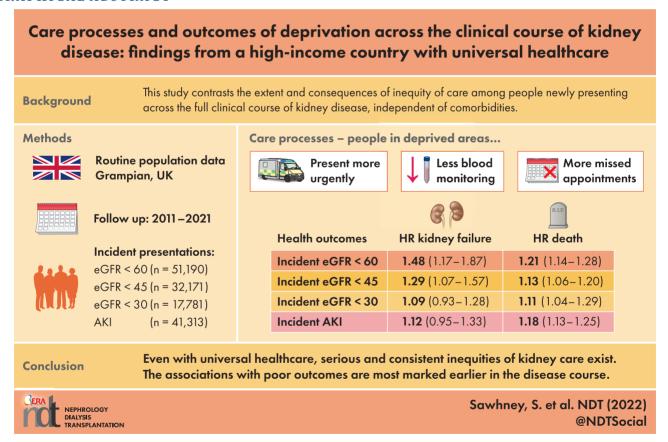


Care processes and outcomes of deprivation across the clinical course of kidney disease: findings from a high-income country with universal healthcare

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GRAPHICAL ABSTRACT



ABSTRACT

Background. No single study contrasts the extent and consequences of inequity of kidney care across the clinical course of kidney disease.

Methods. This population study of Grampian (UK) followed incident presentations of acute kidney injury (AKI) and incident estimated glomerular filtration rate (eGFR) thresholds of <60, <45 and <30 mL/min/1.73 m² in

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KEY LEARNING POINTS

What is already known about this subject?

- There is a recognised lack of evidence for inequity of kidney care that covers a full population across a full clinical course of kidney disease.
- Previous studies have shown associations between deprivation and adverse kidney health in snapshots of health datasets or have shown poorer outcomes in specific disease subsets only (e.g. reduced access to transplantation for kidney failure).

What this study adds?

- Across all incident patients in the Grampian population, healthcare in deprived areas was more reactive (more acute initial
 presentations and emergency hospital visits) and less proactive (less blood monitoring and outpatient attendances, less
 preventative prescribing).
- Associations with mortality and kidney failure were most severe earlier in the clinical course (newly presenting eGFR <60 mL/min/1.73 m² or AKI), in those of working age, and men.
- Age- and sex-adjusted associations were not substantially affected by any further adjustment of known medical factors such as comorbidities.

What impact this may have on practice or policy?

- Despite free universal healthcare in the UK, inequity of kidney care exists for people presenting at every stage of illness, but with particularly large inequity in the initial access to preventative care at early stages in the clinical course.
- An overly medical approach to addressing these inequities (without focusing on social needs, health literacy, behaviour or access) is likely to be insufficient to close this inequity gap.

separate cohorts (2011–2021). The key exposure was arealevel deprivation (lowest quintile of the Scottish Index of Multiple Deprivation). Outcomes were care processes (monitoring, prescribing, appointments, unscheduled care), long-term mortality and kidney failure. Modelling involved multivariable logistic regression, negative binomial regression and cause-specific Cox models with and without adjustment of comorbidities.

Results. There were 41 313, 51 190, 32 171 and 17 781 new presentations of AKI and eGFR thresholds <60, <45 and <30 mL/min/1.73 m². A total of 6.1-7.8% of the population was from deprived areas and (versus all others) presented on average 5 years younger, with more diabetes and pulmonary and liver disease. Those from deprived areas were more likely to present initially in hospital, less likely to receive community monitoring, less likely to attend appointments and more likely to have an unplanned emergency department or hospital admission episode. Deprivation had the greatest association with long-term kidney failure at the eGFR <60 mL/min/1.73 m² threshold {adjusted hazard ratio [HR] 1.48 [95% confidence interval (CI) 1.17-1.87]} and this association decreased with advancing disease severity [HR $1.09 (95\% \text{ CI } 0.93-1.28) \text{ at eGFR} < 30 \text{ mL/min}/1.73 \text{ m}^2)$, with a similar pattern for mortality. Across all analyses the most detrimental associations of deprivation were an eGFR threshold <60 mL/min/1.73 m², AKI, males and those <65 years of age.

Conclusions. Even in a high-income country with universal healthcare, serious and consistent inequities in kidney care exist. The poorer care and outcomes with area-level deprivation were greater earlier in the disease course.

Keywords: AKI, care processes, CKD, epidemiology, health inequalities, prognosis

INTRODUCTION

Equitable health service design for people with long-term illnesses such as kidney disease requires evidence on how inequalities vary across the full clinical course [1]. Time points of greatest relevance are those when improvements in preventative healthcare could still feasibly modify the disease course [2]. Addressing inequalities in integrated care is a core objective of primary care services [3, 4], but the administrative data used to inform policies are typically coarse, providing limited insights into disease severity or progression [5]. This limitation poses a challenge for determining where the most pressing missed opportunities in care exist [3, 4].

Kidney disease, represented by chronic kidney disease (CKD) stages and intervening episodes of acute kidney injury (AKI), is a long-term condition where clinical course and severity can be readily followed through serial blood tests. The population burden of kidney disease is substantial, including associations with multimorbidity, older age and a major effect on mortality, morbidity and healthcare costs [6, 7]. As <10% of those with CKD or AKI receive follow-up by a nephrologist [8–13], prioritisation must occur, with coordination primarily in the community. This means kidney care may be prone to inequitable population-level coordination, but unlike other long-term conditions, a nuanced study across its full clinical course and severity may be objectively possible using blood test data.

