

Toward Improving Prehospital Trauma Triage for Older People

Abdullah Alshibani^{1,2*}, Simon Conroy¹

¹ Department of Health Sciences, College of Life Sciences, University of Leicester, Leicester, United Kingdom; ² Emergency Medical Services Department, College of Applied Medical Sciences, King Saud bin Abdulaziz University for Health Sciences, Riyadh, Saudi Arabia.

***Correspondence to:** Mr. Abdullah Alshibani, Department of Health Sciences, College of Life Sciences, University of Leicester, Leicester LE1 7HA, UK; Tel: +44 (0)116 252 5992; email: anaa3@le.ac.uk.

Portrait photograph

Toward Improving Prehospital Trauma Triage for Older People

Abstract

Background: The proportions of older adults with major trauma is increasing. High-quality care for this population requires accurate and effective prehospital trauma triage decisions.

Objectives: Anatomical and physiological changes with age, comorbidities, and medications use for older adults may affect the accuracy of prehospital trauma triage.

Materials and methods: This narrative review focuses on age-related anatomical and physiological changes, comorbidities, and medications use for older adults with an emphasis on their impact on the accuracy of prehospital trauma triage tools. It also addresses efforts to develop alternative triage criteria to reduce under-triage.

Results: These factors were shown to affect physiological responses to injury and mechanism of injury for older people. Current triage tools poorly predicted injury severity. Geriatric-specific physiological measures and comorbidities significantly improved sensitivity with much lower specificity. Assessing anticoagulant or antiplatelet use in head injury notably improved sensitivity to identify traumatic intracranial haemorrhage, neurosurgery or death with modest decrease in specificity.

Conclusions: Improving paramedics' knowledge about the challenges when assessing older people with silver trauma may reduce under-triage. Assessing frailty could help in predict injury severity. Future research is needed to improve triage decisions for this population.

Keywords: Triage, Injury, Geriatrics, Paramedics, Emergency.

Introduction

Major trauma has historically been conceived as disease condition of younger populations, resulting from high-energy mechanisms of injury [19]; this thinking is reflected in education and training approaches [5]. However, the proportion of patients with major trauma aged ≥ 65 years has progressively increased in the last years [7]. In the United Kingdom (UK), the median age of trauma patients increased from 36.1 years

in 1990 to 53.8 years in 2013 [19]. Therefore, more age-attuned prehospital triage is needed.

Scope of the Problem

Prehospital trauma triage tools are developed to assess injury severity for patients to determine transportation decisions [38]. They are designed to assess the severity of a single event (i.e. injury), which is not the case when it comes to older people who are at greater risk of silver trauma (define/expand). Older trauma patients are significantly under-triaged in the prehospital phase [3, 6, 9, 22, 26, 27, 28, 33, 39]. This might be due to the process of aging, comorbidities, and medications use in older people which could affect the accuracy of prehospital triage [27]. This paper aims to describe the evidence around these factors and how they might affect the accuracy of prehospital trauma triage for older people in order to improve triage decisions for this population.

Age-Related Anatomical and Physiological Changes

Older adults undergo several anatomical and physiological changes with age. These changes in different body systems could affect their response to injury, which may negatively result in inappropriate assessment of injury severity.

For the cardiovascular system, the heart is subjected to structural changes like hypertrophy and sigmoid septum of the left ventricle [10], and also physiological changes such as reduced diastolic filling of the left ventricle [37] and prolonged contraction period [21]. Blood vessels also undergo structural changes including increased length of large vessels, enlarged lumen, and increased walls thickness [45]. This could impact their compliance and distensibility which is the main functional

change of vessels with age [43]. The aorta, for example, has decreased distensibility with age [43]. Such changes were shown to result in lower chance of older adults to present with low blood pressure (hypotension) or increased heart rate (tachycardia), the principal signs of shock [12].

Older adults are at greater risk of impaired respiratory responses with the process of ageing. They have significantly decreased response to hypoxia and hypercarbia [20], decreased pulmonary reserve [13], and decreased ability to effectively clear secretions and fluids from the lungs [13]. Therefore, older adults with chest trauma are at great risk of adverse events including pneumonia, respiratory failure, and death [24].

