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POST-SERVICE LOWER LIMB AMPUTATION IN SCOTTISH MILITARY VETERANS

Short running title: Amputation in veterans

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ABSTRACT

Introduction

Recent attention has focussed on veterans who have lost limbs in conflict, but the number of UK veterans who lose limbs to disease is unknown. We used data from the Trends in Scottish Veterans' Health study to explore post-service lower limb amputation.

Methods

We carried out a retrospective cohort study of 78,000 veterans and 253,000 non-veterans born between 1945 and 1995, matched for age, sex and area of residence. We used survival analysis to examine the risk of amputation in veterans compared with non-veterans, and explored associations with antecedent disease.

Results

We found no difference between veterans and non-veterans in the risk of lower limb amputation, which was recorded in 145 (0.19%) veterans and 464 (0.18%) non-veterans, Cox proportional hazard ratio 1.00, 95% confidence intervals 0.82-1.20, $p=0.961$. Peripheral arterial disease was recorded in two-thirds of both veteran and non-veteran amputees, and type 2 diabetes in 41% of veterans and 33% of non-veterans, with a dual diagnosis in 32% of veterans and 26% of non-veterans. Trauma was an infrequent cause of amputation.

Conclusion

Although in later life veterans are no more likely to lose a limb to disease than non-veterans, the number so affected greatly outweighs those who have lost limbs in conflict. The high public profile of conflict-related limb loss risks eclipsing the needs of veterans with disease-related loss. Support for ageing veterans who have lost limbs due to disease will require

planning with the same care as that afforded to the victims of conflict if inequalities are to be avoided.

KEY MESSAGES

- In the long term, the number of veterans who undergo lower limb amputation for disease far exceeds the number who lose limbs in conflict.
- Military veterans are no more likely to experience lower limb amputation after leaving service than people who have never served.
- The commonest causes recorded for lower limb loss, in both veterans and non-veterans, are peripheral arterial disease followed by type 2 diabetes, accounting for a total of 66% and over 33% of cases respectively.
- Smoking-related diseases (ischaemic heart disease, lung cancer and peripheral arterial disease) are the commonest cause of death in amputees, irrespective of military service.
- Veterans who lose limbs to disease should be afforded the same level of support as combat-related amputees.

INTRODUCTION

Amputation of a limb or part thereof is one of the most visible reminders of the physical consequences of battle injury. Between 2001 and 2019, 301 UK service personnel underwent an amputation as a result of injuries sustained on operations in Afghanistan, with a further 32 as a result of service in Iraq between 2003 and 2019. An additional 142 service personnel underwent amputation for conditions sustained in other locations between 2014 and 2019. Whilst some of these amputations involved digits only, over 110 sustained multiple limb amputations, mainly from Afghanistan. Limb amputation does not automatically result in medical discharge from service; an individual who regains sufficient medical fitness may continue to serve if they so wish, subject to assessment by a Medical Board. However over 250 personnel who have sustained amputations in recent conflicts have been medically discharged and become veterans.(1)

Combat-associated trauma is not the only cause of limb amputation which military personnel and veterans may face.(2) Younger people may face amputation as a consequence of civilian trauma such as road traffic accidents, whilst both diabetes and peripheral arterial disease (PAD) are major causes of limb loss in later life.(3) A study of US veterans showed that 63% of amputations between 1989 and 1998 were for diabetes and 24% for peripheral vascular disease, with overall age-specific rates for veterans exceeding those in the general US population.(4, 5) We have shown that Scottish veterans are at increased risk of PAD compared with the wider population (6), but we had no information on whether this was associated with an increased incidence of limb loss.

Amputation carries substantial costs, both economically in terms of the costs associated with surgery, rehabilitation and prostheses, and personally in respect of loss of independence, altered body image, impact on the family and perhaps changes in employability.(3) It also has important implications for care needs, especially in older people,(7-9) but risks and antecedent causes of post-service amputations in UK veterans are unknown. We used data from the Scottish Veterans Health Study 5-year follow-up, Trends in Scottish Veterans' Health, to examine lower limb amputations in veterans in comparison with age, sex and geographically-matched people with no record of service.