As described in a report into kidney care equity in the UK [14] and consistent with reports from other high-income countries [15–18], 'people from lower socio-economic groups are more likely to develop CKD ... and die earlier'. Nevertheless, existing literature has been limited to isolated snapshots of kidney disease focusing on disease subsets or recruited cohorts rather than following a whole population over time [19–22]. Examples are studies focusing on ethnicity and the prevalence

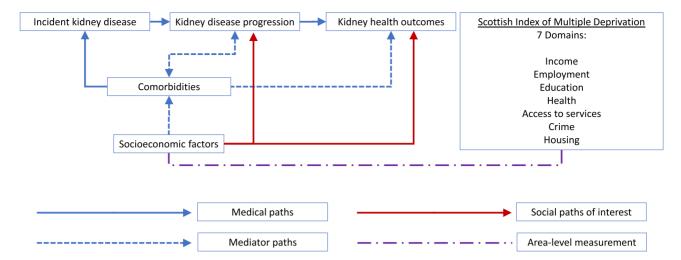


FIGURE 1: Logic model of assumptions for the bidirectional relationships between kidney disease clinical course, comorbidities and socio-economic factors.

of CKD or kidney failure and of access to transplantation [20, 21, 23-28]. The UK report concluded that there was insufficient evidence to make specific recommendations for intervention studies or policy [14, 29]. For instance, without a lens on the full clinical course within a single study, it would be hard to compare the extent of inequitable care of those newly presenting with mild impairment [estimated glomerular filtration rate (eGFR) <60 mL/min/1.73 m²] with those developing severe disease (eGFR < 30 mL/min/1.73 m²). Even in a country with free universal healthcare, such as the UK, recommended kidney care varies by severity: those with severe disease are prioritised for specialist nephrology care, whereas those with mild disease are often exclusively followed up in primary care. In addition, the relative contributions of social and medical factors towards overall health may shift as disease progresses, with medical factors predominating among those with more severe illness. Thus social deprivation may have a 'ceiling effect' among those with advancing kidney disease, but this has not previously been assessed.

The aim of this study was to assess, within the same population-level analysis, the extent to which a socio-economic gradient in care changes across the spectrum of kidney health and the implications for subsequent health outcomes and unscheduled care use. Specifically, we evaluated the care processes and outcomes of people from deprived and non-deprived backgrounds at different stages on a spectrum from mild to severe kidney impairment, independent of comorbidities. We hypothesised that inequitable care and outcomes for those from deprived areas would exist across all stages of kidney disease, but due to a ceiling effect, the magnitude of this inequity would be greater for those at earlier stages of kidney disease when the clinical course may also still be more modifiable.

MATERIALS AND METHODS

Modelling assumptions

Based on broader (non-kidney) chronic disease literature, a complex role exists for socio-economic factors (e.g. income,

employment, education, housing, access to care) in health and healthcare, with food insecurity, financial strain and housing instability highlighted as priorities [30]. For this study, socioeconomic metrics were obtained from the Scottish Index of Multiple Deprivation (SIMD), which covers seven area-level domains of income, employment, education, health, access to services, crime and housing. Socio-economic factors may associate with worse subsequent kidney disease and other comorbidities, but bidirectionally also represent consequences of ill health (Figure 1). For this analysis we assumed the former predominated (forwards directional arrows). We tested this association at different levels of kidney disease severity with and without adjustment of comorbidities as a mediator. We addressed the analysis in stages by assessing the variation in 1year mortality by area; 1-year healthcare processes, kidney failure and mortality by lowest quintile of Scotland for area-level deprivation versus all other levels; long-term kidney failure and mortality by lowest quintile for area-level deprivation versus other levels and, in sensitivity analyses, we evaluated outcomes across all quintiles of area-level deprivation and tested for effect modification by age, sex and acuity of presentation (i.e. evidence of prior or concurrent AKI within the prior 90 days).

Population

The Grampian Laboratory Outcomes Morbidity and Mortality Study provides routine health data for all adult residents in Grampian (adult population 473 226) [9, 31–33]. The study includes complete capture and data linkage (2009–2021 for this analysis) of all hospital admission episodes [Scottish Morbidity Record 01 (SMR01)] [34], emergency department visits (Accident and Emergency Data Mart) [35], outpatient clinic episodes [Scottish Morbidity Record 00 (SMR00)] [34], blood and urine tests from all locations [inpatient, community including primary care or outpatient, National Health Service (NHS), private practice] and prescribed medications [Prescribing Information System (PIS)] [36].

