Traumatic coagulopathy and massive transfusion: improving outcomes and saving blood

Karim Brohi* and Simon Eaglestone

Centre for Trauma Sciences, Blizard Institute, Barts and the London School of Medicine, Queen Mary University of London, London, UK

*Corresponding author k.brohi@qmul.ac.uk

Declared competing interests of authors: Karim Brohi reports that TEM Innovations GmbH (ROTEM®) provided unrestricted support to the programme in the form of equipment and reagents for the observational study [National Institute for Health Research (NIHR) portfolio ID 5637].

Published November 2017 DOI: 10.3310/pgfar05190

Scientific summary

Traumatic coagulopathy and transfusion

Programme Grants for Applied Research 2017; Vol. 5: No. 19 DOI: 10.3310/pgfar05190

NIHR Journals Library www.journalslibrary.nihr.ac.uk

Scientific summary

Introduction

Trauma-induced coagulopathy (TIC) often complicates severe haemorrhage and is associated with significantly worse outcomes for trauma patients. TIC was previously thought to occur late and primarily to be caused by the consumption and dilution of clotting factors. However, the recognition that TIC developed rapidly and was more complex in its aetiology also suggested new management strategies and therapeutic options. There was sparse evidence on the epidemiology of trauma haemorrhage, existing practice patterns, outcomes and costs, as well as little information on the patterns of coagulopathy present on arrival and during haemorrhage and how these responded to transfusion therapy.

Methods

Between 2008 and 2013 we conducted a multimodal programme of work to develop our understanding of coagulopathy and its optimal management. We identified existing evidence, practice patterns and outcomes through systematic reviews of the literature and a national study of trauma haemorrhage, its transfusion management and associated health-care costs. We further examined several point-of-care coagulation tools for their ability to diagnose TIC and to assess the response to blood component therapy. We progressively implemented our findings into practice and assessed the outcomes of trauma patients presenting to our major trauma centre. To examine different approaches to the provision of blood to casualties in a mass casualty event, we constructed a discrete event model based on data from the 2005 London bombings.

Key results

Our national study recruited 442 patients in 22 hospitals and found that the 1-year mortality rate for patients with major haemorrhage [requiring 4+ units of packed red blood cells (PRBCs) in the first 24 hours] was and approached 50% for those with massive haemorrhage (10+ PRBCs). Half of these deaths occurred in the first 24 hours after injury and nearly one-quarter in the first 4 hours. In this critical window the delivery of blood component therapy was often below the recommended thresholds. Studying the pattern of TIC at this time point we determined that loss of fibrinogen and excessive fibrinolysis were key derangements. We were able to determine that rotational thromboelastometry could rapidly identify patients with TIC and high transfusion requirements, based on the clot amplitude at 5 minutes. We were further able to show how existing damage control resuscitation regimens do not maintain haemostatic competence during bleeding. Furthermore, fibrinogen levels started below the recommended range and became dangerously low after 8 units of transfusion without supportive transfusion therapy. Severe fibrinolysis was extremely common (seen in > 60% of patients) and in its most extreme phenotype only detectable by thromboelastometry. In total, the estimated cost of treating a major haemorrhage patient was £20,600 and the estimated cost of treating a massive haemorrhage patient was £24,000. Nationally, the estimated cost of trauma haemorrhage is £85M annually. In mass casualty situations early results show that the only mutable factor that has a large effect on the provision of care in the initial phase of the response is the level of blood stocks held in the receiving hospital.

Conclusions

Outcomes from trauma haemorrhage remain poor at a national level and there are important areas for improvement with regard to the delivery of transfusion therapy within the critical early post-injury phase of care. Implementation of consistent high-dose plasma-based damage control resuscitation improves outcomes but does not correct TIC during haemorrhage. Key features of TIC that may respond to therapy are loss of fibrinogen and excessive fibrinolysis. Thromboelastometry may be able to personalise coagulation therapy using new diagnostic criteria to correct TIC and further improve outcomes for critically injured bleeding trauma patients.

Funding

Funding for this study was provided by the Programme Grants for Applied Research programme of the National Institute for Health Research.

Programme Grants for Applied Research

ISSN 2050-4322 (Print)

ISSN 2050-4330 (Online)

This journal is a member of and subscribes to the principles of the Committee on Publication Ethics (COPE) (www.publicationethics.org/).

Editorial contact: journals.library@nihr.ac.uk

The full PGfAR archive is freely available to view online at www.journalslibrary.nihr.ac.uk/pgfar. Print-on-demand copies can be purchased from the report pages of the NIHR Journals Library website: www.journalslibrary.nihr.ac.uk

Criteria for inclusion in the Programme Grants for Applied Research journal

Reports are published in *Programme Grants for Applied Research* (PGfAR) if (1) they have resulted from work for the PGfAR programme, and (2) they are of a sufficiently high scientific quality as assessed by the reviewers and editors.