METHODS

Trends in Scottish Veterans' Health is a retrospective cohort study of all 78,385 military veterans resident in Scotland who were born between 1 January 1945 and 31 December 1995, and a comparison group of 252,637 individuals with no record of service matched 3:1 for age, sex and postcode sector of residence (mean population 5,000). Veterans were eligible for inclusion if they were registered with National Health Service (NHS) Scotland both pre- and post-service. The study follows on from the Scottish Veterans Health Study, using similar methodology which is fully described elsewhere.⁽¹⁰⁾ Demographic data obtained from electronic NHS registration records were linked at an individual level to routine hospital admissions data (Scottish Morbidity Record SMR01), the Scottish Diabetes Register, and death certificates to provide information on all recorded lower limb amputations and first recorded diagnosis of diabetes (including type), PAD, trauma, and all-cause death. Anonymised data were also obtained from Blesma, The Limbless Veterans charity, on the lower limb amputees of whom they have record in Scotland. Although membership of Blesma is voluntary, it is likely to be reasonably complete in respect of military operational casualties. Dates of entering and leaving the Service, for veterans, were obtained from the Scottish NHS registration record. The maximum period of follow-up was from 1 January 1981 (or date of leaving the Armed Forces, for veterans, if later) to 31 December 2017. Data on the gender balance of the Armed Forces were taken from published Ministry of Defence statistics.⁽¹¹⁾ Individual-level prescribing data were obtained from the NHS Prescribing Information System for a limited range of drugs including nicotine replacement therapy (NRT). As we had no data on lifestyle risk factors, we used NRT prescribing as a proxy measure for smoking.⁽¹⁰⁾ The data extract was pseudo-anonymised and approval for the study was granted by the Public Benefit and Privacy Panel of the

Information Services Division of NHS Scotland. Use of anonymised extracts of Scottish routine data for health research is covered by generic NHS ethics approval (East of Scotland Research Ethics Committee; reference 16/ES/0112). As this was a secondary data study, individual consent was not required.

Socio-economic status

A measure of socio-economic status (SES) is provided by the Scottish Index of Multiple Deprivation (SIMD), which is based on 6,505 datazones, derived from postcode of residence, each having a mean population of 800. Deprivation status is calculated from information on income, employment, health, education (including skills and training), housing, crime, and access to services. The SIMD has been used to derive quintiles of SES for the Scottish population; ranging from 1 (most deprived) to 5 (least deprived).(12) The cohort participants were categorised according to these quintiles using postcode of residence.

Statistical methods

For the purposes of the study, lower limb amputation was defined as any occurrence of the OPCS Classification of Interventions and Procedures version 4 (OPCS-4)(13, 14) codes X01, X07, X08, X081, X09, X095, X11 or X12, including subcodes where appropriate, or OPCS-3 codes 870-875 or 862-864, and a diagnosis of diabetes was defined as ICD-10 E10-E14 or ICD-9 250 at any position in the NHS SMR01 record, or an entry in the Scottish Diabetes Register. For Scottish Diabetes Register entries, we restricted the analysis to type 2 diabetes. For SMR01 records, where type of diabetes was not recorded, we restricted the analysis to cases first diagnosed at age 30 or later to maximise the likelihood of capturing type 2 disease. PAD

was defined as ICD-10 I73-I79 or I702, or ICD-9 443.9 or 440.2. Trauma was defined as ICD-10 S00-S99, T00-T32 or ICD-9 E850-E858, E860-E869 or 960-989.4, and road traffic accident was defined as ICD-10 V00-V99 or ICD-9 E809-E819 or E820-E825.(15)

Cox proportional hazard models (16) were used to examine the association between veteran status and cumulative risk of amputation, overall and in association with specific antecedent conditions, using age as the time dependent variable, age at first record of amputation as the failure time, and death (if no amputation) as the censor time. A landmark analysis was performed using age 40 years as the starting point for PAD. We also examined comorbidity with mental health diagnoses where this was recorded on the inpatient or death record and, in particular, with severe stress or post-traumatic stress disorder (PTSD) (ICD-10 F43 or ICD-9 308 or 309). Hazard ratios and p values were calculated and the *a priori* rejection level was set at 0.05. Proportionality was tested using methodology based on Schoenfeld residuals.(17) The models were run univariably and then repeated adjusting for the potential confounding effect of SES. The analyses were repeated stratifying by 5-year bands of birth year, to examine birth cohort effects, and stratifying by length of service. All analyses were performed using Stata® v.16.

