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1. Introduction

Hemorrhagic shock (HS) is the leading cause of traumarelated deaths in civilian patients surviving to the trauma center, accounting for about 40% of trauma deaths globally [1]. On the battle field, it is the leading cause of death in potentially survivable cases, where it accounts for as much as 80% of these cases [2]. Fluid resuscitation, hemorrhage control, correction of coagulopathy, and rapid transport to the trauma center are key tenets of care for the traumatic HS patient which have been associated with improved outcome [3].

144 The effect of host factors such as age, gender, and in-145 flammatory response on trauma outcomes has been studied 146 extensively. In spite of this, the exact relationship between 147 age and trauma outcomes is far from settled. Several in-148 149 vestigators have documented increasing mortality with age 150 after blunt or penetrating injury. Hollis et al. [4] studied 65,743 151 patients from the database of the United Kingdom Trauma 152 Network and showed that there was an increase in mortality 153 with increasing age at all levels of injury severity. Morris et al. 154 [5] examined 199,737 trauma patients in the state of Califor-155 nia and showed that mortality in the entire cohort increased 156 starting at the age of 40 y, with the mortality rates becoming 157 higher in the elderly at the age of \geq 65 y in patients with 158 minor injuries. They also showed that in patients with 159 160 moderate injury severity, mortality increased in both the 161 middle age (40–64 y) and elderly groups (\geq 65 y). They found 162 that the presence of preexisting medical disease or injury to 163 the head or abdomen was related to increased mortality in 164 middle-aged patients at all injury severity levels. In contrast, 165 Bhullar et al. [6] studied 539 hemodynamically stable patients 166 with blunt splenic trauma who underwent nonoperative 167 management and concluded that age had no effect on out-168 comes, the outcome in this case being failure of nonoperative 169 management. 170

171 Most studies examining the relationship between 172 trauma outcomes and age seem to reach one conclusion as 173 age increases and mortality increases. The specific factors 174 that predict mortality in different age groups are seldom 175 examined. In traumatic HS in particular, there are very 176 sparse data examining the association between age and 177 mortality. This is important because there is some evidence 178 from preclinical studies showing that animals of different 179 ages have different optimal resuscitation blood pressures 180 181 after uncontrolled hemorrhage and these resuscitation 182 pressures influence both survival time and 24-h survival 183 rate [7].

184 Therefore, the need to identify the specific determinants 185 of survival or mortality in different age groups after HS 186 remains valid. We therefore hypothesized that (1) there is a 187 distinct mortality inflection point, that is, the age at which 188 there is a sharp rise in mortality after blunt traumatic HS, 189 and (2) the determinants of survival after blunt traumatic 190 HS vary from one age group to the other. We sought to 191 examine the relationship between age and mortality as well 192 193 as identify the predictors of mortality in multiple age group 194 cohorts. 195

2. Materials and methods

2.1. Study design and patient selection

This is a retrospective analysis of the Inflammation and the Host Response to Injury trauma relational database (Glue Grant database), a large scale collaborative program that involved multiple level I trauma centers in the United States and established to study the inflammatory changes accompanying trauma and burns. The criteria for patient inclusion in the original database development have been published previously [8]. Briefly, these were patients aged >16 y who sustained blunt trauma; were in HS as evidenced by a base deficit \geq 6 or systolic blood pressure (SBP) <90 mm Hg within 60 min of emergency room (ER) arrival; were transfused with blood within 12 h of injury; and had no evidence of severe traumatic brain or spinal cord injury, that is, Glasgow coma scale (GCS) <8 after intensive care unit (ICU) admission and brain computerized tomography scan abnormality within first 12 h after injury. Patients with anticipated survival of <24 h from injury or anticipate survival <28 d because of a preexisting medication were excluded. In our present study, all patients with complete clinical and laboratory data points were included.

2.2. Patient stratification

Patients were initially stratified into eight age groups (16–24 y, 25–34 y, 35–44 y, 45–54 y, 55–64 y, 65–74 y, 75–84 y, and \geq 85 y) to identify the mortality inflection point, that is, the age group at which there is a sharp or sudden change in mortality on a curve showing the relationship between mortality and age group. This initial classification into eight age groups (referred to in this study as age group I) was used solely for determination of the mortality inflection point. For subsequent analyses, patients were restratified into young age (16–44 y), middle age (45–64 y), and elderly (\geq 65 y), using definitions from the US Census Bureau [9]. For clarification purposes, this second stratification into three age groups is referred to as age group II.

2.3. Data variables and outcome measures

The variables analyzed include demographics (age, gender, and ethnicity), injury severity score (ISS), preinjury comorbidities, inhospital complications, Marshall's multipleorgan dysfunction score (MODS), ER serum lactate level, volume of resuscitative fluids (crystalloids, colloids, and blood products), and operative interventions. The primary outcome measure was inhospital mortality. Hospital length of stay and ICU length of stay were the secondary outcome measures.

2.4. Statistical analysis

Continuous variables were described with summary statistics, whereas proportions were used to describe categorical

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