

1 **Association of Neighbourhood Deprivation with Risks of Major Amputation and Death**  
2 **Following Lower Limb Revascularisation**

3

4 Katriina Heikkilä<sup>1,2</sup>, Ian M. Loftus<sup>3</sup>, Sam Waton<sup>4</sup>, Amundeep S. Johal<sup>4</sup>, Jonathan R. Boyle<sup>5</sup>,  
5 David A. Cromwell<sup>1,4</sup>

6

7 <sup>1</sup>Department of Health Services Research and Policy, London School of Hygiene and Tropical  
8 Medicine, London, United Kingdom

9 <sup>2</sup>Finnish Institute for Health and Welfare, Helsinki, Finland

10 <sup>3</sup>St. George's Vascular Institute, St. George's Healthcare NHS Trust, London, United Kingdom

11 <sup>4</sup>Clinical Effectiveness Unit, the Royal College of Surgeons of England, London, United  
12 Kingdom

13 <sup>5</sup>Cambridge Vascular Unit, Cambridge University Hospitals, NHS Trust, Cambridge, United  
14 Kingdom

15

16 **Corresponding author:**

17 Katriina Heikkilä, Department of Health Services Research and Policy, London School of  
18 Hygiene and Tropical Medicine, 15-17 Tavistock Place, London WC1H 9SH, United Kingdom

19 Email: katriina.heikkila@thl.fi

20

21 **Word counts:** Abstract: 249, Manuscript (including figure and table legends): 1,326.

22 **Tables:** 1

23 **Figures:** 1

24 **Key words:** peripheral artery disease, revascularisation, amputation, neighbourhood  
25 deprivation, administrative data

26 **Abstract**

27 **Background and aims** Individual-level socioeconomic deprivation is associated with an  
28 increased risk of adverse patient outcomes following cardiovascular disease interventions, but  
29 the role of area-level socioeconomic circumstances as a predictor for treatment outcomes is  
30 unclear. we have examined the association of neighbourhood socioeconomic deprivation with  
31 risks of major lower limb amputation and death following surgical and endovascular lower  
32 limb revascularisation due to peripheral artery disease (PAD).

33 **Methods** Patients aged 50+ years who underwent surgical or endovascular lower limb  
34 revascularisation for PAD were identified from Hospital Episode Statistics, a nationwide  
35 hospital data warehouse in England. Major amputations and deaths within a year of  
36 revascularisation were ascertained from HES and national mortality register, respectively.  
37 Index of Multiple Deprivation (IMD) was used to measure neighbourhood deprivation.  
38 Flexible parametric competing risks models were used to estimate sub-distribution hazard  
39 ratios (SHRs) for amputation and death.

40 **Results** In all, 65,806 patients underwent endovascular and 20,072 underwent surgical  
41 revascularisation. The covariate-adjusted 1-year risk of major amputation was higher among  
42 patients from the most deprived compared to least deprived neighbourhoods following  
43 endovascular revascularisation (SHR: 1.24, 95% confidence interval, CI:1.10 to 1.38) and  
44 surgical revascularisation (SHR:1.28, 95% CI: 1.09 to 1.51). The risk of death was higher in  
45 most deprived compared to the least deprived neighbourhoods following both procedures.

46 **Conclusions** We found an association between neighbourhood deprivation with amputation  
47 and death outcomes following lower limb revascularisation for PAD. These findings suggest  
48 there may be opportunities for targeted interventions to improve care of PAD patients in  
49 deprived neighbourhoods.

50 **Introduction**

51 Socioeconomic circumstances, at individual- and area-levels, are important determinants of  
52 human health. Many manifestations of atherosclerotic cardiovascular disease, including  
53 coronary heart disease, heart failure and stroke, are socioeconomically patterned, with a higher  
54 prevalence and incidence in socioeconomically disadvantaged groups and areas (1-5).  
55 Individual- and area-level socioeconomic deprivation is also associated with an increased risk  
56 of developing peripheral artery disease (PAD)(6, 7).

57

58 Socioeconomic circumstances impact not only on the incidence of cardiovascular disease but  
59 also on patient outcomes. Studies from the United States and the Netherlands have shown that  
60 individual-level socioeconomic position, marked by low income, is associated with an  
61 increased risk of lower limb amputation among PAD patients (8-10). An association of low  
62 income with an increased risk of death following surgical intervention for PAD has also been  
63 reported in one study (10), but not corroborated in another (11). However, the extent to which  
64 area-level socioeconomic deprivation is implicated in patient outcomes in PAD is largely  
65 unknown. A study of US military veterans suggests that area deprivation is associated with an  
66 increased risk of major amputations among PAD patients (12), but the generalisability of the  
67 findings to other populations or healthcare systems not clear.

