
Supplementary information

Trauma-induced coagulopathy

In the format provided by the
authors and unedited

Supplementary information

Table S1: Civilian studies addressing haemorrhagic deaths

Studies	Year(s) covered by study	Setting	% blunt trauma	% of pre-hospital deaths due to haemorrhage	% of in-hospital deaths due to haemorrhage
Sauaia et al. ¹	1992	Population-based Denver, CO	51%	36%	41%
Shackford et al. ²	1992	Population-based S. Diego, CA	71%	31%	
Stewart et al. ³	1995-2001	Hospital-based S. Antonio, TX	71%	NA	21%*
Tien et al. ⁴	1999-2003	Hospital-based Toronto, Canada	87%	NA	15%
Evans et al. ⁵	2005	Population-based Newcastle, Australia	86%	33%	
Consunji et al. ⁶	2004-2007	Hospital-based Manila, Philippines	41%	NA	28%
Soreide et al. ⁷	2010	Population-based Stavanger, Norway	87%	12.5%***	12.5%***
Kahl et al. ⁸	2000-2011	Hospital-based S. Diego, CA	69%	NA	28%
Kleber et al. ⁹	2010	Population-based Berlin, Germany	87%	14%#	4%#
Trajano et al. ¹⁰	1995, 2000, 2005, 2010	Hospital-based Campinas, Brazil	73%	NA	18%
Davis et al. ¹¹	2011	Population-based Miami-Dade County, FL	53%	34%	NA
Roberts et al. ¹²	2005-2013	Hospital-based Calgary, Canada	94%	NA	27%
Arsilan et al. ¹³	2010-2013	Hospital-based Ankara, Turkey	85%	28%‡	29%
Oyenyi et al. ¹⁴	2005-2006 and 2012-2013	Hospital-based Houston, TX	81-80%	NA	2005-2006: 36% 2007-2013: 25%
Drake et al. ¹⁵ **	2014	Population-based Harris County, TX	62%	20%	20%
Jochems et al. ¹⁶	2007-2016	Hospital-based Utrecht, Netherlands	98%	NA	2007-2012: 9% 2013-2016: 3%
Callcut et al. ¹⁷	2015-2017	Hospital-based 18 centers across US	73%	NA	23%

*Attributed to shock; **18% of the trauma-related deaths occurred after the index hospitalization, of which 5(1%) were attributed to hemorrhage; ***authors estimation based on the time from injury to death reported in original study; # definition excludes 'polytrauma deaths'; ‡ death at scene or during transport but transported to the hospital

Table S2: Epidemiologic studies of trauma-induced coagulopathy (TIC) since 2000

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Raum et al. ¹⁸	2001	1993-1999	PT<60% than normal plasma	1351 adult trauma patients with ISS>15, multicenter	Germany	NA	30	NA	27%	47%	ED	NA	ISS, hypoperfusion
Keller et al. ¹⁹	2001	1999-2000	INR>1.2	56 children with blunt TBI, single center	US	100%	NA	NA	38%	35%	ED	TBI	GCS<=9
Brohi et al. ²⁰	2003	1993-1998	PT>18s, PTT>60s, TT>15s (1.5 x normal)	1088 helicopter-transported patients, single center	United Kingdom	75%	20	18%	24%	46%	ED	TIC less frequent in isolated TBI	hypoperfusion
McLeod et al. ²¹	2003	1995-2000	PT>14s or PTT>34s	10,790 injured adults, single center	US	NA	9	12%	28%	19%	ED	NA	NA
Carrick et al. ²²	2005	2001-2003	PT>14.2s or PTT > 38.4s	184 adults with blunt TBI, single center	US	100%	17		21% upon admission, 41% within 72hours	62%	ED and 72 hours	severity of TBI associated with TIC	Severity of TBI
Affonseca et al. ²³	2007	1998-2003	Prothrombin activity < 70%, PT>16s, PTT 10s>control time, platelets< 150,000/mm ³	301 children with blunt TBI	Brazil	100%	NA	27%	76%	40%	ED	TIC correlated with TBI severity, TIC not related to extra cranial injuries	Severity of TBI
Rugeri et al. ²⁴	2007	2004	INR>1.6 or PTT>60s or platelets< 100,000/mm ³ or fibrinogen <1g/L	88 adults admitted to TRU, single center	US	NA	22	NA	28%	NA	TRU	NA	NA