RESULTS

After data cleansing to remove incomplete and invalid records, 78,157 (99.7%) veterans and 252,637 (100.0%) non-veterans were included in the analysis. The dataset included 7,573 (9.7%) women veterans, reflecting the gender balance of the Armed Forces, and 28,983 (11.5%) non-veteran women. Health outcome data were available for a total of 10,643,252 person-years of follow-up.

During the period of follow-up, 145 (0.19%) veterans and 464 (0.18%) non-veterans underwent a lower limb amputation. The difference was not statistically significant, Cox proportional hazard ratio (HR) 1.00, 95% confidence intervals (CI) 0.82-1.20, $p=0.961$ (Figure 1). The median age at first record of amputation was 55.6 years (IQR 47.2-61.5) for veterans and 54.0 years (IQR 44.2-61.3) for non-veterans. Less than 5% (exact numbers suppressed) of those who had undergone lower limb amputation were women, with no significant difference between veteran and non-veteran women.

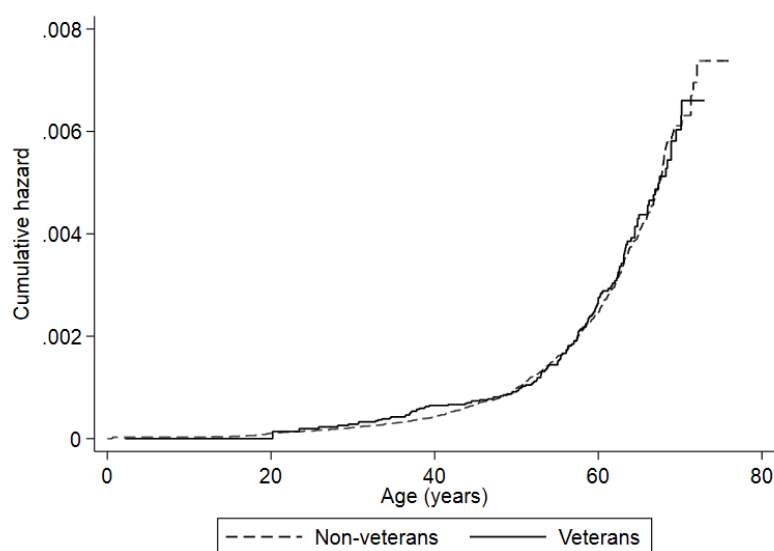


Figure 1. Nelson-Aalen cumulative hazard plot of risk of lower limb amputation in veterans compared with non-veterans

The commonest association with lower limb amputation was PAD, which was recorded in 95 (65.5%) amputations in veterans and 310 (66.8%) in non-veterans. Although PAD was more common overall in veterans than non-veterans (1.34% vs. 0.98%, HR 1.29, 95% CI 1.20-1.39, $p < 0.001$), the risk of progressing to limb loss in veterans was lower than in non-veterans, 9.0% of cases leading to amputation in veterans compared with 12.6% in non-veterans. The reduction in risk was statistically significant, HR 0.66, 95% CI 0.52-0.84, $p = 0.001$. The second most common associated diagnosis was type 2 diabetes, diagnosed in 61 (42.1%) of veteran amputations and 154 (33.1%) in non-veterans. The difference in risk was not statistically significant, HR 1.03, 95% CI 0.77-1.41, $p = 0.807$. There was a substantial overlap; 47 veterans (32.4% of all lower limb amputees) and 123 (26.5%) non-veterans had a dual diagnosis of PAD and type 2 diabetes. Again, the difference in risk was non-significant, HR 0.79, 95% CI 0.56-1.11, $p = 0.174$. Trauma (including road traffic accidents) was a very small contributor, accounting for just over 5% of amputations in veterans and under 5% in non-veterans (exact numbers suppressed); the differences were not statistically significant. Neither birth cohort (Figure 2) nor length of service (Figure 3) influenced risk of lower limb amputation, apart from a small peak at 10-12 years service. Being an Early Service Leaver did not increase the risk of post-service amputation.

