL.00	Read	o/e - diabetic maculopathy present both eyes
2BBM.00	Read	o/e - diabetic maculopathy absent both eyes

1			
2	2BB0.00	Read	o/e - sight threatening diabetic retinopathy
3	2BBP.00	Read	o/e - right eye background diabetic retinopathy
4	2BBQ.00	Read	o/e - left eye background diabetic retinopathy
5	2BBR.00	Read	impaired vision due to diabetic retinopathy
6	2BBR.00	Read	o/e - right eye preproliferative diabetic retinopathy
7	2BBS.00	Read	o/e - left eye preproliferative diabetic retinopathy
8	2BBT.00	Read	o/e - right eye proliferative diabetic retinopathy
9	2BBV.00	Read	o/e - left eye proliferative diabetic retinopathy
10	2BBW.00	Read	o/e - right eye diabetic maculopathy
11	2BBX.00	Read	o/e - left eye diabetic maculopathy
12	2G51000	Read	foot abnormality - diabetes related
13	2G5A.00	Read	o/e - right diabetic foot at risk
14	2G5B.00	Read	o/e - left diabetic foot at risk
15	2G5C.00	Read	foot abnormality - diabetes related
16	2G5d.00	Read	o/e - left diabetic foot at increased risk
17	2G5e.00	Read	o/e - right diabetic foot at increased risk
18	2G5E.00	Read	o/e - right diabetic foot at low risk
19	2G5F.00	Read	o/e - right diabetic foot at moderate risk
20	2G5G.00	Read	o/e - right diabetic foot at high risk
21	2G5H.00	Read	o/e - right diabetic foot - ulcerated
22	2G5I.00	Read	o/e - left diabetic foot at low risk
23	2G5J.00	Read	o/e - left diabetic foot at moderate risk
24	2G5K.00	Read	o/e - left diabetic foot at high risk
25	2G5L.00	Read	o/e - left diabetic foot - ulcerated
26	2G5V.00	Read	o/e - right chronic diabetic foot ulcer
27	2G5W.00	Read	o/e - left chronic diabetic foot ulcer
28	3881	Read	education score - diabetes
29	3882	Read	diabetes well being questionnaire
30	3883	Read	diabetes treatment satisfaction questionnaire
31	42c..00	Read	hba1 - diabetic control
32	42c0.00	Read	hba1 < 7% - good control
33	42c1.00	Read	hba1 7 - 10% - borderline control
34	42c2.00	Read	hba1 > 10% - bad control
35	42W..00	Read	hb. a1c - diabetic control
36	42W1.00	Read	hb. a1c < 7% - good control
37	42W2.00	Read	hb. a1c 7-10% - borderline
38	42W3.00	Read	hb. a1c > 10% - bad control
39	42WZ.00	Read	hb. a1c - diabetic control nos
40	44T9.00	Read	glucometer blood sugar
41	44UZ.00	Read	blood glucose 14+ mmol/l
42	44Uz.11	Read	blood hyperglycaemia nos
43	44V3.00	Read	glucose tol. test diabetic
44	661N400	Read	diabetes self-management plan review
45	66A..00	Read	diabetic monitoring
46	66A1.00	Read	initial diabetic assessment

1			
2	66A2.00	Read	follow-up diabetic assessment
3	66A3.00	Read	diabetic on diet only
4	66A4.00	Read	diabetic on oral treatment
5	66A5.00	Read	diabetic on insulin
6	66A8.00	Read	has seen dietician - diabetes
7	66A9.00	Read	understands diet - diabetes
10	66Aa.00	Read	diabetic diet - poor compliance
11	66AA.11	Read	injection sites - diabetic
12	66Ab.00	Read	diabetic foot examination
13	66Ac.00	Read	diabetic peripheral neuropathy screening
14	66AC.00	Read	blood sugar charts
15	66AD.00	Read	fundoscopy - diabetic check
16	66Ae.00	Read	hba1c target
17	66AE.00	Read	feet examination
18	66Af.00	Read	patient diabetes education review
19	66Ag.00	Read	insulin needles changed daily
20	66AG.00	Read	diabetic drug side effects
21	66Ah.00	Read	insulin needles changed for each injection
22	66AH.00	Read	diabetic treatment changed
23	66AH000	Read	conversion to insulin
24	66AH100	Read	conversion to insulin in secondary care
25	66AH200	Read	conversion to insulin by diabetes specialist nurse
26	66Ai.00	Read	diabetic 6 month review
27	66Ai.00	Read	diabetic - good control
28	66Aj.00	Read	insulin needles changed less than once a day
29	66AJ.00	Read	diabetic - poor control
30	66AJ000	Read	chronic hyperglycaemia
31	66AJ100	Read	brittle diabetes
32	66AJ.11	Read	unstable diabetes
33	66AJ200	Read	loss of hypoglycaemic warning
34	66AJz00	Read	diabetic - poor control nos
35	66Ak.00	Read	diabetic monitoring - lower risk albumin excretion
36	66AK.00	Read	diabetic - cooperative patient
37	66Al.00	Read	diabetic monitoring - higher risk albumin excretion
38	66AL.00	Read	diabetic-uncooperative patient
39	66Am.00	Read	insulin dose changed
40	66AM.00	Read	diabetic - follow-up default
41	66An.00	Read	diabetes type 1 review
42	66AN.00	Read	date diabetic treatment start
43	66Ao.00	Read	diabetes type 2 review
44	66Ap.00	Read	insulin treatment initiated
45	66AP.00	Read	diabetes: practice programme
46	66Aq.00	Read	diabetic foot screen
47	66AQ.00	Read	diabetes: shared care programme
48	66AQ000	Read	unsuitable for diabetes year of care programme