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Maegele et al. ²⁵	2007	2007 (period not specified)	PT (Quick's value) <70%, platelets <100,000/mm ³	8724 adults, multicenter	Germany	96%	30 in TIC; 21 in non-TIC	NA	34%	28%	ED	TIC less frequent in isolated head	ISS, pre-hospital fluids
Niles et al. ²⁶	2008	2003-2004	INR>=1.5	347 combat patients who received RBC, single combat support hospital	US-Iraq conflict	8%	17	25% required massive transfusion	38%	24%	admission	NA	NA
Kashuk et al. ²⁷	2008	2001-2006	INR>1.5	133 adults who received >10 units RBC in 6 hours, single center	US	56%	36	100%	30%	NA	first 6 hours post hospital admission	NA	ED temperature <34°C
Talving et al. ²⁸	2009	2005-2007	platelets <100,000/mm ³ , INR>1.1, PTT>36s	436 adults with TBI	US	32-33%	NA	4%	36% all TBI, 34% in isolated TBI	49%	ED	TBI	GCS<8, cerebral edema, SAH, midline shift
Frith et al. ²⁹	2010	2007-	INR>1.2 and INR>1.5	3646 adults, multicenter	United Kingdom, Germany, Netherlands, Norway, US	90%	22	24%	INR>1.2 36%, INR>1.5 19%	INR>1.2 23%, INR>1.5 28%	ED	NA	Combination of tissue injury and Shock
Wafaisade et al. ³⁰	2010	1993-2007	PT (Quick's value) <70%, platelets <100,000/mm ³	3,114 adults with blunt, isolated TBI, multicenter	Germany	100%	TIC 25 non-TIC 21	Median SBP (mmHg): TIC: 125 non-TIC 134	23%	50%	ED	Isolated TBI	age>=75yrs, pre-hospital fluids>=2000ml, shock at scene or ED, GCS<9 at scene, AIS head

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Wafaisade et al. ³¹	2010	2002-2007	PT (Quick's value) <70%, platelets <100,000/mm ³	1987 adults with ISS>15, multicenter	Germany	95%	TIC 36 non-TIC: 28	Median SBP (mmHg) TIC: 104 non-TIC: 123	36%	26%	ED	NA	ISS, AIS abdomen, shock at scene or ED, ratio of prehospital colloids: crystalloids ≥1.2. prehospital fluids ≥3000 ml
Dirks et al. ³²	2010	2001-2003 and 2005-2007	PTT>35s	253 adults who required ≥1 red blood cells unit, single center	Denmark	85%	2001-03: 20 2005-07: 24	NA	PTT 2001-03: 27% 2005-07: 34%	49%	ED	NA	NA
Jeger et al. ³³	2010	2007-2008	INR>1.5 and PTT>60s or TT>15s	172 adults with ISS>15, single center	Switzerland	NA	25	NA	33%	NA	ED	NA	NA
Tauber et al. ³⁴	2011	2005-2008	FIBTEM MCF<7mm, EXTEM MCF<45mm, EXTEM CFT>200s, EXTEM CT>100s, fibrinogen <150mg/dL or INR>1.5 or PTT>50s, platelets<100000/mm ³	Adults with polytrauma patients, single center	Austria	100%	34	16%	26% fibrinogen <150mg/L, 30% impaired fibrin polymerization, 22% reduced clot firmness INR>1.5 14%, 7% hyperfibrinolysis	Hyperfibrinolysis 57%; FIBTEM MCF<7mm 21%, EXTEM>100 s 46%, EXTEM CFT>200 s 27%, EXTEM MCF<45mm 25%	ED	Hyperfibrinolysis associated with abdominal trauma	TBI, abdominal trauma (hyperfibrinolysis), acidosis, hypothermia, hypoperfusion, ISS
Johansson et al. ³⁵	2011	2010	PTT > 35s, INR > 1.2	80 adults, single center	Denmark	91%	17	Median shock index 0.62	15%	50%	ED	NA	ISS, hypoperfusion