This analysis involved four subset cohorts nested within the Grampian population containing all new presentations of

kidney disease in Grampian between January 2011 and December 2018. In the main analysis the four cohorts represented the first (incident) instance when the eGFR crossed thresholds of above to below 60, 45, 30 ml/min/1.73 m², respectively, and the first (incident) instance of AKI among those with evidence of no prior presentation in 2009-2010. To reflect the real world in which high-risk patients may deteriorate and cross thresholds quickly, these cohort criteria were enabled to be non-mutually exclusive—a person presenting with either a rapid or unmonitored decrease in eGFR to 40 ml/min/1.73 m² could belong to both the <60 and <45 eGFR cohorts, or only the <45 cohort if there was previous evidence of an eGFR of 45-60 ml/min/1.73 m². Similarly, in the main analysis, copresentation with AKI was allowed. In a sensitivity analysis, cohorts were made mutually exclusive by excluding those presenting concurrently with AKI from all other groups and by excluding those progressing rapidly (or unmonitored) through higher eGFR thresholds by adding lower boundaries (45-60, 30-44, <30 ml/min/1.73 m²).

To ensure we studied only new (incident) presentations of kidney impairment without a 'prevalent pool effect' (i.e. inadvertent mixing of prevalent and incident patients) we excluded patients with prior evidence of kidney impairment at the respective subset threshold in 2009 and 2010. We also recognised *a priori* that a narrow interpretation of Kidney Disease: Improving Global Outcomes (KDIGO) criteria with repeat testing over at least 90 days could lead to underascertainment of kidney disease among those with poor access to blood testing (a care process outcome of study that would thus lead to selection bias). To mitigate against this potential selection bias of testing access, we focused on first presentation of kidney impairment at the eGFR thresholds outlined above without requirement for confirmatory testing.

Exposure and comparator

Area-level deprivation was determined based on people living in the lowest quintile of Scotland for the SIMD from 2016 [37]. In statistical analyses, we compared people from deprived areas (lowest quintile of the SIMD) with those from all other non-deprived areas (all other quintiles of the SIMD). Specifically for data visualisation, we grouped people by residential areas, distinguishing areas containing the lowest quintile of the SIMD as 'most deprived', areas of affluence containing the top two quintiles of the SIMD as 'most affluent' and the remaining areas as neither deprivation nor affluence.

Covariates of interest

To determine eGFR, we used the Chronic Kidney Disease Epidemiology Collaboration equation excluding the coefficient for black race [38]. Additionally, the first (incident) instance of AKI was determined using a KDIGO-based AKI algorithm (changes in creatinine of 0.3 mg/dL within 48 hours, a 50% increase within 7 days or a 50% increase from a baseline creatinine estimated from the median of creatinine levels over the previous 90 and 365 days), as described in greater detail elsewhere [9, 31–32]. Additional covariates adjusted in models

were age, sex and comorbidities (cardiovascular disease, coronary heart disease, heart failure, stroke, atrial fibrillation, peripheral arterial disease, diabetes mellitus, hypertension, cancer, liver disease and chronic pulmonary disease) based on previously recorded International Classification of Diseases, 10th Revision hospital episode codes using validated coding dictionaries as listed elsewhere [9, 39–41].

Outcomes

We studied kidney failure and mortality for all people and care processes for surviving at least 1 year from initial presentation. We evaluated mortality at 1 and 3 years and over the long term with the last follow-up on 1 July 2021 (up to 10 years). Onset of kidney failure was either sustained eGFR $<15~\text{mL/min}/1.73~\text{m}^2$ for at least 90 days on at least two blood tests and all intervening tests or onset of chronic kidney replacement therapy (dialysis or transplant).

Care processes assessed were the context of the initial presentation (inpatient or community, from SMR01 episode dates), outpatient serum creatinine test within 90 days, urine protein assessment within 1 year for those with a history of diabetes mellitus (in accordance with screening guidelines) [42], renin–aldosterone–angiotensin system (RAAS) blocker prescription (British National Formulary subsection code 020505 in the PIS) within 1 year among those with a history of myocardial infarction or heart failure (as per UK clinical guidelines) [43, 44], kidney care clinic referral within 1 year (SMR00 specialty code AG for renal medicine) [34], non-attendance at the clinic among those referred to a clinic, emergency department use within 1 year and unscheduled (emergency) hospital admission within 1 year of presentation (SMR01) [34].

Statistical analysis

We reported the number and characteristics of people from deprived areas versus all other areas. To visualise the overall extent of area-level variation and potential contribution of deprivation (and affluence) on mortality at each point in the clinical course, we plotted age- and sex-adjusted 1-year mortality estimates across geographic areas in each kidney subset (AKI and eGFR 45–60, 30–44 and <30 mL/min/1.73 m 2). We calculated estimates using a multilevel logistic regression model with random intercepts for resident area and age and sex as fixed effects using the 'melogit' package in Stata/SE 16 (StataCorp, College Station, TX, USA) [45].

Next, to quantify the influence of deprivation (worst quintile of SIMD versus all other areas) on early outcomes, we used multivariable logistic regression to calculate odds ratios (ORs) for the association of deprivation on mortality adjusting for age and sex and listed comorbidities for all patients. We used the same approach for each of the care processes listed in the outcomes section above. For processes reported as counts (number of emergency department attendances, total unscheduled inpatient care days), data were overdispersed and analysed using negative binomial regression with confidence intervals (CIs) derived from robust standard errors.