1			
2	66AQ100	Read	declined consent for diabetes year of care programme
3	66AR.00	Read	diabetes management plan given
4	66As.00	Read	diabetic on subcutaneous treatment
5	66AS.00	Read	diabetic annual review
6	66AS000	Read	diabetes year of care annual review
7	66At.00	Read	diabetic dietary review
8	66AT.00	Read	annual diabetic blood test
9	66At000	Read	type i diabetic dietary review
10	66At011	Read	type 1 diabetic dietary review
11	66At100	Read	type ii diabetic dietary review
12	66At111	Read	type 2 diabetic dietary review
13	66Au.00	Read	diabetic erectile dysfunction review
14	66AU.00	Read	diabetes care by hospital only
15	66Av.00	Read	diabetic assessment of erectile dysfunction
16	66AV.00	Read	diabetic on insulin and oral treatment
17	66Aw.00	Read	insulin dose
18	66AW.00	Read	diabetic foot risk assessment
19	66AY.00	Read	diabetic diet - good compliance
20	66AZ.00	Read	diabetic monitoring nos
21	66b1.00	Read	diabetic monitoring not required
22	66o..00	Read	further diabetic monitoring
23	6761	Read	diabetic pre-pregnancy counselling
24	679c.00	Read	insulin administration education
25	679L.00	Read	health education - diabetes
26	679L000	Read	education in self management of diabetes
27	679L200	Read	education about diabetes and driving
28	679L211	Read	advice about diabetes and driving
29	679R.00	Read	patient offered diabetes structured education programme
30	67D8.00	Read	provision of diabetes clinical summary
31	68A7.00	Read	diabetic retinopathy screening
32	68A9.00	Read	diabetic retinopathy screening offered
33	68AB.00	Read	diabetic digital retinopathy screening offered
34	7276	Read	pan retinal photocoagulation for diabetes
35	7L10000	Read	continuous subcutaneous infusion of insulin
36	7L10011	Read	subcutaneous infusion with insulin pump
37	7L19800	Read	subcutaneous injection of insulin
38	7L19J00	Read	subcutaneous injection of exenatide
39	889A.00	Read	diab mellit insulin-glucose infus acute myocardial infarct
40	8A12.00	Read	diabetic crisis monitoring
41	8A13.00	Read	diabetic stabilisation
42	8A17.00	Read	self monitoring of blood glucose
43	8B3l.00	Read	diabetes medication review
44	8BAi.00	Read	insulin passport completed
45	8BAj.00	Read	informed dissent not to carry insulin passport
46	8BAm.00	Read	insulin passport checked

1			
2	8BL2.00	Read	patient on maximal tolerated therapy for diabetes
3	8CA4100	Read	pt advised re diabetic diet
4	8CAQ.00	Read	advice about blood glucose control
5	8CE0100	Read	insulin alert patient information booklet given
6	8CE0200	Read	insulin passport given
7	8CMW700	Read	diabetes clinical pathway
10	8CP2.00	Read	transition of diabetes care options discussed
11	8CR2.00	Read	diabetes clinical management plan
12	8CS0.00	Read	diabetes care plan agreed
13	8H2J.00	Read	admit diabetic emergency
14	8H3O.00	Read	non-urgent diabetic admission
15	8H4e.00	Read	referral to diabetes special interest general practitioner
16	8H4F.00	Read	referral to diabetologist
17	8H7C.00	Read	refer
18	8H7f.00	Read	referral to diabetes nurse
19	8H7r.00	Read	refer to diabetic foot screener
20	8HBG.00	Read	diabetic retinopathy 12 month review
21	8HBH.00	Read	diabetic retinopathy 6 month review
22	8Hg4.00	Read	discharged from care of diabetes specialist nurse
23	8HgC.00	Read	discharged from diabetes shared care programme
24	8HHy.00	Read	referral to diabetic register
25	8Hj0.00	Read	referral to diabetes structured education programme
26	8Hj3.00	Read	referral to dafne diabetes structured education programme
27	8Hj4.00	Read	referral to desmond diabetes structured education programme
28	8Hj5.00	Read	referral to xpert diabetes structured education programme
29	8HKE.00	Read	diabetology d.v. requested
30	8HI1.00	Read	referral for diabetic retinopathy screening
31	8HI4.00	Read	referral to community diabetes specialist nurse
32	8Hlc.00	Read	referral to community diabetes service
33	8HLE.00	Read	diabetology d.v. done
34	8HME.00	Read	listed for diabetology admissn
35	8HTe.00	Read	referral to diabetes preconception counselling clinic
36	8HTE100	Read	referral to community diabetes clinic
37	8HTi.00	Read	referral to multidisciplinary diabetic clinic
38	8HTk.00	Read	referral to diabetic eye clinic
39	8HVU.00	Read	private referral to diabetologist
40	8I2P.00	Read	sulphonylureas contraindicated
41	8I2S.00	Read	glitazones contraindicated
42	8I3k.00	Read	insulin therapy declined
43	8I3W.00	Read	diabetic foot examination declined
44	8I3X.00	Read	diabetic retinopathy screening refused
45	8I57.00	Read	patient held diabetic record declined
46	8I6F.00	Read	diabetic retinopathy screening not indicated
47	8I6G.00	Read	diabetic foot examination not indicated
48	8I7B.00	Read	metformin not tolerated

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1			
2	8I7C.00	Read	sulphonylureas not tolerated
3	8I81.00	Read	did not complete diabetes structured education programme
4	8I82.00	Read	did not complete dafne diabetes structured education program
5	8I83.00	Read	did not complete desmond diabetes structured educat program
6	8I84.00	Read	did not complete xpert diabetes structured education program
7	8IAs.00	Read	diabetic dietary review declined
8	8IE2.00	Read	diabetes care plan declined
9	8IEa.00	Read	referral to dafne diabetes structured educn prog declined
10	8IEQ.00	Read	referral to community diabetes specialist nurse declined
11	8OA3.00	Read	provision of written information about diabetes and driving
12	9360	Read	patient held diabetic record issued
13	93C4.00	Read	patient consent given for addition to diabetic register
14	9b92000	Read	diabetic medicine
15	9h4..00	Read	exception reporting: diabetes quality indicators
16	9h41.00	Read	excepted from diabetes qual indicators: patient unsuitable
17	9h42.00	Read	excepted from diabetes quality indicators: informed dissent
18	9kL..00	Read	insulin initiation - enhanced services administration
19	9m00.00	Read	eligible for diabetic retinopathy screening
20	9M00.00	Read	informed consent for diabetes national audit
21	9m0A.00	Read	declined diabetic retinopathy screening
22	9M10.00	Read	informed dissent for diabetes national audit
23	9N0m.00	Read	seen in diabetic nurse consultant clinic
24	9N0n.00	Read	seen in community diabetes specialist clinic
25	9N0o.00	Read	seen in community diabetic specialist nurse clinic
26	9N1i.00	Read	seen in diabetic foot clinic
27	9N1o.00	Read	seen in multidisciplinary diabetic clinic
28	9N1Q.00	Read	seen in diabetic clinic
29	9N1v.00	Read	seen in diabetic eye clinic
30	9N2d.00	Read	seen by diabetologist
31	9N2i.00	Read	seen by diabetic liaison nurse
32	9N4I.00	Read	dna - did not attend diabetic clinic
33	9N4p.00	Read	did not attend diabetic retinopathy clinic
34	9NiA.00	Read	did not attend diabetes structured education programme
35	9NiC.00	Read	did not attend dafne diabetes structured education programme
36	9NiD.00	Read	did not attend desmond diabetes structured education program
37	9NiE.00	Read	did not attend xpert diabetes structured education programme
38	9NiZ.00	Read	did not attend diabetes foot screening
39	9N14.00	Read	seen by general practitioner special interest in diabetes
40	9NM0.00	Read	attending diabetes clinic
41	9NN8.00	Read	under care of diabetologist
42	9NN9.00	Read	under care of diabetes specialist nurse
43	9NND.00	Read	under care of diabetic foot screener
44	9OL..00	Read	diabetes monitoring admin.
45	9OL1.00	Read	attends diabetes monitoring
46	9OL..11	Read	diabetes clinic administration