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Greuters et al. ³⁶	2011	2003-2007	PTT >40s, INR >1.2, platelets <120,000/mm ³	107 adults with isolated TBI	Netherlands	93%	25	NA	ED: 24%; 24 hours postinjury : 40%	NA	ED, 24 hours postinjury	Isolated TBI	NA
Patregnani et al. ³⁷	2012	2002-2009	INR ≥ 1.5	744 children in US combat support hospitals in Iraq and Afghanistan	U.S. combat support hospitals in Iraq and Afghanistan	17%	10	38%	27%	22%	ED	NA	ISS, shock
Hendrickson et al. ³⁸	2012	2006-2010	PT >15.9s, PTT >42.1s, fibrinogen <180 mg/dL, platelets <185,000/mm ³	102 children requiring blood products, single center	US	80%	20	NA	77%	NA (increased mortality odds associated with abnormal PT, PTT, and platelet count, but not fibrinogen)	ED	TBI	ISS, GCS
Mitra et al. ³⁹	2012	2005-2008	INR >1.5 or PTT >60 s	772 adults with ISS >15 who required ≥1 red blood cells unit, single center	Australia	92%	29	37% required massive transfusion	25%	38%	ED	NA	NA
Peiniger et al. ⁴⁰	2012	2002-2008	PT test (Quick value) <70%; PTT >38s, platelets <100,000/mm ³	200 children with isolated blunt TBI, multicenter	Germany	100%	NA (51% GCS <=8)	38%	GCS <=8: 44% GCS >8: 14%	GCS <=8: 52%	ED	TBI	GCS <=8

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Genét et al. ⁴¹	2013	NA	INR>1.2, PTT> 35s	80 adult trauma patients, single center	Denmark	91%	17	median shock index 0.62	TBI+other trauma: 47% isolated TBI 13% non-TBI 5%	NA	ED	NA	NA
Chhabra et al. ⁴²	2013	2008-2010	INR>1.5 and PTT>1.5 x control time	208 adults with isolated TBI, single center	India	NA	NA	NA	46%	30%	ED and 72 hours	effaced basal cisterns on CT scan	GCS ≤ 8, effaced basal cisterns on CT scan, Hemoglobin <10 g/dl and D-dimer >1 µg/dl
Xu et al. ⁴³	2013	2008-2009	PT > 18s, INR > 1.6, PTT > 60s	223 adults with blunt trauma, ISS>15, single center	China	100%	33	14% base deficit >6mEq/L	23%	37%	ICU		Base deficit ≥6 mEq/L, GCS <=8, platelet count
Alexiou et al. ⁴⁴	2014	2008-2012	PTT>40s, INR > 1.2, platelet <120,000 /mm ³	149 adults with isolated TBI	Greece	NA	NA	NA	23%	NA	24 hours postadmission	Isolated TBI	Glucose >151 mg/dl, Hemoglobin <12.4 mg/dL
Neal et al. ⁴⁵	2014	2003-2010	INR>1.5	1897 adults with blunt trauma, multicenter	US	100%	34	52%	22%	NA	ED	NA	Time from injury, age, hypoperfusion, serious comorbidity
MacLeod et al. ⁴⁶	2014	2008-2009	PT>13.3s	701 adult trauma patients, single center	US	74%	9	Pre-hospital median SBP 130 mmHg	16%	9% in PT 13.4-15.5s; 26% in PT 15.6-17.5s; 33% in PT 18-20s; 36% in PT>20s	ED	NA	TBI, ISS and hypotension