68

69 The aim of our investigation was to examine the association of neighbourhood deprivation with  
70 the risks of major lower limb amputation and death among patients undergoing endovascular  
71 or surgical revascularisation for lower limb PAD. To do this, we used data from Hospital  
72 Episode Statistics (HES), a data warehouse containing records of all patients admitted to  
73 National Health Service hospitals in England.

74

75 **Patients and methods**

76 Our analyses were based on data from women and men aged 50 years and older, who underwent  
77 endovascular (angioplasty with or without stent) or surgical (endarterectomy, profundaplasty  
78 or bypass) revascularisation for PAD between 1<sup>st</sup> Jan 2010 and 31<sup>st</sup> Dec 2015. The outcomes  
79 were major lower limb amputation (above the ankle; ipsilateral or contralateral) or death from  
80 any cause within one year of the revascularisation. The predictor of interest was neighbourhood  
81 deprivation, operationalised as the Index of Multiple Deprivation (IMD), a ranked score based  
82 on information on income, employment, education, health, crime, housing and the environment  
83 (13). Covariates were patient age and sex, indication for the intervention and the Royal College  
84 of Surgeons (RCS) Charlson comorbidity score (14).

85

86 All data were obtained from HES, apart from deaths, which were ascertained from the Office  
87 for National Statistics death register. Revascularisations and amputations were identified using  
88 a combination of International Classification of Disease (ICD) version 10 diagnostic codes  
89 indicating PAD and Office for Population Censuses and Surveys (OPCS) procedure codes  
90 indicating surgical or endovascular lower limb revascularisation (Online Supplement, Tables  
91 S1-S3). IMD was analysed as quintiles, from 1<sup>st</sup> (least deprived) to 5<sup>th</sup> (most deprived). The  
92 RCS Charlson score (0, 1 or 2+ comorbidities) and the indication for intervention (intermittent  
93 claudication or limb ischaemia *without* record of tissue loss, and severe limb ischaemia *with* a  
94 record of tissue loss) were defined using ICD-10 codes (Online Supplement, Tables S4 and  
95 S5). Flexible parametric competing risks regression was used to estimate sub-distribution  
96 hazard ratios (SHRs) for major amputation and death, with the other outcome as the competing  
97 risk (15). All analyses were conducted using Stata MP 15 (Stata Corporation, College Station,  
98 Texas, US).

99

100 This study is exempt from United Kingdom National Research Ethics Committee approval as  
101 it involved secondary analysis of anonymised data. HES data were made available by NHS  
102 Digital (Copyright© 2015).

103

## 104 **Results**

105 In all, 65,806 patients underwent endovascular and 20,072 underwent surgical lower limb  
106 revascularisation in 2010-2015. Patients' median age was 71 years (interquartile range: 63 to  
107 88); 65% were men. Endovascular revascularisations were slightly more common among  
108 patients living in affluent neighbourhoods and surgical interventions among those living in  
109 deprived neighbourhoods (Table). During the year after revascularisation, 4,937 patients  
110 underwent a major amputation (4.9% in the endovascular and 8.4% of those in the surgical  
111 group) and 9,682 died (11.0% and 12.2% in the endovascular and surgical groups,  
112 respectively). Following both revascularisation types, the SHRs for major amputation  
113 increased with greater neighbourhood deprivation, independently of covariates (Figure). The  
114 SHRs for death following either procedure were also higher among patients living in more  
115 deprived neighbourhoods (Figure). To explore potential interactions between procedure type  
116 and IMD, we modelled covariate-adjusted associations of procedure type with amputation and  
117 death, stratified by IMD quintile. In these analyses, the risks of amputation and death were  
118 consistently lower among patients undergoing endovascular revascularisation compared to  
119 those undergoing surgical procedures in all IMD quintiles (Table).

120

## 121 **Discussion**

122 Our findings, based on data from a nationwide, administrative dataset in England, suggest that  
123 neighbourhood socioeconomic deprivation is associated with an increased risk of major  
124 amputation following endovascular and surgical revascularisation alike. These observations

125 are in agreement with those reported in US military veterans and small case-control study of  
126 PAD patients at one district hospital in England, which suggest that neighbourhood deprivation  
127 is associated with an increased risk of major amputation in this patient group (12, 16).  
128 However, previous studies have not examined the association separately by revascularisation  
129 type.