1			
2	9OL2.00	Read	refuses diabetes monitoring
3	9OL3.00	Read	diabetes monitoring default
4	9OL4.00	Read	diabetes monitoring 1st letter
5	9OL5.00	Read	diabetes monitoring 2nd letter
6	9OL6.00	Read	diabetes monitoring 3rd letter
7	9OL7.00	Read	diabetes monitor.verbal invite
8	9OL8.00	Read	diabetes monitor.phone invite
9	9OL9.00	Read	diabetes monitoring deleted
10	9OLA.00	Read	diabetes monitor. check done
11	9OLA.11	Read	diabetes monitored
12	9OLB.00	Read	attended diabetes structured education programme
13	9OLD.00	Read	diabetic patient unsuitable for digital retinal photography
14	9OLE.00	Read	attended desmond structured programme
15	9OLF.00	Read	diabetes structured education programme completed
16	9OLG.00	Read	attended xpert diabetes structured education programme
17	9OLH.00	Read	attended dafne diabetes structured education programme
18	9OLJ.00	Read	dafne diabetes structured education programme completed
19	9OLK.00	Read	desmond diabetes structured education programme completed
20	9OLL.00	Read	xpert diabetes structured education programme completed
21	9OLM.00	Read	diabetes structured education programme declined
22	9OLN.00	Read	diabetes monitor invitation by sms (short message service)
23	9OLZ.00	Read	diabetes monitoring admin.nos
24	9Oy..00	Read	diabetes screening administration
25	9Oy0000	Read	diabetic foot screening invitation
26	9Oy0200	Read	diabetic foot screening invitation first letter
27	9Oy0300	Read	diabetic foot screening invitation second letter
28	9Oy0400	Read	diabetic foot screening invitation third letter
29	C10..00	Read	diabetes mellitus
30	C100.00	Read	diabetes mellitus with no mention of complication
31	C100000	Read	diabetes mellitus
32	C100011	Read	insulin dependent diabetes mellitus
33	C100100	Read	diabetes mellitus
34	C100111	Read	maturity onset diabetes
35	C100112	Read	non-insulin dependent diabetes mellitus
36	C100z00	Read	diabetes mellitus nos with no mention of complication
37	C101.00	Read	diabetes mellitus with ketoacidosis
38	C101000	Read	diabetes mellitus
39	C101100	Read	diabetes mellitus
40	C101y00	Read	other specified diabetes mellitus with ketoacidosis
41	C101z00	Read	diabetes mellitus nos with ketoacidosis
42	C102.00	Read	diabetes mellitus with hyperosmolar coma
43	C102000	Read	diabetes mellitus
44	C102100	Read	diabetes mellitus
45	C102z00	Read	diabetes mellitus nos with hyperosmolar coma
46	C103.00	Read	diabetes mellitus with ketoacidotic coma

1			
2	C103000	Read	diabetes mellitus
3	C103100	Read	diabetes mellitus
4	C103y00	Read	other specified diabetes mellitus with coma
5	C103z00	Read	diabetes mellitus nos with ketoacidotic coma
6	C104.00	Read	diabetes mellitus with renal manifestation
7	C104000	Read	diabetes mellitus
8	C104100	Read	diabetes mellitus
9	C104.11	Read	diabetic nephropathy
10	C104y00	Read	other specified diabetes mellitus with renal complications
11	C104z00	Read	diabetes mellitus with nephropathy nos
12	C105.00	Read	diabetes mellitus with ophthalmic manifestation
13	C105000	Read	diabetes mellitus
14	C105100	Read	diabetes mellitus
15	C105y00	Read	other specified diabetes mellitus with ophthalmic complicatn
16	C105z00	Read	diabetes mellitus nos with ophthalmic manifestation
17	C106.00	Read	diabetes mellitus with neurological manifestation
18	C106000	Read	diabetes mellitus
19	C106100	Read	diabetes mellitus
20	C106.11	Read	diabetic amyotrophy
21	C106.12	Read	diabetes mellitus with neuropathy
22	C106.13	Read	diabetes mellitus with polyneuropathy
23	C106y00	Read	other specified diabetes mellitus with neurological comps
24	C106z00	Read	diabetes mellitus nos with neurological manifestation
25	C107.00	Read	diabetes mellitus with peripheral circulatory disorder
26	C107000	Read	diabetes mellitus
27	C107100	Read	diabetes mellitus
28	C107.11	Read	diabetes mellitus with gangrene
29	C107.12	Read	diabetes with gangrene
30	C107200	Read	diabetes mellitus
31	C107300	Read	idm with peripheral circulatory disorder
32	C107400	Read	niddm with peripheral circulatory disorder
33	C107z00	Read	diabetes mellitus nos with peripheral circulatory disorder
34	C108.00	Read	insulin dependent diabetes mellitus
35	C108000	Read	insulin-dependent diabetes mellitus with renal complications
36	C108011	Read	type i diabetes mellitus with renal complications
37	C108012	Read	type 1 diabetes mellitus with renal complications
38	C108100	Read	insulin-dependent diabetes mellitus with ophthalmic comps
39	C108.11	Read	idm-insulin dependent diabetes mellitus
40	C108112	Read	type 1 diabetes mellitus with ophthalmic complications
41	C108.12	Read	type 1 diabetes mellitus
42	C108.13	Read	type i diabetes mellitus
43	C108200	Read	insulin-dependent diabetes mellitus with neurological comps
44	C108211	Read	type i diabetes mellitus with neurological complications
45	C108212	Read	type 1 diabetes mellitus with neurological complications
46	C108300	Read	insulin dependent diabetes mellitus with multiple complicatn