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Moore et al. ⁴⁷	2014	2010-2013	TEG LY30: hyperfibrinolysis ($\geq 3\%$)	180 adults with ISS >15 , single center	US	79%	29	Median SBP 104mmHg	Hyperfibrinolysis 18%	Hyperfibrinolysis 45%	field or ED	NA	NA
Tonglet et al. ⁴⁸	2014	2012-2013	20% abnormality in ROTEM, INR >1.3 , or fibrinogen $<150\text{mg/dL}$ at admission or 3 hours later.	82 adults with severe blunt trauma, single center	Belgium	100%	13	13%	13%	55%	ED	NA	NA
Kutcher et al. ⁴⁹	2015	2003-2011	INR >1.2	1,537 adults with blunt trauma and hemorrhagic shock, multicenter	US	100%	33	65%	38%	23%	ED		Age $<20\text{yrs}$, GCS, transport time, ISS, lower pH, lower temperature, 2-4L of pre-hospital fluids if high lactate, >4 L pre-hospital fluids regardless of lactate
de Oliveira et al. ⁵⁰	2015	2007	INR >1.5 , PTT $>60\text{s}$, platelets $<100,000/\text{mm}^3$	345 adults with ISS >15 , single center	Canada	NA	Isolated TBI: 21 TBI+other injuries: 37 No TBI: 25	NA	Isolated TBI: 13% TBI+other injuries: 31% No TBI: 23%	Isolated TBI: 66% TBI+other injuries: 48% No TBI: 19%	ED		ISS, severe trauma with no TBI, hypoperfusion

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Peltan et al. ⁵¹	2015	2003-2010	INR>1.2 and INR>1.5	1031 adults with blunt trauma requiring ≥ 1 RBC, multicenter	US	100%	33	23%	50% INR >1.2 and 21% INR >1.5	19% INR >1.2 and 27% INR >1.5	ED	Femur fractures associated with INR>1.5	Age, hypotension, ISS, APACHE II, BMI, lung contusion, femur fracture
Khan et al. ⁵²	2015	2009-2013	ROTEM CA5 ≤ 35 mm	106 trauma patients who received >3 RBC, multicenter	UK, Norway, Denmark, Germany, Netherlands	87%	34	46%	40%	NA	ED	NA	NA
Hagemo et al. ⁵³	2015	2007-2011	INR>1.2	808 adults, multicenter	UK, Denmark and Norway	82%	16	6% required massive transfusion	11%	NA	ED	NA	NA
Holcomb et al. ⁵⁴	2015	2012-2013	INR>1.5	680 adults who required ≥ 1 red blood cells unit, multicenter (RCT)	US	53%	26	39%	17%	NA	ED	NA	NA
Strumwasser et al. ⁵⁵	2016	2004-2009	INR ≥ 1.5 , PTT> 36s, platelets <100,000 /mm ³	20,126 (7.6% pediatric, 92.4% adult) trauma patients, single center	US	87% in children; 77% in adults	ISS>15: 15% children; 17% in adults	0.8% in children; 3% in adults	5.8% in children and 8.4% in adults	14% in children and 18% in adults	ED	TBI associated with TIC especially in the elderly	TBI, age, hypoperfusion, tissue injury
Peltan et al ⁵⁶	2016	2008-2013	INR > 1.5	Derivation: 1963 adults with blunt trauma, multicenter; Validation: 285 adults with blunt	US	100%	no TIC 17, TIC 26 (derivation) and 32	Derivation: 28% SBP<130mmHg; Validation: 37% SBP<117mmHg	Derivation 6%; Validation 9%	Derivation 46%	ED	NA	Injury severity and hypoperfusion

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS (validation)	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Moore et al. ⁵⁷	2016	2010-2013	Hyperfibrinolysis (HF) (LY30 >=3.0%)	2540 adults with ISS>15, multicenter	US	83%	25	19%	18% with hyperfibrinolysis	Hyperfibrinolysis 34%	ED	NA	Hypotension, penetrating mechanism and higher ISS
David et al. ⁵⁸	2017	2011-2015	Fibrinogen <150mg/dL, INR> 1.5, platelets <100,000 /mm ³	485 adult trauma patients	France	93%	26	Median shock Index 0.76	23%	54%	ED	NA	Vasopressor administration, ISS, shock index, fluid volume
Koami et al. ⁵⁹	2017	2013-2015	JAAM DIC criteria>3	72 adults with blunt trauma, single center	Japan	100%	29 TIC, 12 non-TIC	8%	29% (patients with anti-coagulants and with liver disease not excluded)	19%	ED	NA	shock, ISS, fluid volume
Mistral et al. ⁶⁰	2017	2015-2016	INR>1.1	98 adults with severe trauma (shock, transfusion or high risk injuries)	France	88%	NA	19%	53%	25%	ED	NA	NA
Mador et al. ⁶¹	2017	2007	INR>1.5, Abnormal TEG	628 adult trauma patients, single center	Canada	96%	25	28% required transfusion/first 24 hours	Abnormal TEG 13% INR>1.5	NA	ED	NA	Older age