130

131 The IMD is a weighted score, consisting of information on income (22.5% weight),  
132 employment (22.5%), education (13.5%), health and disability (13.5%), crime (9.3%), housing  
133 and services (9.3%) and living environment (9.3%). We hypothesise that the association  
134 between the quintiles of IMD and outcomes of lower limb revascularisation are mainly driven  
135 by the two key domains, income and employment. In a similar vein, it may be that our finding  
136 of endovascular revascularisation being slightly more common than surgical revascularisation  
137 among PAD patients living in less deprived neighbourhoods reflects these patients' ability to  
138 manage their disease better. This hypothesis is supported by the notion that patients from less  
139 deprived neighbourhoods undergo their first revascularisation at an older age and have a  
140 smaller number of comorbidities than those living in more deprived neighbourhoods (Table).

141

142 Previous research suggests that individual-level socioeconomic deprivation is a risk factor for  
143 adverse patient outcomes following revascularisation for PAD (8-10). Individual-level  
144 socioeconomic position, however, does not present an easy interventional target. Area-level  
145 socioeconomic circumstances, on the other hand, could be targets in themselves, or be used as  
146 indicators of areas where interventions, such as smoking cessation initiatives or diabetic  
147 footcare programmes to improve prognosis for PAD patients, should be directed. As our  
148 investigation is based on a register data, albeit from a nationwide dataset with good coverage  
149 of procedures and outcomes in England, our findings alone cannot form a basis for specific

150 policy or healthcare interventions. Together with the larger burden of disease and risk factors  
151 in deprived areas, as well as the later presentation to health services among PAD patients from  
152 deprived neighbourhoods (17), our findings and those of previous studies suggest that there  
153 might be opportunities for targeted health interventions to improve the care and outcomes for  
154 PAD patients living in deprived neighbourhoods.

155

156 An important strength of our research is that we used individual-level data from HES, which  
157 captures information on all revascularisation procedures conducted in NHS hospitals in  
158 England and has been shown to have good coverage, accuracy and completeness of  
159 cardiovascular procedure and outcome data for the study period. (18, 19) We used flexible  
160 parametric competing risks models to examine the risks of major amputation and death  
161 separately from one another, which is important in order to accurately estimate the risks of  
162 these competing outcomes. Unfortunately, HES does not include data on individual-level  
163 socioeconomic position, pack-years of smoking, obesity or physical activity, and we were  
164 therefore unable to examine their roles in the association between neighbourhood deprivation  
165 and outcomes of revascularisation. Previous studies of other cardiovascular outcomes have,  
166 however, demonstrated that area-level socioeconomic deprivation contributes to the risk of  
167 these outcomes independently of established clinical and behaviour-related risk factors and  
168 individual-level socioeconomic position (20, 21).

169

## 170 **Conclusion**

171 Our findings suggest that area deprivation is associated with an increased risk of major  
172 amputation following endovascular and surgical revascularisation.

173 **Conflict of interest**

174 None declared.

175

176 **Financial Support**

177 This study was undertaken as part of the work by the National Vascular Registry to evaluate  
178 the clinical outcomes achieved by English National Health Service vascular units. The National  
179 Vascular Registry is commissioned by the Healthcare Quality Improvement Partnership as part  
180 of the National Clinical Audit Program. The funders had no role in the study design, data  
181 collection or analysis, preparation of the manuscript or decision to publish the findings.