1			
2	C108311	Read	type i diabetes mellitus with multiple complications
3	C108400	Read	unstable insulin dependent diabetes mellitus
4			
5	C108411	Read	unstable type i diabetes mellitus
6	C108412	Read	unstable type 1 diabetes mellitus
7			
8	C108500	Read	insulin dependent diabetes mellitus with ulcer
9	C108511	Read	type i diabetes mellitus with ulcer
10	C108512	Read	type 1 diabetes mellitus with ulcer
11			
12	C108600	Read	insulin dependent diabetes mellitus with gangrene
13	C108700	Read	insulin dependent diabetes mellitus with retinopathy
14	C108711	Read	type i diabetes mellitus with retinopathy
15			
16	C108712	Read	type 1 diabetes mellitus with retinopathy
17	C108800	Read	insulin dependent diabetes mellitus - poor control
18	C108811	Read	type i diabetes mellitus - poor control
19			
20	C108812	Read	type 1 diabetes mellitus - poor control
21	C108900	Read	insulin dependent diabetes maturity onset
22	C108911	Read	type i diabetes mellitus maturity onset
23			
24	C108912	Read	type 1 diabetes mellitus maturity onset
25	C108A00	Read	insulin-dependent diabetes without complication
26	C108A11	Read	type i diabetes mellitus without complication
27			
28	C108B00	Read	insulin dependent diabetes mellitus with mononeuropathy
29	C108B11	Read	type i diabetes mellitus with mononeuropathy
30			
31	C108C00	Read	insulin dependent diabetes mellitus with polyneuropathy
32	C108D00	Read	insulin dependent diabetes mellitus with nephropathy
33	C108D11	Read	type i diabetes mellitus with nephropathy
34			
35	C108E00	Read	insulin dependent diabetes mellitus with hypoglycaemic coma
36	C108E11	Read	type i diabetes mellitus with hypoglycaemic coma
37			
38	C108E12	Read	type 1 diabetes mellitus with hypoglycaemic coma
39	C108F00	Read	insulin dependent diabetes mellitus with diabetic cataract
40	C108F11	Read	type i diabetes mellitus with diabetic cataract
41			
42	C108G00	Read	insulin dependent diab mell with peripheral angiopathy
43	C108H00	Read	insulin dependent diabetes mellitus with arthropathy
44			
45	C108H11	Read	type i diabetes mellitus with arthropathy
46	C108J00	Read	insulin dependent diab mell with neuropathic arthropathy
47	C108J11	Read	type i diabetes mellitus with neuropathic arthropathy
48			
49	C108J12	Read	type 1 diabetes mellitus with neuropathic arthropathy
50	C108y00	Read	other specified diabetes mellitus with multiple comps
51			
52	C108z00	Read	unspecified diabetes mellitus with multiple complications
53	C109.00	Read	non-insulin dependent diabetes mellitus
54	C109000	Read	non-insulin-dependent diabetes mellitus with renal comps
55	C109011	Read	type ii diabetes mellitus with renal complications
56			
57	C109012	Read	type 2 diabetes mellitus with renal complications
58	C109100	Read	non-insulin-dependent diabetes mellitus with ophthalm comps
59	C109.11	Read	niddm - non-insulin dependent diabetes mellitus
60			
	C109111	Read	type ii diabetes mellitus with ophthalmic complications
	C109112	Read	type 2 diabetes mellitus with ophthalmic complications

1			
2	C109.12	Read	type 2 diabetes mellitus
3	C109.13	Read	type ii diabetes mellitus
4			
5	C109200	Read	non-insulin-dependent diabetes mellitus with neuro comps
6	C109211	Read	type ii diabetes mellitus with neurological complications
7			
8	C109212	Read	type 2 diabetes mellitus with neurological complications
9	C109300	Read	non-insulin-dependent diabetes mellitus with multiple comps
10	C109312	Read	type 2 diabetes mellitus with multiple complications
11			
12	C109400	Read	non-insulin dependent diabetes mellitus with ulcer
13	C109411	Read	type ii diabetes mellitus with ulcer
14	C109412	Read	type 2 diabetes mellitus with ulcer
15			
16	C109500	Read	non-insulin dependent diabetes mellitus with gangrene
17	C109511	Read	type ii diabetes mellitus with gangrene
18	C109512	Read	type 2 diabetes mellitus with gangrene
19			
20	C109600	Read	non-insulin-dependent diabetes mellitus with retinopathy
21	C109611	Read	type ii diabetes mellitus with retinopathy
22	C109612	Read	type 2 diabetes mellitus with retinopathy
23			
24	C109700	Read	non-insulin dependent diabetes mellitus - poor control
25	C109711	Read	type ii diabetes mellitus - poor control
26	C109712	Read	type 2 diabetes mellitus - poor control
27			
28	C109900	Read	non-insulin-dependent diabetes mellitus without complication
29	C109912	Read	type 2 diabetes mellitus without complication
30			
31	C109A00	Read	non-insulin dependent diabetes mellitus with mononeuropathy
32	C109A11	Read	type ii diabetes mellitus with mononeuropathy
33			
34	C109B00	Read	non-insulin dependent diabetes mellitus with polyneuropathy
35	C109B11	Read	type ii diabetes mellitus with polyneuropathy
36	C109C00	Read	non-insulin dependent diabetes mellitus with nephropathy
37	C109C11	Read	type ii diabetes mellitus with nephropathy
38	C109C12	Read	type 2 diabetes mellitus with nephropathy
39			
40	C109D00	Read	non-insulin dependent diabetes mellitus with hypoglyca coma
41	C109D11	Read	type ii diabetes mellitus with hypoglycaemic coma
42	C109D12	Read	type 2 diabetes mellitus with hypoglycaemic coma
43			
44	C109E00	Read	non-insulin depend diabetes mellitus with diabetic cataract
45	C109E11	Read	type ii diabetes mellitus with diabetic cataract
46	C109E12	Read	type 2 diabetes mellitus with diabetic cataract
47			
48	C109F00	Read	non-insulin-dependent d m with peripheral angiopath
49	C109F11	Read	type ii diabetes mellitus with peripheral angiopathy
50	C109F12	Read	type 2 diabetes mellitus with peripheral angiopathy
51			
52	C109G00	Read	non-insulin dependent diabetes mellitus with arthropathy
53	C109G11	Read	type ii diabetes mellitus with arthropathy
54	C109G12	Read	type 2 diabetes mellitus with arthropathy
55			
56	C109H00	Read	non-insulin dependent d m with neuropathic arthropathy
57	C109H11	Read	type ii diabetes mellitus with neuropathic arthropathy
58	C109H12	Read	type 2 diabetes mellitus with neuropathic arthropathy
59			
60	C109J00	Read	insulin treated type 2 diabetes mellitus
	C109J11	Read	insulin treated non-insulin dependent diabetes mellitus

