# nature portfolio

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# **Reporting Summary**

Nature Portfolio wishes to improve the reproducibility of the work that we publish. This form provides structure for consistency and transparency in reporting. For further information on Nature Portfolio policies, see our <u>Editorial Policies</u> and the <u>Editorial Policy Checklist</u>.

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For	all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.
n/a	Confirmed
	$\square$ The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement
$\boxtimes$	A statement on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly
	The statistical test(s) used AND whether they are one- or two-sided  Only common tests should be described solely by name; describe more complex techniques in the Methods section.
	A description of all covariates tested
	🔀 A description of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons
	A full description of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) AND variation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)
	For null hypothesis testing, the test statistic (e.g. <i>F</i> , <i>t</i> , <i>r</i> ) with confidence intervals, effect sizes, degrees of freedom and <i>P</i> value noted <i>Give P values as exact values whenever suitable.</i>
$\boxtimes$	For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings
$\boxtimes$	For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes
$\boxtimes$	$\square$ Estimates of effect sizes (e.g. Cohen's $d$ , Pearson's $r$ ), indicating how they were calculated
	Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.

### Software and code

Policy information about availability of computer code

Data collection

Data extraction for this study was conducted using the Data Extraction for Epidemiological Research (DExtER) tool: Gokhale, K.M., et al. Data extraction for epidemiological research (DExtER): a novel tool for automated clinical epidemiology studies. Eur J Epidemiol 36, 165-178 (2021).

Data analysis

All analysis were conducted in Stata (StataCorp, Texas)

For manuscripts utilizing custom algorithms or software that are central to the research but not yet described in published literature, software must be made available to editors and reviewers. We strongly encourage code deposition in a community repository (e.g. GitHub). See the Nature Portfolio guidelines for submitting code & software for further information.

#### Data

Policy information about availability of data

All manuscripts must include a data availability statement. This statement should provide the following information, where applicable:

- Accession codes, unique identifiers, or web links for publicly available datasets
- A description of any restrictions on data availability
- For clinical datasets or third party data, please ensure that the statement adheres to our policy

Access to anonymized patient data from IMRD is subject to a data sharing agreement and protocol approval from the IMRD Scientific Review Committee. The study-specific analysable dataset is therefore not publicly available, however can be shared after obtaining approvals through contact with the corresponding

author (Prof Dipak Kotecha; d.kotecha@bham.ac.uk; 60-days response time for decisions). Details about IMRD applications and data access are available on the IQVIA Medical Research Data website.

### Research involving human participants, their data, or biological material

Policy information about studies with <u>human participants or human data</u>. See also policy information about <u>sex, gender (identity/presentation)</u>, and sexual orientation and race, ethnicity and racism.

Reporting on sex and gender

This study collected the biological sex of all participants included in the study using the designated category in the participants medical record. A participants chosen gender was not taken into account due to a lack of this information in UK primary care records. Adjusted models of analysis adjusted for sex difference between the two comparator groups.

Reporting on race, ethnicity, or other socially relevant groupings

Information about a patients socioeconomic status and ethnicity as these have the potential to impact on the rates of outcomes accessed in this study. Socioeconomic status was recorded as the Townsend deprivation index categorized into quintiles. Ethnicity was categorised into: (1) White Caucasian; (2) Black Afro-Caribbean; (3) South Asian; (4) Mixed or multiple ethnicities; and (5) Other ethnic group. Patients with missing Townsend and ethnicity data were aggregated into a separate missing category within that variable. Both variables were included in all adjusted analysis for each study outcome.

Population characteristics

Overall median age of 58.2 years (IQR 51.2-63.1), 18.0% women, 2.1% with diabetes and 16.5% with hypertension. See Table 1 for further details.

Recruitment

Participants were included in this study if they were registered at an eligible general practice for at least a year during the study period and met the trials inclusion and exclusion criteria. IMRD database encompasses 6% of the population of the UK, with a slight skew towards areas with younger and more affluent patients. Despite this, the prevalence of diseases is consistent with wider UK data, and is generalisable for demographics and major condition prevalence.5 The NHS is a publicly-funded healthcare system and individual clinicians do not receive personal funding for coded healthcare events.

Ethics oversight

Data collection for IMRD was approved by the National Health Service South-East Multicentre Research Ethics Committee in 2003. Under the terms of this approval, each study protocol undergoes independent review from the Scientific Review Committee (approval July 2017; reference number SRC 17THIN061).

Note that full information on the approval of the study protocol must also be provided in the manuscript.

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∑ Life sciences	Behavioural & social sciences	Ecological, evolutionary & environmental sciences

For a reference copy of the document with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>

### Life sciences study design

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Sample size No sample size calculation performed.

Data exclusions

No data was excluded from the final analysis instead missing data was categorised into a specific category to avoid creating bias in regression models and censorship occurred due to death or de-registration from their registered practice or due to ceasing of practice data contribution to IMRD

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Attempts at replication were successful - see 'Subgroup and Sensitivity analyses' section in main results for details.

Randomization

Replication

Patients with a recorded diagnosis of AF were included in the exposed cohort these. For each exposed patient, up to 4 patients without an AF diagnosis were randomly selected after matching for age  $(\pm 1 \text{ year})$ , sex and the health authority region of the patient's registered general practice. The same index date of the exposed patient was assigned to the corresponding matched unexposed patients to avoid immortal time bias

Blinding

Blinding is not relevant possible for this population-based, retrospective, matched open cohort study.

## Reporting for specific materials, systems and methods

We require information from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, system or method listed is relevant to your study. If you are not sure if a list item applies to your research, read the appropriate section before selecting a response.

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Materials & experime	ental systems	Methods
n/a Involved in the study		n/a Involved in the study
Antibodies		ChIP-seq
Eukaryotic cell lines		Flow cytometry
Palaeontology and a	archaeology	MRI-based neuroimaging
Animals and other o	organisms	•
Clinical data		
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Novel plant genotypes	NA	
Authentication	NA	