Finally, to assess the long-term risks of kidney failure and death (competing outcomes), we fitted long-term cause-specific Cox models for those with and without deprivation. Based on our *a priori* logic model (Figure 1), we theorised that comorbidities were a mediator of poor health outcomes. Accordingly, we fitted two models: one with age and sex adjustment and one with additional adjustment for comorbidities (to determine the age- and sex-adjusted role of deprivation, independent of known existing comorbidities).

We recognise that the influence of socio-economic factors may span the full spectrum from deprivation to affluence; however, for clarity we focused on deprived areas versus all other areas in the main analysis. In sensitivity analyses we included deprivation within five categories of SIMD quintiles to determine whether any association of deprivation with health outcomes was present only for those in the deprived quintile versus all others or if there was a continuum of association from deprivation to affluence. We used likelihood ratio tests of nested models to evaluate interactions with age (working age <65 years versus elderly ≥65 years) and sex. Finally, as incident presentations could arise from both gradual and rapid deteriorations, we tested for an interaction with prior/concurrent AKI. Where the interaction *P*-value was <.1, we reported results separately in a supplementary table. All analyses were performed using Stata/SE 16 (StataCorp) [45].

RESULTS

Characteristics

Table 1 reports characteristics of people newly presenting between January 2011 and December 2018 with AKI and eGFR 45–60, 30–44 and <30 mL/min/1.73 m². Within these four subset cohorts there were 41 313, 51 190, 32 171 and 17 781 people, respectively. The respective proportions of people from deprived areas were 7.8%, 6.0%, 6.5% and 7.1%. Across cohorts, 54.4–57.7% were female and those from deprived areas presented on average 5 years younger. A greater proportion of those from deprived areas had diabetes, liver disease and pulmonary disease (Table 1).

Care processes

Table 2 reports the association of deprivation with care processes within each subset cohort. Across the spectrum of kidney disease, those from deprived areas (versus other areas) were more likely to first present in an inpatient setting rather than through outpatient monitoring. Those from deprived areas were less likely to have subsequent blood test monitoring to evaluate for chronicity. While there were similar levels of renal outpatient referral, those from deprived settings attended their planned appointments less often, but were more likely to attend the emergency department or have an unplanned admission over the following year.

Mortality

Table 3 reports the association of deprivation with 1-year and long-term health outcomes. For those presenting with an

eGFR <60 mL/min/1.73 m² there was a significant (age- and sex-adjusted) excess risk of death among those from deprived areas at 1 year [OR 1.33 (CI 1.20–1.47)], which remained after additional adjustment for comorbidities [adjusted OR 1.28 (CI 1.15–1.42)]. Excess mortality was also present over a long-term follow-up to 10 years [Cox model adjusted hazard ratio (HR) 1.21 (CI 1.14–1.28)]. At each time horizon the excess mortality was attenuated for eGFR <30 mL/min/1.73 m² [1-year adjusted OR 1.19 (CI 1.05–1.35); Cox model long-term adjusted mortality HR 1.11 (CI 1.04–1.19)].

This pattern was also notable in Figure 2A–C of area-level variation in 1-year mortality, with greater variation in mortality among those presenting with eGFR <60 mL/min/1.73 m² (Figure 2A) than eGFR <30 mL/min/1.73 m² (Figure 2C). The separation in estimated mortalities between deprived (red) and affluent (blue) areas was also most evident in eGFR <60 mL/min/1.73 m². The same pattern remained evident at 3 years (Supplementary Figure 1a–c) and with AKI (Supplementary Figure 2a, b).

Kidney failure

Table 3 also reports the long-term HRs of deprived areas (versus other areas) for kidney failure up to 10 years after initial presentation. The greatest excess of kidney failure was among those presenting with eGFR <60 mL/min/1.73 m 2 [HR 1.48 (CI 1.17–1.87)] and the association was attenuated among those presenting with an eGFR <30 mL/min/1.73 m 2 [HR 1.09 (CI 0.93–1.28)].

Sensitivity analyses

We evaluated associations of mortality and kidney failure with deprivation across all quintiles of the SIMD (Supplementary Table 1). Findings were materially consistent, with the greatest rate of kidney failure and death in the most deprived quintile and the lowest rate in the most affluent quintile. We detected no interactions with prior/recurrent AKI (all P-values >.1), but did detect significant interactions with age and sex. As reported in Supplementary Table 2, the associations between deprivation and kidney failure and death were greater among males than females and greater among those of working age (<65 years) than the elderly (≥65 years). Finally, we also created mutually exclusive cohorts by excluding patients with AKI from eGFR threshold cohorts and excluding high-risk individuals with rapid eGFR decreases from a higher eGFR threshold cohort. This led to smaller cohort sizes but not materially different findings, with poorer outcomes for those from deprived areas and the greatest impact in the eGFR 45-60 mL/min/1.73 m² group (Supplementary Table 3).