In regard to the nervous system, brain atrophy was shown to increase with age and accelerate after the age of 60 years [15]; allowing more space between brain and skull which could accumulate a high volume of blood without presenting any sign of increased intracranial pressure [36]. This, as a result, could delay the diagnosis of intracranial bleeding among older people [36]. The structure and function of the blood vessels are also impaired with aging including arterial stiffness and endothelial cells dysfunction [46]. These cerebrovascular changes could increase the vulnerability of intracranial hemorrhage for this population post injury.

Ageing is associated with progressive loss of skeletal muscle mass and strength (sarcopenia) which could negatively impact mobility and balance [16]; increasing the risk of severe injury from falls [40]. Osteoporosis increases the risk of fractures [17], from even low energy mechanisms (i.e. low-level falls) in older people [11].

Comorbidities

Comorbidities can significantly change the physiological responses to trauma for older people. High blood pressure (hypertension) is a common comorbidity for older people and is a significant risk factor of cardiovascular and cerebrovascular diseases [23]. Its presence could impair the response to shock post injury for this population. Older adults are at great risk of developing Chronic Obstructive Pulmonary Disease [8] which could adversely affect the response to hypoxia and hypercarbia post injury. Chronic Kidney Disease is also another common comorbidity among older people and could lead to decreased renal function and increased risk of Adverse Drug Reactions (ADRs) [25].

Medication use

The use of medications is more common in older people than younger populations, particularly, for those with comorbidities. Older people are more common to use anticoagulants which could significantly increase their risk of intracranial hemorrhage following head trauma [2]. As hypertension is also common in this population, they are also more likely to take antihypertensive drugs to control their blood pressure.

Antihypertensive drugs were shown to increase the risk of serious fall injuries in older people [41]. Such important medications and others that are commonly used in this population could impact their response to injury and lead to inappropriate assessment of injury severity.

Apart from the medications themselves, the term 'polypharmacy', which is defined as using multiple medications and/or administering additional medications that are not clinically indicated which represents unnecessary medications use, is common among older people [14]. This issue continues to increase in this population and is a well-

known risk factor of their death and disability [14]. Polypharmacy is significantly associated with the incidence of ADRs in older people; adding a greater risk of poor outcomes [1]. Other issues in this population with medications use include disparity between the labeled dosage and the actual used dosage, drug interactions, and underuse of medications [31]. All of these issues should be considered during the assessment and management of injured older patients.

The impact on the accuracy of prehospital triage

These which factors? were shown to have an impact on the accuracy prehospital trauma triage including physiological responses and mechanism of injury. The findings of physiological responses of older people in prehospital trauma triage are summarised in Table 1. A recent study found that even when injured older patients met the physiological criteria, they had low chance of being transported to a Trauma Centre (TC); 24% of hypotensive patients (SBP<90mmHg), 23% of those with abnormal RR (<10 or >29) , and 26% of those with a GCS <13 were transported to a TC [26].

Table 1 Physiological responses of older people post injury.

Physiological Variables	Findings from Prehospital Trauma Triage Studies
Systolic Blood Pressure (SBP)	<ul style="list-style-type: none"> • Patients aged >55 years were less likely to present with shock compared to younger adults (SBP <90 mm Hg) [SBP, mm Hg, mean (±SD):144 (33) vs. 131 (29)] [9]. • Patients aged >55 years with major trauma were less likely to have SBP <90 mm Hg than younger adults [n (%): 234/3054 (7.7) vs. 565/4407 (12.8), p =0.001] [6]. • There are decreasing rates of trauma patients presenting with hypotension (SBP <90 mm Hg) with age [22].

Heart Rate	<ul style="list-style-type: none"> • Patients aged >55 likely to have tachycardia than younger adults [Heart Rate (HR), beat per minute, mean (\pmSD): 82.7 (20) vs. 91.7 (25)] [9]. • Patients aged >55 years with major trauma were less likely to have HR >124 than younger adults [n (%): 163/3054 (5.3) vs. 609/4407 (13.8), p =0.001] [6]. • There are decreasing rates of trauma patients presenting with tachycardia (HR >100) with age [22].
Respiratory Rate (RR)	<ul style="list-style-type: none"> • Patients aged >55 years with major trauma were less likely to have abnormal RR (RR <12 or >24) than younger adults [n (%): 395/3054 (12.9) vs. 904/4407 (20.5), p =0.001] [6].
Glasgow Coma Scale (GCS)	<ul style="list-style-type: none"> • Older trauma patients aged >55 years, compared to their younger counterparts, had higher GCS [mean (\pmSD): 14.2 (2.4) vs. 13.6 (3.5)] [9]. • Patients aged >55 years with major trauma were less likely to have GCS <13 (RR <12 or >24) than younger adults [n (%): 574/3054 (18.8) vs. 1125/4407 (25.5), p =0.001] [6]. • These findings are consistent with the findings of other studies which showed higher GCS among patients aged \geq65 years with trauma brain injury compared to younger adults [18, 34]