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2	C109J12	Read	insulin treated type ii diabetes mellitus
3	C109K00	Read	hyperosmolar non-ketotic state in type 2 diabetes mellitus
4	C10B.00	Read	diabetes mellitus induced by steroids
5	C10B000	Read	steroid induced diabetes mellitus without complication
6	C10C.00	Read	diabetes mellitus autosomal dominant
7	C10C.11	Read	maturity onset diabetes in youth
8	C10C.12	Read	maturity onset diabetes in youth type 1
9	C10D.00	Read	diabetes mellitus autosomal dominant type 2
10	C10D.11	Read	maturity onset diabetes in youth type 2
11	C10E.00	Read	type 1 diabetes mellitus
12	C10E000	Read	type 1 diabetes mellitus with renal complications
13	C10E012	Read	insulin-dependent diabetes mellitus with renal complications
14	C10E100	Read	type 1 diabetes mellitus with ophthalmic complications
15	C10E.11	Read	type i diabetes mellitus
16	C10E111	Read	type i diabetes mellitus with ophthalmic complications
17	C10E112	Read	insulin-dependent diabetes mellitus with ophthalmic comps
18	C10E.12	Read	insulin dependent diabetes mellitus
19	C10E200	Read	type 1 diabetes mellitus with neurological complications
20	C10E212	Read	insulin-dependent diabetes mellitus with neurological comps
21	C10E300	Read	type 1 diabetes mellitus with multiple complications
22	C10E311	Read	type i diabetes mellitus with multiple complications
23	C10E312	Read	insulin dependent diabetes mellitus with multiple complicat
24	C10E400	Read	unstable type 1 diabetes mellitus
25	C10E411	Read	unstable type i diabetes mellitus
26	C10E412	Read	unstable insulin dependent diabetes mellitus
27	C10E500	Read	type 1 diabetes mellitus with ulcer
28	C10E511	Read	type i diabetes mellitus with ulcer
29	C10E512	Read	insulin dependent diabetes mellitus with ulcer
30	C10E600	Read	type 1 diabetes mellitus with gangrene
31	C10E611	Read	type i diabetes mellitus with gangrene
32	C10E700	Read	type 1 diabetes mellitus with retinopathy
33	C10E711	Read	type i diabetes mellitus with retinopathy
34	C10E712	Read	insulin dependent diabetes mellitus with retinopathy
35	C10E800	Read	type 1 diabetes mellitus - poor control
36	C10E811	Read	type i diabetes mellitus - poor control
37	C10E812	Read	insulin dependent diabetes mellitus - poor control
38	C10E900	Read	type 1 diabetes mellitus maturity onset
39	C10E911	Read	type i diabetes mellitus maturity onset
40	C10E912	Read	insulin dependent diabetes maturity onset
41	C10EA00	Read	type 1 diabetes mellitus without complication
42	C10EA11	Read	type i diabetes mellitus without complication
43	C10EA12	Read	insulin-dependent diabetes without complication
44	C10EB00	Read	type 1 diabetes mellitus with mononeuropathy
45	C10EC00	Read	type 1 diabetes mellitus with polyneuropathy
46	C10EC11	Read	type i diabetes mellitus with polyneuropathy