DISCUSSION

Health service design is most successful when health conditions of greatest burden can be targeted at time points that enable prevention or modification of the disease course [1]. Beyond focusing solely on snapshots of disease prevalence, a detailed whole population view over the clinical course is

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Table 1. Characteristics of people with new-onset kidney disease according to each of the four subset cohort definitions

,			,)e	eGFR <45 ml/min/1.73 m	[/min/1.7	73 m ²								
Characteristics	eGFR <	eGFR <60 ml/min/1.73 m ² ($n =$	1.73 m ² (<i>i</i>	$n = 51 \ 190$		$(n = 32\ 171)$	2 171)		eGFR <	$GFR < 30 \text{ ml/min/1.73 m}^2 (n = 17.781)$.73 m ² (<i>i</i>	n = 17 781		AKI(n =	Λ KI ($n = 41313$)	
	Depr	Deprived areas	All othe	her areas	Depri	Deprived areas	All ot	All other areas	Depri	Deprived areas	All ot	All other areas	Depriv	Deprived areas	All ot	All other areas
	и	%	и	%	и	%	и	%	и	%	и	%	и	%	и	%
Total	3135	6.1	48 055	93.9	2080	6.5	30 091	93.5	1257	7.1	16 524	92.9	3210	7.8	38 103	92.2
Female	1693	54.0	26 363	54.9	1125	54.1	16 468	54.7	655	52.1	8876	53.7	1724	53.7	20 619	54.1
Ethnicity																
White	861	27.5	13 512	28.1	546	26.3	8167	27.1	311	24.7	4365	26.4	825	25.7	10 454	27.4
Non-white	33	1.1	267	9.0	15	0.7	130	0.4	10	8.0	65	0.4	46	1.4	296	8.0
Unknown	2241	71.5	24 276	71.3	1519	73.0	21 794	72.4	936	74.5	12 094	73.2	2339	72.9	27 353	71.8
Age (years), mean (95% CI)	29	66.4-67.6	71.2	71.1-71.4	70.3	69.7-71.0	75	74.8-75.1	72.4	71.5-73.2	76.7	76.5-76.9	63.1	62.4-63.7	69	68.8-69.1
AKI on presenting date	405	12.9	4746	6.6	435	20.9	5617	18.7	380	30.2	4802	29.1	3210	100	38 103	100
Atrial fibrillation	254	8.1	3968	8.3	265	12.7	4335	14.4	229	18.2	3223	19.5	421	13.1	5825	15.3
Cancer	352	11.2	6388	13.3	321	15.4	5282	17.6	216	17.2	3328	20.1	545	17.0	8135	21.4
Chronic obstructive pulmonary disease	563	18.0	5509	11.5	464	22.3	4438	14.7	295	23.5	2798	16.9	795	24.8	0029	17.6
Coronary heart disease	319	10.2	4171	8.7	302	14.5	3933	13.1	235	18.7	2852	17.3	446	13.9	5472	14.4
Diabetes	407	13.0	4575	9.5	381	18.3	4715	15.7	306	24.3	3507	21.2	287	18.3	6757	17.7
Heart failure	178	5.7	2249	4.7	209	10.0	2759	9.2	205	16.3	2534	15.3	316	8.6	4121	10.8
Hypertension	884	28.2	12 862	26.8	827	39.8	12 020	39.9	614	48.8	8043	48.7	1159	36.1	15 327	40.2
Liver disease	206	9.9	1469	3.1	168	8.1	1296	4.3	133	10.6	947	5.7	284	8.8	2029	5.3
Peripheral arterial disease	170	5.4	1949	4.1	173	8.3	2055	8.9	147	11.7	1596	9.7	277	8.6	3028	7.9
Stroke	179	5.7	2491	5.2	178	8.6	2531	8.4	135	10.7	1786	10.8	287	8.9	3648	9.6

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Table 2. Healthcare processes for new-onset kidney disease in each of the subset cohorts

	3)	OR (95% CI)	1.15 (1.06–1.25)	0.84 (0.77-0.92)	0.95 (0.77–1.18)	0.81	1.06 (0.88–1.28)	1.49 (1.34–1.66)
	AKI $(n = 41\ 313)$	Events/cases Events/cases (deprived), not deprived), n/N	24 564/38 103	1370/2346 17 383/27 241	3110/4671	2691/4649	1429/27 241	3909/27 241
		Events/case: $(deprived)$, n/N	2096/3210	1370/2346	268/412	211/371	135/2346	540/2346
	(n = 17.781)	OR (95% CI)	1.27 (1.12–1.43)	0.80 (0.68-0.95)	1.18 (0.83–1.68)	0.92	0.92 (0.74–1.14)	1.40 (1.15–1.70)
nix adjusted)	eGFR <30 ml/min/1.73 m ² ($n = 17.781$)	Events/cases (not deprived), n/N	9271/16 524	7890/10 435	1617/2247	1288/2300	1461/10 435	1316/10 435
ner areas (case n	eGFR <30 m	Events/cases (deprived), (n/N)	800/1257	562/769	145/189	95/170	131769	156/769
Deprived areas versus all other areas (case mix adjusted)	eGFR <45 ml/min/1.73 m ² ($n = 32 \text{ 171}$)	OR (95% CI)	1.22 (1.11–1.34)	0.87 (0.87-0.98)	1.07 (0.82–1.41)	0.79 (0.61–1.03)	1.04 (0.84–1.28)	1.60
Deprived ar	/min/1.73 m	Events/cases (not deprived), n/N	13 318 /30 091	15 757 /22 914	2472 /3488	2409	1289 /22 914	2203 /22 914
	eGFR <45 ml	Events/cases (deprived), n/N	1075/2080	1046/1542	208/285	167/267	122/1542	272/1542
	$(n = 51 \ 190)$	OR (95% CI)	1.25 (1.16–1.36)	0.97	0.87	0.91	0.88 (0.70–1.10)	1.74 (1.55–1.95)
	eGFR < 60 ml/min/1.73 m ² ($n = 51$	events/cases Events/cases (deprived), (not deprived), (n/N	15 055/48 055	1596/2636 24 460/41 660	2603/3692	2939/4264	1199/41 660	3169/41 660
	eGFR <60 m	Events/cases (deprived), n/N	1232/3135	1596/2636	223/331	220/315	93/2636	400/2636
		sses (among unless	Presents first as an inpatient (among all cases)	Outpatient blood monitoring within 90 days	Diabetes urine protein screen within 1 year	RAAS prescription after MI or heart failure within 1 year	Specialty renal clinic referral within 1 year	At least one specialty clinic non-attendance within 1 year
		Healthcare processes (a 1-year survivors unless otherwise stated)	Context	Monitoring			Referrals	