182

183 **Acknowledgements**

184 We thank David C. Mitchell for comments on an earlier version of the manuscript.



185 **References**

- 186 1. Addo J, Ayerbe L, Mohan KM, Crichton S, Sheldenkar A, Chen R, et al. Socioeconomic status  
187 and stroke: an updated review. *Stroke* 2012;**43**(4):1186-1191.
- 188 2. Galobardes B, Smith GD, Lynch JW. Systematic review of the influence of childhood  
189 socioeconomic circumstances on risk for cardiovascular disease in adulthood. *Ann Epidemiol*  
190 2006;**16**(2):91-104.
- 191 3. Manrique-Garcia E, Sidorchuk A, Hallqvist J, Moradi T. Socioeconomic position and incidence  
192 of acute myocardial infarction: a meta-analysis. *J Epidemiol Community Health* 2011;**65**(4):301-309.
- 193 4. Stringhini S, Viswanathan B, Gedeon J, Paccaud F, Bovet P. The social transition of risk  
194 factors for cardiovascular disease in the African region: evidence from three cross-sectional surveys  
195 in the Seychelles. *Int J Cardiol* 2013;**168**(2):1201-1206.
- 196 5. Vathesatogkit P, Batty GD, Woodward M. Socioeconomic disadvantage and disease-specific  
197 mortality in Asia: systematic review with meta-analysis of population-based cohort studies. *J*  
198 *Epidemiol Community Health* 2014;**68**(4):375-383.
- 199 6. Pujades-Rodriguez M, Timmis A, Stogiannis D, Rapsomaniki E, Denaxas S, Shah A, et al.  
200 Socioeconomic deprivation and the incidence of 12 cardiovascular diseases in 1.9 million women and  
201 men: implications for risk prediction and prevention. *PLoS One* 2014;**9**(8):e104671.
- 202 7. Vart P, Coresh J, Kwak L, Ballew SH, Heiss G, Matsushita K. Socioeconomic Status and  
203 Incidence of Hospitalization With Lower-Extremity Peripheral Artery Disease: Atherosclerosis Risk in  
204 Communities Study. *J Am Heart Assoc* 2017;**6**(8).
- 205 8. Durham CA, Mohr MC, Parker FM, Bogey WM, Powell CS, Stoner MC. The impact of  
206 socioeconomic factors on outcome and hospital costs associated with femoropopliteal  
207 revascularization. *J Vasc Surg* 2010;**52**(3):600-606; discussion 606-607.
- 208 9. Henry AJ, Hevelone ND, Belkin M, Nguyen LL. Socioeconomic and hospital-related predictors  
209 of amputation for critical limb ischemia. *J Vasc Surg* 2011;**53**(2):330-339 e331.
- 210 10. Ultee KHJ, Goncalves FB, Hoeks SE, Rouwet EV, Boersma E, Stolker RJ, et al. Low  
211 Socioeconomic Status is an Independent Risk Factor for Survival After Abdominal Aortic Aneurysm  
212 Repair and Open Surgery for Peripheral Artery Disease. *European Journal of Vascular and*  
213 *Endovascular Surgery* 2015;**50**(5):615-622.
- 214 11. Sinnamon AJ, Sonnenberg EM, Bartlett EK, Meise CK, Wang GJ, Kelz RR. The influence of  
215 socioeconomic factors on gender disparities in lower extremity bypass. *J Surg Res* 2014;**188**(2):537-  
216 544.
- 217 12. Arya S, Binney Z, Khakharia A, Brewster LP, Goodney P, Patzer R, et al. Race and  
218 Socioeconomic Status Independently Affect Risk of Major Amputation in Peripheral Artery Disease. *J*  
219 *Am Heart Assoc* 2018;**7**(2).
- 220 13. Centre HaSCI. Hospital Episode Statistics (HES). 2015 [cited 2015; Available from:  
221 <http://www.hscic.gov.uk/hes>
- 222 14. Armitage JN, van der Meulen JH, Royal College of Surgeons Co-morbidity Consensus G.  
223 Identifying co-morbidity in surgical patients using administrative data with the Royal College of  
224 Surgeons Charlson Score. *Br J Surg* 2010;**97**(5):772-781.
- 225 15. Mozumder SI, Rutherford MJ, Lambert PC. A flexible parametric competing-risks model using  
226 a direct likelihood approach for the cause-specific cumulative incidence function. *Stata Journal*  
227 2017;**17**(2):462-489.
- 228 16. Ferguson HJ, Nightingale P, Pathak R, Jayatunga AP. The influence of socio-economic  
229 deprivation on rates of major lower limb amputation secondary to peripheral arterial disease. *Eur J*  
230 *Vasc Endovasc Surg* 2010;**40**(1):76-80.
- 231 17. Schultz WM, Kelli HM, Lisko JC, Varghese T, Shen J, Sandesara P, et al. Socioeconomic Status  
232 and Cardiovascular Outcomes: Challenges and Interventions. *Circulation* 2018;**137**(20):2166-2178.

- 233 18. Kivimaki M, Batty GD, Singh-Manoux A, Britton A, Brunner EJ, Shipley MJ. Validity of  
234 Cardiovascular Disease Event Ascertainment Using Linkage to UK Hospital Records. *Epidemiology*  
235 2017;**28**(5):735-739.
- 236 19. Herrett E, Shah AD, Boggon R, Denaxas S, Smeeth L, van Staa T, et al. Completeness and  
237 diagnostic validity of recording acute myocardial infarction events in primary care, hospital care,  
238 disease registry, and national mortality records: cohort study. *BMJ* 2013;**346**:f2350.
- 239 20. Diez Roux AV, Merkin SS, Arnett D, Chambless L, Massing M, Nieto FJ, et al. Neighborhood of  
240 residence and incidence of coronary heart disease. *N Engl J Med* 2001;**345**(2):99-106.
- 241 21. Tunstall-Pedoe H, Woodward M, estimation Sgor. By neglecting deprivation, cardiovascular  
242 risk scoring will exacerbate social gradients in disease. *Heart* 2006;**92**(3):307-310.