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2	C10EC12	Read	insulin dependent diabetes mellitus with polyneuropathy
3	C10ED00	Read	type 1 diabetes mellitus with nephropathy
4	C10ED12	Read	insulin dependent diabetes mellitus with nephropathy
5	C10EE00	Read	type 1 diabetes mellitus with hypoglycaemic coma
6	C10EE12	Read	insulin dependent diabetes mellitus with hypoglycaemic coma
7			
8	C10EF00	Read	type 1 diabetes mellitus with diabetic cataract
9	C10EF12	Read	insulin dependent diabetes mellitus with diabetic cataract
10			
11	C10EG00	Read	type 1 diabetes mellitus with peripheral angiopathy
12	C10EH00	Read	type 1 diabetes mellitus with arthropathy
13	C10EJ00	Read	type 1 diabetes mellitus with neuropathic arthropathy
14	C10EK00	Read	type 1 diabetes mellitus with persistent proteinuria
15	C10EL00	Read	type 1 diabetes mellitus with persistent microalbuminuria
16	C10EL11	Read	type i diabetes mellitus with persistent microalbuminuria
17	C10EM00	Read	type 1 diabetes mellitus with ketoacidosis
18	C10EM11	Read	type i diabetes mellitus with ketoacidosis
19	C10EN00	Read	type 1 diabetes mellitus with ketoacidotic coma
20	C10EN11	Read	type i diabetes mellitus with ketoacidotic coma
21	C10EP00	Read	type 1 diabetes mellitus with exudative maculopathy
22	C10EP11	Read	type i diabetes mellitus with exudative maculopathy
23	C10EQ00	Read	type 1 diabetes mellitus with gastroparesis
24	C10ER00	Read	latent autoimmune diabetes mellitus in adult
25	C10F.00	Read	type 2 diabetes mellitus
26	C10F000	Read	type 2 diabetes mellitus with renal complications
27	C10F011	Read	type ii diabetes mellitus with renal complications
28	C10F100	Read	type 2 diabetes mellitus with ophthalmic complications
29	C10F.11	Read	type ii diabetes mellitus
30	C10F111	Read	type ii diabetes mellitus with ophthalmic complications
31	C10F200	Read	type 2 diabetes mellitus with neurological complications
32	C10F211	Read	type ii diabetes mellitus with neurological complications
33	C10F300	Read	type 2 diabetes mellitus with multiple complications
34	C10F311	Read	type ii diabetes mellitus with multiple complications
35	C10F400	Read	type 2 diabetes mellitus with ulcer
36	C10F411	Read	type ii diabetes mellitus with ulcer
37	C10F500	Read	type 2 diabetes mellitus with gangrene
38	C10F511	Read	type ii diabetes mellitus with gangrene
39	C10F600	Read	type 2 diabetes mellitus with retinopathy
40	C10F611	Read	type ii diabetes mellitus with retinopathy
41	C10F700	Read	type 2 diabetes mellitus - poor control
42	C10F711	Read	type ii diabetes mellitus - poor control
43	C10F900	Read	type 2 diabetes mellitus without complication
44	C10F911	Read	type ii diabetes mellitus without complication
45	C10FA00	Read	type 2 diabetes mellitus with mononeuropathy
46	C10FA11	Read	type ii diabetes mellitus with mononeuropathy
47	C10FB00	Read	type 2 diabetes mellitus with polyneuropathy
48	C10FB11	Read	type ii diabetes mellitus with polyneuropathy
49			
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2	C10FC00	Read	type 2 diabetes mellitus with nephropathy
3	C10FC11	Read	type ii diabetes mellitus with nephropathy
4			
5	C10FD00	Read	type 2 diabetes mellitus with hypoglycaemic coma
6	C10FD11	Read	type ii diabetes mellitus with hypoglycaemic coma
7			
8	C10FE00	Read	type 2 diabetes mellitus with diabetic cataract
9	C10FE11	Read	type ii diabetes mellitus with diabetic cataract
10	C10FF00	Read	type 2 diabetes mellitus with peripheral angiopathy
11	C10FF11	Read	type ii diabetes mellitus with peripheral angiopathy
12			
13	C10FG00	Read	type 2 diabetes mellitus with arthropathy
14	C10FG11	Read	type ii diabetes mellitus with arthropathy
15			
16	C10FH00	Read	type 2 diabetes mellitus with neuropathic arthropathy
17	C10FJ00	Read	insulin treated type 2 diabetes mellitus
18	C10FJ11	Read	insulin treated type ii diabetes mellitus
19			
20	C10FK00	Read	hyperosmolar non-ketotic state in type 2 diabetes mellitus
21	C10FK11	Read	hyperosmolar non-ketotic state in type ii diabetes mellitus
22			
23	C10FL00	Read	type 2 diabetes mellitus with persistent proteinuria
24	C10FL11	Read	type ii diabetes mellitus with persistent proteinuria
25			
26	C10FM00	Read	type 2 diabetes mellitus with persistent microalbuminuria
27	C10FM11	Read	type ii diabetes mellitus with persistent microalbuminuria
28	C10FN00	Read	type 2 diabetes mellitus with ketoacidosis
29	C10FN11	Read	type ii diabetes mellitus with ketoacidosis
30			
31	C10FP00	Read	type 2 diabetes mellitus with ketoacidotic coma
32	C10FP11	Read	type ii diabetes mellitus with ketoacidotic coma
33			
34	C10FQ00	Read	type 2 diabetes mellitus with exudative maculopathy
35	C10FR00	Read	type 2 diabetes mellitus with gastroparesis
36			
37	C10FS00	Read	maternally inherited diabetes mellitus
38	C10G.00	Read	secondary pancreatic diabetes mellitus
39	C10G000	Read	secondary pancreatic diabetes mellitus without complication
40			
41	C10M.00	Read	lipomatrophic diabetes mellitus
42	C10N.00	Read	secondary diabetes mellitus
43	C10N000	Read	secondary diabetes mellitus without complication
44			
45	C10N100	Read	cystic fibrosis related diabetes mellitus
46	C10y.00	Read	diabetes mellitus with other specified manifestation
47	C10y100	Read	diabetes mellitus
48			
49	C10yy00	Read	other specified diabetes mellitus with other spec comps
50	C10yz00	Read	diabetes mellitus nos with other specified manifestation
51			
52	C10z.00	Read	diabetes mellitus with unspecified complication
53	C10z000	Read	diabetes mellitus
54	C10z100	Read	diabetes mellitus
55			
56	C10zy00	Read	other specified diabetes mellitus with unspecified comps
57	C10zz00	Read	diabetes mellitus nos with unspecified complication
58			
59	C110000	Read	iatrogenic hyperinsulinism
60	C110.11	Read	insulin coma
	C11y000	Read	steroid induced diabetes
	C314.11	Read	renal diabetes

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2	C350011	Read	bronzed diabetes
3	Cyu2.00	Read	[x]diabetes mellitus
4	Cyu2000	Read	[x]other specified diabetes mellitus
5	Cyu2300	Read	[x]unspecified diabetes mellitus with renal complications
6	F171100	Read	autonomic neuropathy due to diabetes
7	F345000	Read	diabetic mononeuritis multiplex
8	F35z000	Read	diabetic mononeuritis nos
9	F372.00	Read	polyneuropathy in diabetes
10	F372000	Read	acute painful diabetic neuropathy
11	F372100	Read	chronic painful diabetic neuropathy
12	F372.11	Read	diabetic polyneuropathy
13	F372.12	Read	diabetic neuropathy
14	F372200	Read	asymptomatic diabetic neuropathy
15	F381300	Read	myasthenic syndrome due to diabetic amyotrophy
16	F381311	Read	diabetic amyotrophy
17	F3y0.00	Read	diabetic mononeuropathy
18	F420.00	Read	diabetic retinopathy
19	F420000	Read	background diabetic retinopathy
20	F420100	Read	proliferative diabetic retinopathy
21	F420200	Read	preproliferative diabetic retinopathy
22	F420300	Read	advanced diabetic maculopathy
23	F420400	Read	diabetic maculopathy
24	F420500	Read	advanced diabetic retinal disease
25	F420600	Read	non proliferative diabetic retinopathy
26	F420700	Read	high risk proliferative diabetic retinopathy
27	F420800	Read	high risk non proliferative diabetic retinopathy
28	F420z00	Read	diabetic retinopathy nos
29	F440700	Read	diabetic iritis
30	F464000	Read	diabetic cataract
31	G73y000	Read	diabetic peripheral angiopathy
32	K01x100	Read	nephrotic syndrome in diabetes mellitus
33	K01x111	Read	kimmelstiel - wilson disease
34	K08yA00	Read	proteinuric diabetic nephropathy
35	K08yA11	Read	clinical diabetic nephropathy
36	K27y700	Read	erectile dysfunction due to diabetes mellitus
37	Kyu0300	Read	[x]glomerular disorders in diabetes mellitus
38	L180500	Read	pre-existing diabetes mellitus
39	L180600	Read	pre-existing diabetes mellitus
40	L180X00	Read	pre-existing diabetes mellitus
41	M037200	Read	cellulitis in diabetic foot
42	M21yC00	Read	insulin lipohypertrophy
43	M21yC11	Read	insulin site lipohypertrophy
44	M271000	Read	ischaemic ulcer diabetic foot
45	M271100	Read	neuropathic diabetic ulcer - foot
46	M271200	Read	mixed diabetic ulcer - foot