These factors were also found to affect the mechanism of injury (i.e. increase the risk of low-level falls) which is now the most common mechanism of injury in the UK (falls <2 meters) [42]. The assessment of triage patterns for injured adults showed increasing rates of patients who had falls with age (from 12% of patients aged 16-25 years to 77% of patients aged >65 years, p <0.05) and decreasing rates of motor vehicle accidents (from 52% of patients aged 16-25 years to 16% of patients aged >65 years, p <0.05) [22]. Indeed, most injuries among patients aged \geq 65 years occurred at home usually due to falls from standing height (62%) [3]. An earlier study showed that trauma triage tool failed to identify major trauma resulting from falls (94% under-triage) [33]. Falls were related to 70% of the hospitalisations of these patients and 45% of those with

major trauma had falls [33]. Most injured patients aged ≥ 55 years due to falls were transported to non-TC (63.4% vs. 33.9%, $p < 0.001$) [39].

Adjusting prehospital trauma triage

Recent literature has investigated modifying and developing specific trauma triage criteria for older people. The application of the criterion (SBP < 110 mm Hg) instead of (SBP < 90 mm Hg) in the prehospital trauma triage tool was assessed for older adults [4]. It was shown to reduce the rate of under-triage by 4% and increase the rate of over-triage by 4% [4]. As the risk of death for older trauma patients who had a SBP < 110 mm Hg is similar to those with a SBP < 90 mm Hg, this may highlight the importance of applying this criterion for direct transport of these patients to TCs [4]. Is this clinically important?

A recent study developed specific criteria for prehospital trauma triage of older adults aged ≥ 55 years [28]. It showed that HR was not associated with major trauma (Injury Severity Score [ISS] > 15) in adjusted models ($p = 0.48$) and was excluded from further analysis [28]. In the revised triage tool, the study showed that GCS ≤ 14 was the most predictive variable of ISS > 15 [28]. Replacing current criterion (GCS ≤ 13) with GCS ≤ 14 in the trauma triage tool could decrease the rate of under-triage (sensitivity from 78.6% to 84.1%) with similar increase in the rate of over-triage (specificity from 75.5% to 68.4%) [28]. Adding RR < 10 or > 24 (including the need for assisted ventilation) was the second most predictive physiological measure as it improved sensitivity from 78.6% to 84.5% but had more lower specificity from 75.5% to 66.9% [28]. However, the change of under-triage for respiratory status compared to GCS was small (sensitivity, 84.5% and 84.1%, respectively); indicating that many patients with abnormal RR were already

identified with abnormal GCS [28]. Shock index >1.0 and SBP <110 or >200 mm Hg improved sensitivity by similar change (from 78.6% to 86.4% and 86.3%, respectively) with also similar much lower decrease in the specificity (from 75.5% to 60.4% and 60.7%, respectively); resulting in lower predictive value than GCS and RR [28].

Adding the count of comorbidities was assessed in a developed triage tool for injured older adults aged ≥ 65 years [27]. Recursive partitioning identified predictors in order; any current criterion in the triage tool, GCS ≤ 14 , abnormal geriatric specific physiological criteria (RR <10 or >24 , SBP <110 or >200 mmHg, and HR ≤ 60 or ≥ 110 beat per minute), and comorbidity count ≥ 2 [27]. This triage tool had 90.3% sensitivity (95% Confidence Interval [CI], 86.8%–93.7%) and 17.0% specificity (95% CI, 15.8%–18.1%) to identify patients with ISS >15 or require major non-orthopaedic surgery compared to current triage tool which had sensitivity of 36.6% (95% CI, 31.2–42.0%) and specificity, 90.1% (95% CI, 89.2%–91.0%) [27]. It was also found to have sensitivity and specificity to predict short and long-term mortality similar to other definitions of high-risk patients compared to current triage tool which was poor [27]. Anticoagulant use was assessed in this study and found not to be a primary predictor in recursive partitioning analyses. Adding this criterion to the developed triage tool showed 94.1% sensitivity (95% CI, 90.1%–98.1%) and 14.0% specificity (95% CI, 12.6%–15.4%). When comorbidity count was replaced by medication use, the triage tool showed 78.9% sensitivity (95% 71.5%–86.2%) and 40.8% specificity (95% CI, 38.7%–42.9). This showed that geriatric-specific physiology measure and comorbidity count are primary predictors of major trauma for this population.