Table 2 Continued

	eGFR < 60 n	$_{\text{eGFR}} < 60 \text{ m}/\text{min}/1.73 \text{ m}^2 (n = 51)$	190)	eGFR <45 m	1/min/1 73 m	eGFR < 45 ml/min/1 73 m² (n = 32.171) eGFR < 30 ml/min/1 73 m² (n = 17.781)	Deptived at east versus at 10 the 1 areas (case 111x adjusted) finite/[73 m^2 ($n = 32.171$) eGFR $< 30 \text{ m}/\text{min}/[73 \text{ m}^2$	//////////////////////////////////////	'n = 17 781)		AKI (n = 41313)	
										'		
Healthcare processes (among F 1-year survivors unless otherwise stated)	Events/cases (deprived), n/N	Events/cases Events/cases (deprived), (not deprived), n/N	OR (95% CI)	Events/cases (deprived), n/N	Events/cases (not deprived), n/N	; OR (95% CI)	Events/cases (deprived), n/N	Events/cases Events/cases (deprived), (not deprived), n/N	OR (95% CI)	Events/cases (deprived), (n/N	Events/cases Events/cases (deprived), (not deprived), n/N	OR (95% CI)
Emergency department use within	414/1772	2511/27 521	1.44 (1.28–1.62)	252/1019	2963 /14 858	1.26 (1.09–1.47)	132/506	1585/6628	1.09 (0.89–1.35)	467/1518	4007/17 547	1.44 (1.28–1.62)
Unscheduled hospitalisation within 1 year	776/2636	9027/41 660	1.38 (1.26–1.51)	581/1542	7099 /22 914	1.25 (1.12–1.40)	350/769	4394/10 435	1.08 (0.93–1.26)	966/2346	9557/27 241	1.34 (1.23–1.46)
Emergency department days within	0.40 (1.08)	0.25 (0.83)	RR 1.45 (1.28-1.64)	0.40 (1.00)	0.30 (0.82)	RR 1.21 (1.04–1.41)	0.44 (1.11)	0.37	RR 1.10 (0.89-1.36)	0.61 (1.49)	0.41 (1.30)	RR 1.34 (1.17-1.52)
1 year, mean (SD) Unscheduled acute hospital days within 1 year, mean	5.1 (17.1)	3.7 (14.5)	RR 1.31 (1.13–1.51)	6.8 (19.1)	6.1 (18.6)	RR 1.11 (0.96–1.30)	10.2 (26.2)	9.1 (22.2)	RR 1.13 (0.94–1.35)	8.3 (23.2)	7.0 (19.7)	RR 1.28 (1.13–1.45)

MI, myocardial infarction; OR, odds ratio (from logistic regression); RR, rate ratio (from negative binomial regression with robust CIs). Emergency department data available from 2014 onwards
Statistically significant values are in bold.

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Table 3. Mortality and kidney failure outcomes for new-onset kidney disease in each of the subset cohorts