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2	N030000	Read	diabetic cheiroarthropathy
3	N030011	Read	diabetic cheiropathy
4	N030100	Read	diabetic charcot arthropathy
5			
6	R054200	Read	[d]gangrene of toe in diabetic
7			
8	R054300	Read	[d]widespread diabetic foot gangrene
9	R105712	Read	[d]hyperglycaemia
10	Ryu8A00	Read	[x]hyperglycaemia
11			
12	TJ23.00	Read	adverse reaction to insulins and antidiabetic agents
13	TJ23000	Read	adverse reaction to insulins
14	TJ23200	Read	adverse reaction to chlorpropamide
15	TJ23300	Read	adverse reaction to glibenclamide
16	TJ23400	Read	adverse reaction to gliclazide
17	TJ23500	Read	adverse reaction to glipizide
18			
19	TJ23800	Read	adverse reaction to tolazamide
20	TJ23900	Read	adverse reaction to tolbutamide
21			
22	TJ23A00	Read	adverse reaction to metformin hydrochloride
23			
24	TJ23B00	Read	adverse reaction to glucagon
25	TJ23z00	Read	adverse reaction to insulins and antidiabetic agents nos
26			
27	U602300	Read	[x]insul/oral hypoglyc drugs caus adverse eff therapeut use
28	U602311	Read	[x] adverse reaction to insulins and antidiabetic agents
29	U602312	Read	[x] adverse reaction to insulins
30	U602315	Read	[x] adverse reaction to glibenclamide
31	U602316	Read	[x] adverse reaction to gliclazide
32	U602317	Read	[x] adverse reaction to glipzide
33	U602318	Read	[x] adverse reaction to gliquidone
34	U60231A	Read	[x] adverse reaction to tolazamide
35	U60231B	Read	[x] adverse reaction to tolbutamide
36	U60231C	Read	[x] adverse reaction to metformin hydrochloride
37	U60231E	Read	[x] adverse reaction to insulins and antidiabetic agents nos
38			
39	ZC2C800	Read	dietary advice for diabetes mellitus
40	ZC2C900	Read	dietary advice for type i diabetes
41	ZC2CA00	Read	dietary advice for type ii diabetes
42			
43	ZL22500	Read	under care of diabetic liaison nurse
44	ZL62500	Read	referral to diabetes nurse
45	ZL62600	Read	referral to diabetic liaison nurse
46			
47	ZLA2500	Read	seen by diabetic liaison nurse
48	ZLD7500	Read	discharge by diabetic liaison nurse
49	ZRB4.00	Read	diabetes clinic satisfaction questionnaire
50	ZRB4.11	Read	csq - diabetes clinic satisfaction questionnaire
51	ZRB5.00	Read	diabetes treatment satisfaction questionnaire
52	ZRB5.11	Read	dtsq - diabetes treatment satisfaction questionnaire
53			
54	ZRB6.00	Read	diabetes wellbeing questionnaire
55	ZRB6.11	Read	dwbq - diabetes wellbeing questionnaire
56			
57	ZRBa.00	Read	education score - diabetes
58			
59	ZRbH.00	Read	perceived control of insulin-dependent diabetes
60			

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ZV65312	Read	[v]dietary counselling in diabetes mellitus
ZV6DA00	Read	[v]admitted for commencement of insulin
ZV6DB00	Read	[v]admitted for conversion to insulin
E10	ICD-10	insulin-dependent diabetes mellitus
E10.0	ICD-10	insulin-dependent diabetes mellitus
E10.1	ICD-10	insulin-dependent diabetes mellitus
E10.2	ICD-10	insulin-dependent diabetes mellitus
E10.3	ICD-10	insulin-dependent diabetes mellitus
E10.4	ICD-10	insulin-dependent diabetes mellitus
E10.5	ICD-10	insulin-dependent diabetes mellitus
E10.6	ICD-10	insulin-dependent diabetes mellitus
E10.7	ICD-10	insulin-dependent diabetes mellitus
E10.8	ICD-10	insulin-dependent diabetes mellitus
E10.9	ICD-10	insulin-dependent diabetes mellitus
E11	ICD-10	non-insulin-dependent diabetes mellitus
E11.0	ICD-10	non-insulin-dependent diabetes mellitus
E11.1	ICD-10	non-insulin-dependent diabetes mellitus
E11.2	ICD-10	non-insulin-dependent diabetes mellitus
E11.3	ICD-10	non-insulin-dependent diabetes mellitus
E11.4	ICD-10	non-insulin-dependent diabetes mellitus
E11.5	ICD-10	non-insulin-dependent diabetes mellitus
E11.6	ICD-10	non-insulin-dependent diabetes mellitus
E11.7	ICD-10	non-insulin-dependent diabetes mellitus
E11.8	ICD-10	non-insulin-dependent diabetes mellitus
E11.9	ICD-10	non-insulin-dependent diabetes mellitus
E13	ICD-10	other specified diabetes mellitus
E13.0	ICD-10	other specified diabetes mellitus
E13.1	ICD-10	other specified diabetes mellitus
E13.2	ICD-10	other specified diabetes mellitus
E13.3	ICD-10	other specified diabetes mellitus
E13.4	ICD-10	other specified diabetes mellitus
E13.5	ICD-10	other specified diabetes mellitus
E13.6	ICD-10	other specified diabetes mellitus
E13.7	ICD-10	other specified diabetes mellitus
E13.8	ICD-10	other specified diabetes mellitus
E13.9	ICD-10	other specified diabetes mellitus
E14	ICD-10	unspecified diabetes mellitus
E14.0	ICD-10	unspecified diabetes mellitus
E14.1	ICD-10	unspecified diabetes mellitus
E14.2	ICD-10	unspecified diabetes mellitus
E14.3	ICD-10	unspecified diabetes mellitus
E14.4	ICD-10	unspecified diabetes mellitus
E14.5	ICD-10	unspecified diabetes mellitus
E14.6	ICD-10	unspecified diabetes mellitus
E14.7	ICD-10	unspecified diabetes mellitus

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2	E14.8	ICD-10	unspecified diabetes mellitus
3	E14.9	ICD-10	unspecified diabetes mellitus
4	G59.0	ICD-10	diabetic mononeuropathy
5	G63.2	ICD-10	diabetic polyneuropathy
6	H28.0	ICD-10	diabetic cataract
7	H36.0	ICD-10	diabetic retinopathy
8	M14.2	ICD-10	diabetic arthropathy
9	N08.3	ICD-10	glomerular disorders in diabetes mellitus
10	Y42.3	ICD-10	insulin and oral hypoglycaemic [antidiabetic] drugs
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Death
Date of death registered in CPRD

The RECORD statement – checklist of items, extended from the STROBE statement, that should be reported in observational studies using routinely collected health data.