Anticoagulant or antiplatelet use was assessed to be added to current prehospital trauma triage tool when assessing patients aged ≥ 55 years with head injury [29]. The study showed that applying physiological, anatomical, and mechanism of injury criteria in the triage tool had poor sensitivity in identifying traumatic intracranial haemorrhage (26/131; 19.8%, 95% CI 5.5-51.2%) and in-hospital death or neurosurgery (14/41; 34.1%, 95% CI 21.6-49.5%) [29]. Adding the criterion of anticoagulant or antiplatelet use in the triage tool had improved its sensitivity to identify traumatic Intracranial Haemorrhage (tICH) (78/131; 59.5%, 95% CI 51.0-67.6%) and death or neurosurgery (29/41; 70.7%, 95% CI 55.5-82.4%) with modest decrease in specificity from (1843/1979; 93.1%, 95% CI 91.2-94.7%) to (1329/1979; 67.2%, 95% CI 61.1-72.7%) [29].

Recommendations for Practice and Future Research

Older adults were shown to have unique factors that could impact their response to injury and reduce their chance to meet current triage criteria. Therefore, paramedics should focus on improving their knowledge about the changes in anatomy and physiology with age, the impact of comorbidities and medications use on response to injury and assessing injury severity for this population. This will aid paramedics to improve their judgments and triage decisions. A recent systematic review showed that paramedic judgment could decrease the rate of under-triage for trauma patients especially for those who did not meet the triage criteria [44]. The assessment of 'frailty' could also help in determining the severity of injury for these patients. It was shown to be an independent predictor of adverse outcomes for older trauma patients [32, 35].

Applying simple frailty assessment tools like the Clinical Frailty Scale (CFS), which was

determined to be feasible, reliable, and accurate to apply in emergency care [30], could be used in prehospital phase. The CFS was shown recently to independently predict 30-day mortality, inpatient delirium, and increased care level at discharge in older people aged ≥ 65 years post injury [35].

Apart from this, further research is needed to improve prehospital triage decisions for injured older patients to assess and look into: 1) the accuracy of different triage tools as most studies were published in the United States of America and Australia, 2) the development of application of more accurate geriatric-specific trauma triage criteria as most developed criteria were more highly sensitive but lack acceptable specificity, and 3) the destination compliance for positively triaged patients as older patients who met the triage criteria seemed less likely to be transported to a TC.

Conclusion

Older adults have unique factors related to the anatomical and physiological changes with age, comorbidities, and medications use which were shown to affect their physiological responses to injury and mechanism of injury. Current triage criteria had poor sensitivity to capture older people with major trauma. Geriatric-specific physiology measures and comorbidity count significantly improved sensitivity of prehospital triage with much lower specificity. Adding the criterion of anticoagulant or antiplatelet use for those with head injury could significantly improve the sensitivity for predicting tICH and death or neurosurgery with modest decrease in specificity. Paramedics' knowledge about the challenges when assessing and managing these patients could improve their judgments. Assessing frailty using, for example, the CFS could aid in assessing injury

severity. More research is needed to improve early and appropriate identification and triage decisions for this population.

Strengths and weaknesses of this paper

Implications for research

Implications for clinical practice

Compliance with Ethics Guidelines

Conflict of interest: The authors declare that they have no competing interests.

Patients' rights and animal protection statements: Not applicable.

Author Contributions: AA developed the structure of the article and drafted the manuscript. SC reviewed and edited the structure of the study, discussed the available literature around the main heading with AA, monitored the progress of the article, and critically reviewed and edited the manuscript. All authors read and approved the final manuscript.