					D	eprived area	Deprived areas versus all other areas (case mix adjusted)	er areas (case	mix adjusted)				
		eGFR <60 ml/min/1.73	/min/1.73 m ² ($m^2 (n = 51 190)$	eGFR <	eGFR < 45 ml/min/1.73 m ² $(n = 32 \ 171)$		eGFR <30 ml	/min/1.73 m ²	AKI eGFR <30 ml/min/1.73 m ² $(n = 17.781)(n = 41.313)$	AKI $(n = 41313)$		
Health outcomes (among all patients)		Events/ cases Events/ (deprived), cases (not	Events/ , cases (not deprived), n/N	Estimate (95% CI)	Events/ Events/ cases cases (not (deprived), deprived), n/N n/N	Events/ cases (not deprived),	Estimate (95% CI)	Events/ cases (deprived),	Events/ cases (not deprived)	Estimate (Events/ Events/ cases cases (deprived),(not deprived),Estimate	Events/ cases ot deprived	,Estimate (95% CI)
1-year mortality OR	Age and sex		6395	1.33	538	7177	1.22	488	6809	1.21	864	10 862	1.18
(l'ogistic regression model)	adjusted	/3135		(1.20–1.47)	/2080	/30 091	(1.10-1.35)	/1257	/16 524	(1.07-1.37)	/3210	/38 103	(1.09-1.29)
	Fully adjusted	499	6395	1.28	538	7177	1.19	488	6809	1.19	864	10 862	1.20
	•	/3135	/48 055	(1.15-1.43)	/2080	/30 091	(1.06-1.32)	/1257	/16 524	(1.05-1.35)	/3210	/38 103	(1.09-1.31)
Long-term mortality HR Age and sex	Age and sex	1167	16 432	1.27	1094	15 712	1.18	823	11 044	1.15	1644	19 964	1.22
(Cox model)	adjusted	/3135	/48 055	(1.20-1.35)	/2080	/30 091	(1.11-1.26)	/1257	/16 524	(1.07-1.23)	/3210	/38 103	(1.16-1.28)
	Fully adjusted	1167	16 432	1.21	1094	15 712	1.13	823	11 044	1.11	1644	19 964	1.18
		/3135	/48 055	(1.14-1.28)	/2080	/30 091	(1.06-1.20)	/1257	/16 524	(1.04-1.19)	/3210	/38 103	(1.13-1.25)
Long-term kidney failure Age and sex	e Age and sex	80	716	1.50	115	11111	1.28	170	1854	1.07	153	1854	1.12
HR (cause-specific Cox adjusted model)	adjusted	/3135	/48 055	(1.19–1.90)	/2080	/30 091	(1.06-1.56)	/1257	/16 524	(0.91–1.26)	/3210	/38 103	(0.95-1.32)
	Fully adjusted	80	716	1.48	115	1111	1.29	170	1854	1.09	153	1854	1.12
		/3135	/48 055	(1.17-1.87)	/2080	/30 091	(1.07-1.57)	/1257	/16 524	(0.93-1.28)	/3210	/38 103	(0.95-1.33)

Covariates adjusted in models were age at first presentation, sex and comorbidities (cancer, coronary heart disease, heart failure, stroke, atrial fibrillation, peripheral arterial disease, diabetes mellitus, hypertension, liver disease and chronic pulmonary

Significant values is; bold type where statistically significant.

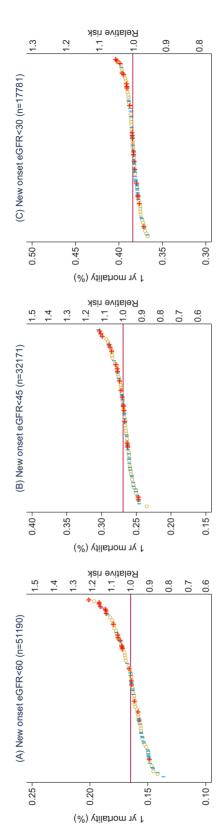


FIGURE 2: Age- and sex-standardised 1-year mortality risk of people with new-onset (A) eGFR <60 ml/min/1.73 m², (B) eGFR <45 ml/min/1.73 m² and (C) eGFR <30 ml/min/1.73 m² grouped by localities of deprivation (red "+"), affluence (blue "I"), and all other areas (yellow "0")

needed but is difficult to obtain. Here we provide populationlevel insights into the care and long-term outcomes attributable to socio-economic deprivation across the spectrum of the AKI/CKD clinical course. Those from deprived backgrounds developed kidney impairment on average 5 years younger and had a higher burden of comorbidities. For all people newly presenting with AKI or CKD at any severity threshold, we found greater mortality, kidney failure and use of unscheduled healthcare among those from deprived versus non-deprived areas. Even though healthcare in the UK is free at the point of use, uptake of healthcare in deprived areas was more reactive (more likely to present in an acute setting, more emergency hospital visits) and less proactive (e.g. less outpatient blood monitoring, less preventative prescribing). Despite the poorer health outcomes and greater comorbidities, there was no difference in renal outpatient clinic referral and those referred were substantially less likely to attend appointments.

Of particular importance, the greatest associations between deprivation and clinical outcomes (especially kidney failure) were early in the clinical course at first presentation with an eGFR <60 ml/min/1.73 m² or AKI. This is plausible because those with poorer kidney function may already have recognised advanced disease, whereby prevention may be less of an option. In addition, adjustment of comorbidities (a potential mediator of poor outcomes) made little difference to risk estimates. This implies that an overly medical approach to addressing inequities (without focusing on social needs, health literacy, behaviour or access) may be insufficient to close the inequity gap. One process we report here was the substantially increased clinic non-attendance in deprived areas. Adverse prognostic implications of clinic non-attendance have previously been reported for those attending diabetes clinics and driven by multifactorial reasons [46, 47]. Evidence of inequity in access to transplantation is also recognised [28], and poor health literacy has been associated with poorer kidney health outcomes in research mainly limited to kidney failure [48]. We also found the poorest outcomes among men and those of working age, which supports the need for further work understanding barriers to accessing kidney care. The extent to which these factors are causal and could be modified by an intervention is uncertain and the current lack of qualitative research is an acknowledged limiting factor [24].