	Item No.	STROBE items	Location in manuscript where items are reported	RECORD items	Location in manuscript where items are reported
Title and abstract					
	1	(a) Indicate the study's design with a commonly used term in the title or the abstract (b) Provide in the abstract an informative and balanced summary of what was done and what was found	a) 1, 3 b) 3	RECORD 1.1: The type of data used should be specified in the title or abstract. When possible, name of the databases used should be included. RECORD 1.2: If applicable, the geographic region and time frame within which the study took place should be reported in the title or abstract. RECORD 1.3: If linkage between databases was conducted for the study, this should be clearly stated in the title or abstract.	1.1) 1, 3 1.2) 1, 3 1.3) 3
Introduction					
Background rationale	2	Explain the scientific background and rationale for the investigation being reported	5		
Objectives	3	State specific objectives, including any prespecified hypotheses	5		
Methods					
Study Design	4	Present key elements of study design early in the paper	5-6		
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	5-7		

Participants	6	(a) <i>Cohort study</i> - Give the eligibility criteria, and the sources and methods of selection of participants. Describe methods of follow-up <i>Case-control study</i> - Give the eligibility criteria, and the sources and methods of case ascertainment and control selection. Give the rationale for the choice of cases and controls <i>Cross-sectional study</i> - Give the eligibility criteria, and the sources and methods of selection of participants (b) <i>Cohort study</i> - For matched studies, give matching criteria and number of exposed and unexposed <i>Case-control study</i> - For matched studies, give matching criteria and the number of controls per case	a) 5-7 b) Not applicable	RECORD 6.1: The methods of study population selection (such as codes or algorithms used to identify subjects) should be listed in detail. If this is not possible, an explanation should be provided. RECORD 6.2: Any validation studies of the codes or algorithms used to select the population should be referenced. If validation was conducted for this study and not published elsewhere, detailed methods and results should be provided. RECORD 6.3: If the study involved linkage of databases, computer use of a flow diagram or other graphical display to demonstrate the data linkage process, including the number of individuals with linked data at each stage.	6.1) 5-7 6.2) 5-6, 15 6.3) Not needed
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable.	5-7	RECORD 7.1: A complete list of codes and algorithms used to classify exposures, outcomes, confounders, and effect modifiers should be provided. If these cannot be reported, an explanation should be provided.	5-7
Data sources/ measurement	8	For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe comparability of assessment methods if there is more than one group	5-7		

Bias	9	Describe any efforts to address potential sources of bias	8-10		
Study size	10	Explain how the study size was arrived at	6-7		
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen, and why	5-10		
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding (b) Describe any methods used to examine subgroups and interactions (c) Explain how missing data were addressed (d) <i>Cohort study</i> - If applicable, explain how loss to follow-up was addressed <i>Case-control study</i> - If applicable, explain how matching of cases and controls was addressed <i>Cross-sectional study</i> - If applicable, describe analytical methods taking account of sampling strategy (e) Describe any sensitivity analyses	8-10		
Data access and cleaning methods		5-7		RECORD 12.1: Authors should describe the extent to which the investigators had access to the database population used to create the study population.	12.1) 5-6 12.2) 5-7

				RECORD 12.2: Authors should provide information on the data cleaning methods used in the study.	
Linkage		5-6		RECORD 12.3: State whether the study included person-level, institutional-level, or other data linkage across two or more databases. The methods of linkage and methods of linkage quality evaluation should be provided.	12.3) 5-6
Results					
Participants	13	(a) Report the numbers of individuals at each stage of the study (e.g., numbers potentially eligible, examined for eligibility, confirmed eligible, included in the study, completing follow-up, and analysed) (b) Give reasons for non-participation at each stage. (c) Consider use of a flow diagram	6, 10-13	RECORD 13.1: Describe in detail the selection of the persons included in the study (i.e., study population selection) including filtering based on data quality, data availability, and linkage. The selection of included persons can be described in the text and/or by means of the study flow diagram.	13.1) 6, 10-13
Descriptive data	14	(a) Give characteristics of study participants (e.g., demographic, clinical, social) and information on exposures and potential confounders (b) Indicate the number of participants with missing data for each variable of interest (c) Cohort study - summarise follow-up time (e.g., average and total amount)	10-11, Table 1		
Outcome data	15	Cohort study - Report numbers of outcome events or summary measures over time Case-control study - Report numbers in each exposure	10-13		

		category, or summary measures of exposure <i>Cross-sectional study</i> - Report numbers of outcome events or summary measures			
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (e.g., 95% confidence interval). Make clear which confounders were adjusted for and why they were included (b) Report category boundaries when continuous variables were categorized (c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	10-13		
Other analyses	17	Report other analyses done—e.g., analyses of subgroups and interactions, and sensitivity analyses	10-13		
Discussion					
Key results	18	Summarise key results with reference to study objectives	13		
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias	13-15	RECORD 19.1: Discuss the implications of using data that were not created or collected to answer the specific research question(s). Include discussion of misclassification bias, unmeasured confounding, missing data, and changing eligibility over time, as they pertain to the study being reported.	13-15
Interpretation	20	Give a cautious overall interpretation of results considering objectives,	18,19		

		limitations, multiplicity of analyses, results from similar studies, and other relevant evidence			
Generalisability	21	Discuss the generalisability (external validity) of the study results	18		
Other Information					
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	20		
Accessibility of protocol, raw data, and programming code		..	20, Appendix, protocol made available for reviewers	RECORD 22.1: Authors should provide information on how to access any supplemental information such as the study protocol, raw data, or programming code.	20, Appendix, protocol made available for reviewers

*Reference: Benchimol EI, Smeeth L, Guttman A, Harron K, Moher D, Petersen I, Sørensen HT, von Elm E, Langlois SM, the RECORD Working Committee. The REporting of studies Conducted using Observational Routinely-collected health Data (RECORD) Statement. *PLoS Medicine* 2015; in press.

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Correction: Adherence to guidelines for creatinine and potassium monitoring and discontinuation following renin-angiotensin system blockade: a UK general practice-based cohort study

Schmidt M, Mansfield KE, Bhaskaran K, *et al.* Adherence to guidelines for creatinine and potassium monitoring and discontinuation following renin-angiotensin system blockade: a UK general practice-based cohort study. *BMJ Open* 2017;**7**:e012818. doi: 10.1136/bmjopen-2016-012818

In the 3rd column of Table 5 the heading ‘Serum potassium increase $\geq 30\%^*$ ’ should read ‘Serum potassium increase >6 mmol/L’.

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BMJ Open 2017;**7**:e012818corr1. doi:10.1136/bmjopen-2016-012818corr1



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