References

1. Ahmed B, Nanji K, Mujeeb R et al. (2014) Effects of polypharmacy on adverse drug reactions among geriatric outpatients at a tertiary care hospital in Karachi: a prospective cohort study. *PLoS One* 9:e112133
2. Batchelor JS, Grayson A (2012) A meta-analysis to determine the effect of anticoagulation on mortality in patients with blunt head trauma. *British journal of neurosurgery* 26:525-530
3. Brown E, Tohira H, Bailey P et al. (2019) Older age is associated with a reduced likelihood of ambulance transport to a trauma centre after major trauma in Perth. *Emergency Medicine Australasia* 31:763-771
4. Brown JB, Gestring ML, Forsythe RM et al. (2015) Systolic blood pressure criteria in the National Trauma Triage Protocol for geriatric trauma: 110 is the new 90. *The journal of trauma and acute care surgery*. 78:352-359
5. Collicott PE, Hughes I (1980) Training in advanced trauma life support. *Jama* 243:1156-1159
6. Cox S, Morrison C, Cameron P et al. (2014) Advancing age and trauma: triage destination compliance and mortality in Victoria, Australia. *Injury* 45:1312-1319
7. Dinh MM, Roncal S, Byrne CM et al. (2013) Growing trend in older patients with severe injuries: mortality and mechanisms of injury between 1991 and 2010 at an inner city major trauma centre. *ANZ journal of surgery* 83:65-69
8. Divo MJ, Martinez CH, Mannino DM (2014) Ageing and the epidemiology of multimorbidity. In:*Eur Respiratory Soc*
9. Garwe T, Stewart K, Stoner J et al. (2017) Out-of-hospital and Inter-hospital Under-triage to Designated Tertiary Trauma Centers among Injured Older Adults: A 10-year Statewide Geospatial-Adjusted Analysis. *Prehospital Emergency Care* 21:734-743
10. Gentile-Lorente D, Salvado-Usach T (2016) Sigmoid septum: A variant of the ventricular hypertrophy or of the hypertrophic cardiomyopathy? *Archivos de cardiologia de Mexico* 86:110-122
11. Greenspan SL, Myers ER, Maitland LA et al. (1994) Fall severity and bone mineral density as risk factors for hip fracture in ambulatory elderly. *Jama* 271:128-133
12. Gribbin B, Pickering TG, Sleight P et al. (1971) Effect of age and high blood pressure on baroreflex sensitivity in man. *Circulation research* 29:424-431
13. Haas CF, Loik PS, Gay SE (2007) Airway clearance applications in the elderly and in patients with neurologic or neuromuscular compromise. *Respiratory care* 52:1362-1381
14. Hajjar ER, Cafiero AC, Hanlon JT (2007) Polypharmacy in elderly patients. *The American journal of geriatric pharmacotherapy* 5:345-351
15. Hedman AM, Van Haren NE, Schnack HG et al. (2012) Human brain changes across the life span: a review of 56 longitudinal magnetic resonance imaging studies. *Human brain mapping* 33:1987-2002
16. Janssen I, Heymsfield SB, Ross R (2002) Low relative skeletal muscle mass (sarcopenia) in older persons is associated with functional impairment and physical disability. *Journal of the American Geriatrics Society* 50:889-896