Programme Grants for Applied Research programme

The Programme Grants for Applied Research (PGfAR) programme, part of the National Institute for Health Research (NIHR), was set up in 2006 to produce independent research findings that will have practical application for the benefit of patients and the NHS in the relatively near future. The Programme is managed by the NIHR Central Commissioning Facility (CCF) with strategic input from the Programme Director.

The programme is a national response mode funding scheme that aims to provide evidence to improve health outcomes in England through promotion of health, prevention of ill health, and optimal disease management (including safety and quality), with particular emphasis on conditions causing significant disease burden.

For more information about the PGfAR programme please visit the website: http://www.nihr.ac.uk/funding/programme-grants-for-applied-research.htm

This report

The research reported in this issue of the journal was funded by PGfAR as project number RP-PG-0407-10036. The contractual start date was in July 2008. The final report began editorial review in August 2014 and was accepted for publication in June 2017. As the funder, the PGfAR programme agreed the research questions and study designs in advance with the investigators. The authors have been wholly responsible for all data collection, analysis and interpretation, and for writing up their work. The PGfAR editors and production house have tried to ensure the accuracy of the authors' report and would like to thank the reviewers for their constructive comments on the final report document. However, they do not accept liability for damages or losses arising from material published in this report.

This report presents independent research funded by the National Institute for Health Research (NIHR). The views and opinions expressed by authors in this publication are those of the authors and do not necessarily reflect those of the NHS, the NIHR, CCF, NETSCC, PGfAR or the Department of Health. If there are verbatim quotations included in this publication the views and opinions expressed by the interviewees are those of the interviewees and do not necessarily reflect those of the authors, those of the NHS, the NIHR, NETSCC, the PGfAR programme or the Department of Health.

© Queen's Printer and Controller of HMSO 2017. This work was produced by Brohi et al. under the terms of a commissioning contract issued by the Secretary of State for Health. This issue may be freely reproduced for the purposes of private research and study and extracts (or indeed, the full report) may be included in professional journals provided that suitable acknowledgement is made and the reproduction is not associated with any form of advertising. Applications for commercial reproduction should be addressed to: NIHR Journals Library, National Institute for Health Research, Evaluation, Trials and Studies Coordinating Centre, Alpha House, University of Southampton Science Park, Southampton SO16 7NS, UK.

Published by the NIHR Journals Library (www.journalslibrary.nihr.ac.uk), produced by Prepress Projects Ltd, Perth, Scotland (www.prepress-projects.co.uk).

Programme Grants for Applied Research Editor-in-Chief

Professor Paul Little Professor of Primary Care Research, University of Southampton, UK

NIHR Journals Library Editor-in-Chief

Professor Tom Walley Director, NIHR Evaluation, Trials and Studies and Director of the EME Programme, UK

NIHR Journals Library Editors

Professor Ken Stein Chair of HTA and EME Editorial Board and Professor of Public Health, University of Exeter Medical School, UK

Professor Andrée Le May Chair of NIHR Journals Library Editorial Group (HS&DR, PGfAR, PHR journals)

Dr Martin Ashton-Key Consultant in Public Health Medicine/Consultant Advisor, NETSCC, UK

Professor Matthias Beck Chair in Public Sector Management and Subject Leader (Management Group), Queen's University Management School, Queen's University Belfast, UK

Dr Tessa Crilly Director, Crystal Blue Consulting Ltd, UK

Dr Eugenia Cronin Senior Scientific Advisor, Wessex Institute, UK

Dr Peter Davidson Director of the NIHR Dissemination Centre, University of Southampton, UK

Ms Tara Lamont Scientific Advisor, NETSCC, UK

Dr Catriona McDaid Senior Research Fellow, York Trials Unit, Department of Health Sciences, University of York, UK

Professor William McGuire Professor of Child Health, Hull York Medical School, University of York, UK

Professor Geoffrey Meads Professor of Wellbeing Research, University of Winchester, UK

Professor John Norrie Chair in Medical Statistics, University of Edinburgh, UK

Professor John Powell Consultant Clinical Adviser, National Institute for Health and Care Excellence (NICE), UK

Professor James Raftery Professor of Health Technology Assessment, Wessex Institute, Faculty of Medicine, University of Southampton, UK

Dr Rob Riemsma Reviews Manager, Kleijnen Systematic Reviews Ltd, UK

Professor Helen Roberts Professor of Child Health Research, UCL Institute of Child Health, UK

Professor Jonathan Ross Professor of Sexual Health and HIV, University Hospital Birmingham, UK

Professor Helen Snooks Professor of Health Services Research, Institute of Life Science, College of Medicine, Swansea University, UK

Professor Jim Thornton Professor of Obstetrics and Gynaecology, Faculty of Medicine and Health Sciences, University of Nottingham, UK

Professor Martin Underwood Director, Warwick Clinical Trials Unit, Warwick Medical School, University of Warwick, UK

Please visit the website for a list of members of the NIHR Journals Library Board: www.journalslibrary.nihr.ac.uk/about/editors

Editorial contact: journals.library@nihr.ac.uk