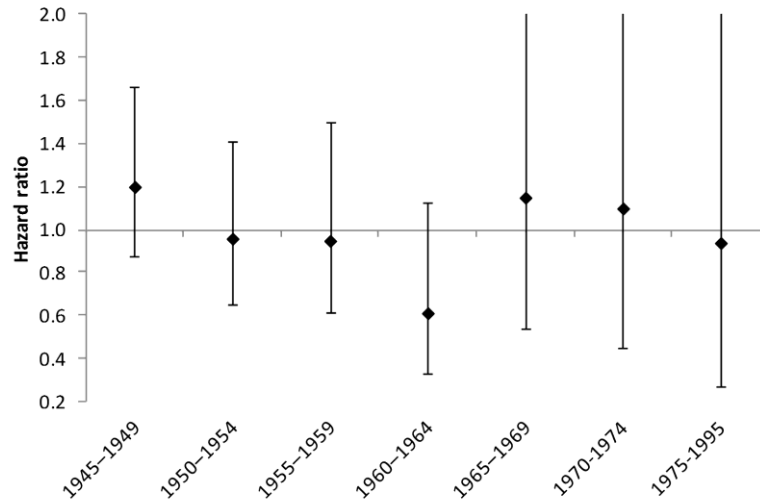


Figure 2. Hazard ratios for lower limb amputation by birth cohort, veterans vs. non-veterans. Error bars represent 95% confidence intervals

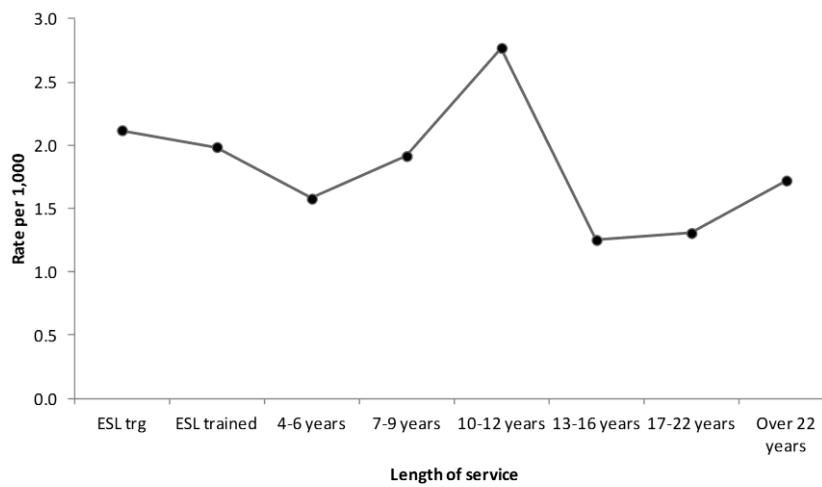


Figure 3. Lower limb amputation in veterans, crude rate by length of service
Categories represent common terms of military engagement

By the end of follow-up, 68 (46.9%) veterans who had undergone lower limb amputation had died, as had 211 (45.5%) non-veterans. The hazard ratio for all-cause mortality was

0.98, 95% CI 0.96-1.01, $p=0.288$. The commonest cause of death, for both veterans and non-veterans, was acute myocardial infarction or ischaemic heart disease, followed by lung cancer and peripheral arterial disease.

Blesma data showed 38 veterans living in Scotland with service-related lower limb amputations who would have met the age criteria for our study. Of these, 32 were aged under 50 and so were likely to have been associated with recent conflicts, whilst only 6 were older. Of the veterans registered with Blesma who had disease-related lower limb amputations (all ages), 67.7% were self-declared as due to PAD and 29.0% to diabetes.

We used NRT as a proxy for smoking prevalence; veterans were significantly more likely than non-veterans to have received a prescription for NRT, 8,812 (11.2%) veterans compared with 24,775 (9.8%) non-veterans, OR 1.15, 95% CI 1.12-1.17, $p<0.001$. However, NRT prescription rates for amputees did not differ between veterans and non-veterans, 17.2% for both.

There was an increased risk of amputation in veterans with a record of severe stress or PTSD (4.1% of veteran amputees compared with 2.8% of non-veterans), and with mood disorder (13.1% of veteran amputees compared with 9.5% of non-veterans), although neither achieved statistical significance owing to small numbers, OR 1.48, 95% CI 0.57-3.81, $p=0.419$ for stress/PTSD and OR 1.38, 95% CI 0.83-2.29, $p=0.211$ for mood disorder. There was no difference for anxiety (6.9% vs. 6.5%).