17. Kannus P, Palvanen M, Niemi S et al. (1996) Increasing number and incidence of osteoporotic fractures of the proximal humerus in elderly people. *Bmj* 313:1051-1052
18. Kehoe A, Smith J, Bouamra O et al. (2016) Older patients with traumatic brain injury present with a higher GCS score than younger patients for a given severity of injury. *Emergency medicine journal* 33:381-385
19. Kehoe A, Smith J, Edwards A et al. (2015) The changing face of major trauma in the UK. *Emerg Med J* 32:911-915
20. Kronenberg RS, Drage CW (1973) Attenuation of the ventilatory and heart rate responses to hypoxia and hypercapnia with aging in normal men. *The Journal of clinical investigation* 52:1812-1819
21. Lakatta E, Gerstenblith G, Angell CS et al. (1975) Prolonged contraction duration in aged myocardium. *The Journal of clinical investigation* 55:61-68
22. Lehmann R, Beekley A, Casey L et al. (2009) The impact of advanced age on trauma triage decisions and outcomes: a statewide analysis. *American Journal of Surgery* 197:571-575
23. Lionakis N, Mendrinou D, Sanidas E et al. (2012) Hypertension in the elderly. *World journal of cardiology* 4:135
24. Lottipour S, Kaku SK, Vaca FE et al. (2009) Factors associated with complications in older adults with isolated blunt chest trauma. *Western journal of emergency medicine* 10:79
25. Mallappallil M, Friedman EA, Delano BG et al. (2014) Chronic kidney disease in the elderly: evaluation and management. *Clinical practice (London, England)* 11:525
26. Meyers MH, Wei TL, Cyr JM et al. (2019) The Triage of Older Adults with Physiological Markers of Serious Injury Using a State-Wide Prehospital Plan. *Prehospital and Disaster Medicine* 34:497-505
27. Newgard CD, Lin A, Eckstrom E et al. (2019) Comorbidities, anticoagulants, and geriatric-specific physiology for the field triage of injured older adults. *The journal of trauma and acute care surgery* 86:829
28. Newgard CD, Richardson D, Holmes JF et al. (2014) Physiological field triage criteria for identifying seriously injured older adults. *Prehospital Emergency Care* 18:461-470
29. Nishijima DK, Gaona SD, Waechter T et al. (2017) Out-of-hospital triage of older adults with head injury: a retrospective study of the effect of adding "anticoagulation or antiplatelet medication use" as a criterion. *Annals of emergency medicine* 70:127-138. e126
30. O'caoimh R, Costello M, Small C et al. (2019) Comparison of frailty screening instruments in the emergency department. *International journal of environmental research and public health* 16:3626
31. Ostrom JR, Hammarlund ER, Christensen DB et al. (1985) Medication usage in an elderly population. *Medical Care*:157-164
32. Pecheva M, Phillips M, Hull P et al. (2020) The impact of frailty in major trauma in older patients. *Injury*

33. Phillips S, Rond lli PC, Kelly SM et al. (1996) The failure of triage criteria to identify geriatric patients with trauma: Results from the Florida Trauma Triage Study. *Journal of Trauma* 40:278-283
34. Rau C-S, Wu S-C, Chen Y-C et al. (2017) Effect of age on Glasgow Coma Scale in patients with moderate and severe traumatic brain injury: an approach with propensity score-matched population. *International journal of environmental research and public health* 14:1378
35. Rickard F, Ibitoye S, Deakin H et al. (2020) The Clinical Frailty Scale predicts adverse outcome in older people admitted to a UK major trauma centre. *Age and ageing*
36. Rose J, Valtonen S, Jennett B (1977) Avoidable factors contributing to death after head injury. *Br Med J* 2:615-618
37. Salmasi A-M, Alimo A, Jepson E et al. (2003) Age-associated changes in left ventricular diastolic function are related to increasing left ventricular mass. *American journal of hypertension* 16:473-477
38. Sasser SM, Hunt RC, Faul M et al. (2012) Guidelines for field triage of injured patients: recommendations of the National Expert Panel on Field Triage, 2011. *MMWR: Recommendations and Reports* 61:1-20
39. Staudenmayer KL, Hsia RY, Mann NC et al. (2013) Triage of elderly trauma patients: A population-based perspective. *Journal of the American College of Surgeons* 217:569-576
40. Sterling DA, O'connor JA, Bonadies J (2001) Geriatric falls: injury severity is high and disproportionate to mechanism. *Journal of Trauma and Acute Care Surgery* 50:116-119
41. Tinetti ME, Han L, Lee DS et al. (2014) Antihypertensive medications and serious fall injuries in a nationally representative sample of older adults. *JAMA internal medicine* 174:588-595
42. Trauma Audit and Research Network (Tarn) (2017) Major Trauma In Older People. In, Salford/United Kingdom
43. Van Der Heijden-Spek JJ, Staessen JA, Fagard RH et al. (2000) Effect of age on brachial artery wall properties differs from the aorta and is gender dependent: a population study. *Hypertension* 35:637-642
44. Van Rein EA, Van Der Sluijs R, Raaijmakers AM et al. (2018) Compliance to prehospital trauma triage protocols worldwide: A systematic review. *Injury* 49:1373-1380
45. Virmani R, Avolio A, Mergner W et al. (1991) Effect of aging on aortic morphology in populations with high and low prevalence of hypertension and atherosclerosis. Comparison between occidental and Chinese communities. *The American journal of pathology* 139:1119
46. Xu X, Wang B, Ren C et al. (2017) Age-related impairment of vascular structure and functions. *Aging and disease* 8:590