243

244 **Figure legend**

245 Figure. Outcomes of lower limb revascularisation, by quintiles of Index of Multiple

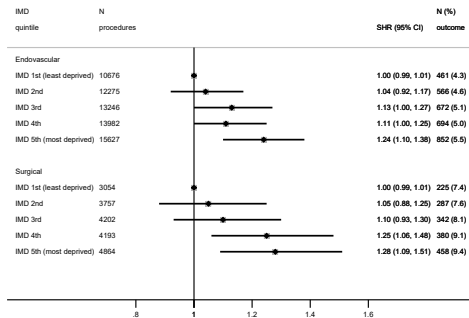
246 Deprivation: (A) Major amputation (B) Death from any cause

**Table. Procedures, patient characteristics and outcomes by IMD quintile**

<b>N (%) patients</b>	<b>IMD 1<sup>st</sup></b> <i>Least deprived</i>	<b>IMD 2<sup>nd</sup></b>	<b>IMD 3<sup>rd</sup></b>	<b>IMD 4<sup>th</sup></b>	<b>IMD 5<sup>th</sup></b> <i>Most deprived</i>	<b>p trend</b>
All patients	13,730	16,032	17,450	18,175	20,491	
<b>Procedure type</b>						
Endovascular (n=65,806)	10,676 (77.8)	12,275 (76.6)	13,246 (75.9)	13,982 (76.9)	15,627 (76.3)	0.023
Surgical (n=20,072)	3,054 (22.2)	3,757 (23.4)	4,202 (24.1)	4,193 (23.1)	4,864 (23.7)	0.023
<b>Patient characteristics</b>						
Age (mean, SD)	73.8 (10.1)	72.8 (10.2)	72.1 (10.3)	70.3 (10.4)	68.4 (10.3)	<0.0001
Women	4,944 (36.0)	5,752 (35.9)	6,303 (36.1)	6,373 (35.1)	7,014 (34.2)	<0.0001
Indication for intervention <sup>a</sup>						
IC/SLI <i>without</i> tissue loss	10,422 (75.9)	12,180 (76.0)	13,153 (75.4)	13,900 (76.5)	15,825 (77.2)	0.001
SLI <i>with</i> tissue loss	3,308 (24.1)	3,852 (24.0)	4,297 (24.6)	4,275 (23.5)	4,666 (22.8)	
RCS Charlson score						
0	8,239 (60.0)	9,248 (57.7)	9,851 (56.5)	10,226 (56.3)	11,331 (55.3)	<0.0001
1	4,175 (30.4)	5,028 (31.4)	5,690 (32.6)	5,892 (32.4)	6,817 (33.3)	
2+	1,316 (9.6)	1,756 (11.0)	1,909 (10.9)	2,057 (11.3)	2,343 (11.4)	
<b>Major lower limb amputation</b>						
Endovascular	461 (4.3)	566 (4.6)	672 (5.1)	694 (5.0)	852 (5.5)	<0.0001
Surgical	225 (7.4)	287 (7.6)	342 (8.1)	380 (9.1)	458 (9.4)	<0.0001
<b>Death from any cause</b>						
Endovascular	1,220 (11.4)	1,447 (11.8)	1,493 (11.3)	1,507 (10.8)	1,558 (10.0)	<0.0001
Surgical	399 (13.1)	486 (12.9)	507 (12.1)	481 (11.5)	584 (12.0)	0.1
<b>SHR<sup>b</sup> for amputation (95% CI)</b>						
Endovascular	0.57 (0.48 to 0.66)	0.57 (0.49 to 0.66)	0.59 (0.52 to 0.68)	0.53 (0.47 to 0.60)	0.58 (0.52 to 0.66)	
Surgical	1 (ref. cat.)	1 (ref. cat.)	1 (ref. cat.)	1 (ref. cat.)	1 (ref. cat.)	
<b>SHR<sup>b</sup> for death (95% CI)</b>						
Endovascular	0.78 (0.70 to 0.87)	0.81 (0.73 to 0.89)	0.85 (0.77 to 0.94)	0.84 (0.76 to 0.94)	0.78 (0.71 to 0.86)	
Surgical	1 (ref. cat.)	1 (ref. cat.)	1 (ref. cat.)	1 (ref. cat.)	1 (ref. cat.)	

248 <sup>a</sup> IC: intermittent claudication; SLI: severe limb ischaemia. <sup>b</sup> Adjusted for patient age, sex, indication for intervention and RCS Charlson score.

## (A) Major amputation



## (B) Death from any cause