DISCUSSION

We have shown that in long-term follow-up (up to 57 years post-service), the risk of post-service lower limb amputation in veterans does not differ from non-veterans. Two-thirds of cases were associated with PAD in both veterans and non-veterans, whilst there was an association with type 2 diabetes in over 40% of veteran amputees and one-third of non-veterans. PAD in patients with type 2 diabetes was common. Perhaps surprisingly, trauma was a rare cause of post-service amputation; although this cohort overlaps the period of intense military operations in Iraq and Afghanistan, it appears that few of the in-service amputees who became veterans and settled in Scotland have needed to undergo further amputation. Nor do veterans appear to be at increased risk of traumatic injury leading to amputation, notwithstanding the well-documented association between deployment and risk of road traffic accident (18), and between military service and risky driving.(19, 20)

Although the number of veterans who settle in Scotland having sustained an in-service amputation is not formally recorded, a figure of approximately 10% of the UK-wide figures is widely used for planning purposes in various contexts since approximately 10% of the Armed Forces are recruited from Scotland.(21) Therefore it is likely that there are around 30-40 veteran amputees from recent conflicts living in Scotland. This estimate is supported by our analysis of the Blesma data. Prior to the start of hostilities in Iraq and Afghanistan, amputation as a consequence of military service would have been uncommon. Therefore, our finding that 145 veterans in Scotland had undergone post-service lower limb amputation, mostly due to disease, demonstrates that diabetes and peripheral vascular disease account for more limb loss in veterans than conflict.

Smoking is widely recognised as a risk factor for lower limb loss. Studies in the US have shown that veterans are more likely to smoke than non-veterans,(22) and UK studies in the 1960s and 1970s, when the older veterans in our study were serving, showed that serving soldiers were more likely to smoke, and to smoke more heavily, than civilians.(23, 24) Therefore it might be anticipated that amputation for vascular reasons is more common. We have shown that PAD is more common in veterans, both in our earlier study (6) and confirmed in this follow-up, but that paradoxically fewer amputations take place in veterans with PAD than non-veterans. We suggest that this may be due to earlier presentation, and hence greater likelihood of limb-saving treatment, as veterans may be less likely to ignore the symptoms of claudication than non-veterans.

The strength of this study is that we had access to follow-up data on over 330,000 people, including over 78,000 veterans, over a period of up to 37 years. The data were drawn from Scottish computerised healthcare records and so are likely to be complete in respect of surgical procedures carried out in NHS facilities. The use of record linkage to analyse individual level data directly derived from health records allowed a robust cohort study design to be employed. It was possible to do subgroup analyses by sex, birth cohort, and length of service, contributing to the understanding of factors influencing veterans' health.

Some loss to follow-up of subjects is likely to have occurred due to migration away from Scotland. Such losses could not be quantified and the assumption has been made that there was no systematic difference in this respect between veterans and non-veterans. No data were available for smoking rates so we have inferred likelihood of smoking from other

studies, and from NRT prescribing data. We had no data on the Service (Royal Navy, Army, Royal Air Force) to which a veteran belonged. Veterans with Reserve service only could not be identified from NHS records and were therefore included amongst the non-veterans; this would have had the effect of reducing any observed differences between veterans and non-veterans although the numbers involved are likely to be too small to have an important impact on our findings.

CONCLUSIONS

Lower limb amputation brings care needs, especially as amputees age when mobility and ability to manage a prosthesis become impaired. Recent attention has focussed on the long term needs of the amputee veterans of recent conflicts. We have shown that although in later life veterans are no more likely to lose a limb to disease than are people who have never served, the number of veterans so affected will greatly outweigh those who have lost limbs in conflict. The burden on care provision for ageing veterans is considerable, and will require planning with the same care as that afforded to the victims of conflict, who generally benefit from a higher profile and greater perceived public support, if major inequalities within the veteran community are to be avoided. This highlights an important aspect of the Armed Forces Covenant; notwithstanding the right to priority treatment for those injured in service, there should be no difference in the provision of long-term support (actual or perceived), either in comparison with the civilian community or within the veteran community itself.

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DECLARATION OF INTERESTS

BPB is a military veteran and Honorary Civilian Consultant Advisor (Army) in Veterans' Health and Epidemiology. Neither the Army nor the Ministry of Defence had any input to this paper, and the views and opinions expressed are solely those of the authors. No other interests are declared by the authors.

AUTHOR CONTRIBUTIONS

BPB conceived the idea and designed the study, with advice from JPP and DFM. BPB carried out the data analysis, which was overseen by DFM, and interpreted the findings. BPB wrote the first draft of the report, which was critically reviewed and edited by all authors. All authors approved the final article. BPB is the guarantor for overall content.

DATA SHARING

The study remains in progress and the data are not currently available for sharing.

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