Our broad finding of inequity of kidney care and outcomes is consistent with literature covering kidney diseases, cardiovascular disease and diabetes [14, 17, 21, 24]. The novel clinical course perspective of this analysis addresses a gap of existing literature [24] and spotlights a possible need for greater attention early (upstream) in the disease course and to non-medical factors. To restate a core public health principle, 'so long as we continue to fight the battle downstream...we are doomed to frustration, repeated failure, and perhaps ultimately to a sicker society' [2]. To determine the potential for interventions, further granular assessment of contributing non-medical factors should be guided by qualitative work to understand the lived experiences of those with early CKD and AKI. For instance, more than onethird of people in our analysis did not receive timely blood and urine monitoring in outpatient or community settings,

even though this is advocated in clinical guidelines for both groups [49, 50]. Other potential factors may include the implications of health literacy, lifestyle, trust, stigmatisation and how high-risk individuals engage with healthcare along potentially complex individual care pathways. A first step at presentation of CKD/AKI is the initial recognition and communication of its existence. Tensions exist here between early communication to promote self-management versus concerns about causing unnecessary anxiety or unnecessary disruption [51–53]. Collectively these factors will need to be considered within the context of existing practice [54] and primary care stakeholders planning the integration of community and outpatient care of those with long-term conditions and multimorbidity [3, 4].

Strengths of this study include a population approach with data capture of all in the region, in all contexts and with long recruitment and follow-up periods. The analysis included both outcomes and care processes to contrast outcomes at all levels of kidney severity, including AKI. Our analysis focused on care in a high-income country with integrated universal healthcare. Thus the stark findings are despite the availability of care in Scotland, which is based on clinical need rather than the ability to pay. Healthcare has evolved in Scotland, England, Wales and Northern Ireland, and each faces similar challenges, with rising demand, tight budget limits and clinicians in each nation following the National Institute for Health and Care Excellence referral guidelines for AKI and CKD. Thus our analysis can be expected to be generalisable for care in England, Wales and Northern Ireland [55]. However, it can be assumed that the challenges for countries with fragmented or private healthcare provision may be even greater.

Of note, in 2004 the Quality and Outcomes Framework (QOF), a pay-for-performance scheme, was introduced across the UK, which used a range of quality indicators to incentivise general practices to improve long-term condition management, practice organisation and patient experience of care [56]. Our analysis predates the coronavirus 2019 pandemic and the introduction of new policies in the new primary care contract in Scotland (in 2018) [4]. This included removal of the QOF in Scotland and a shift to investment in general practitioner clusters, professional groupings of general practices focused on improving quality of care to improve 'wellbeing, health and reduce health inequalities' [4]. Accordingly additional UK-wide analyses are warranted in the future.

A further limitation is the use of an area-level rather than individual-level deprivation measure. This may have led to underestimation of the full extent of the influence of deprivation because areas may contain residents from a mixture of backgrounds and social statuses, leading to bias towards the null. We also note that while we focused on socio-economic differences, we did not evaluate ethnic differences. As reported in Table 1, ethnicity is poorly recorded in administrative health datasets in Scotland, unlikely to be missing at random, and therefore not imputable. The importance of ethnicity is reported in detail in other studies, whereas in Grampian an overwhelming (>95% in the previous census) white European population predominates. Similarly, health literacy, migrant status, body weight, obesity, smoking

and alcohol intake are poorly recorded in hospital episode morbidity data and therefore not included in this analysis, but all would be relevant to understand health behaviour. Finally, our study used real-world data captured for clinical reasons rather than in a protocolised fashion. This is a strength because it reflects existing clinical practice, but it is not possible to interpret the clinical context and indications for blood test monitoring (which would be needed to more precisely understand and separate acute from chronic presentations and respective care requirements) nor disentangle the bidirectional relationship between kidney disease and accumulating non-kidney comorbidities, for which routine health data sources can provide only limited longitudinal information about disease severity at different time points.

In conclusion, by following the clinical course of kidney disease in a high-income country, we noted serious and consistent inequities of care and outcomes that were independent of comorbidities and greatest early in the disease course among those newly presenting with mild CKD and AKI. Greater understanding is now needed about how people initially present and how they access the support they require. This should include non-medical factors and incorporate lived experiences of those with early CKD or affected by AKI.

SUPPLEMENTARY DATA

Supplementary data are available at *ndt* online.

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AUTHORS' CONTRIBUTIONS

S.S. was responsible for the research idea and study design. S.S. and C.B. were responsible for data acquisition. S.S., D.Bl., N.F., S.M., M.R. and C.B. were responsible for data analysis/interpretation. S.S., D.Bo. and M.N. were responsible for statistical analysis. Each author contributed important intellectual content during manuscript drafting or revision, accepts personal accountability for the author's own contributions and agrees to ensure that questions pertaining to the accuracy or integrity of any portion of the work are appropriately investigated and resolved.

DATA AVAILABILITY STATEMENT

Datasets cannot be made available to other researchers due to ethical considerations. Information on how researchers may make requests to obtain similar datasets from the health research dataset custodians will be provided upon request.

CONFLICT OF INTEREST STATEMENT

All the authors have declared no competing interests.

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