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Liras et al. ⁶²	2017	2010-2016	TIC: ACT \geq 128s, angle \leq 65°, MA \leq 55 mm, LY30 \geq 3%	956 children with severe trauma, single center	US	72%	14	SBP TIC 120mmHg non-TIC 122mmHg	Overall 57%; TBI 62%; Age \leq 5yrs 65%	Overall 12%; TBI: 31%; Age \leq 5yrs: 11%	ED	No differences in injury patterns between TIC and non-TIC	Overall: field intubation, ED hypotension; TBI: head AIS, field and ED hypotension; Age \leq 5yrs: ED hypotension
Moore et al. ⁶³	2018	2014-2017	INR $>$ 1.3; Abnormal TEG (ACT $>$ 128s, MA $<$ 55mm, Angle \leq 65°, LY30 $>$ 3%)	125 adult trauma patients with pre-hospital shock, single center (RCT)	US	50%	27	100%	Field: INR $>$ 1.3 6%; Abnl TEG 44% ED: INR $>$ 1.3 34%; Abnl 56%	Field INR $>$ 1.3; Abnl TEG 0 ED: INR $>$ 1.3 32%; Abnl TEG 20%	Field and ED	NA	NA
Dwivedi et al. ⁶⁴	2018	2015-2016	Platelets $<$ 100,000/mm ³ , bleeding or clotting time $>$ lab upper range, INR $>$ 1.5, PTT $>$ 10s above control, D-dimer $>$ 0.5 mg/l, and FDP $>$ 10 mg/l	152 adults and 48 children with TBI, single center	India	97% in adults, 96% in children	NA	NA	Adult 61%; Children 29%	Adult 63%; Children 50%	ED	TBI	GCS \leq 8
Sperry et al. ⁶⁵	2018	2014-2017	INR $>$ 1.3; Abnormal TEG (ACT $>$ 128s, MA $<$ 55mm, Angle \leq 65°, Hyperfibrinolysis LY30 $>$ 3%)	501 air-transported adults in shock, multicenter (RCT)	US	82%	22	100%	INR $>$ 1.3 45%; Abnl TEG 59%	ED: INR $>$ 1.3 34%; Abnl TEG 29%	ED	NA	NA

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Fröhlich et al. ⁶⁶	2019	2002-2013	Quick's value < 70%, platelets < 100,000/mm ³	61,212 adults, multicenter	Germany	95%	22 in 2002 and 18 in 2013	Mean ED SBP 124mmHg in 2002; 132mmHg in 2013	25%	NA	ED	NA	NA
Baksaas-Aasen et al. ⁶⁷	2019	2008-2014	INR > 1.2	2287 trauma activation patients, multicenter	6 major trauma centers in 5 countries: United Kingdom, Denmark, Germany, Netherlands, Norway	85%	13	15% required >3 RBC/12 hours	7%	NA	ED	NA	NA
Sumistawski et al. ⁶⁸	2019	2011-2017	Conventional: INR ≥ 1.4 and/or PTT ≥ 35 s TEG: ACT >128 s, angle <65°, or MA <55 mm	839 trauma activation patients, multicenter	US	51%	17	Base deficit between 4 (no TIC) and 7mEq/L (TIC)	33%	26%	ED	TBI associated with TIC	NA
Ishii et al. ⁶⁹	2019	2013-2015	Fibrinogen <130mg/dL, D-dimer >=118µg/mL	666 adults with blunt trauma, multicenter	Japan	100%	26	20%	17%	44%	ED	TBI more frequent in TIC	Overall: field intubation, ED hypotension; TBI: head AIS, field and ED hypotension;

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Albert et al. ⁷⁰	2019	2014-2016	INR) ≥ 1.27, PT ≥ 16.7s, PTT ≥ 28.8s	120 patients with isolated TBI, age 20-50 years, single center	India	NA	NA	28%	42%	44%	ED	NA	NA
Samuels et al. ⁷¹	2019	2014-2018	TEG ACT>128 s, MA <55 mm, angle <65°; LY30 > 7.6% Conventional coagulation assays: INR>1.3 and PTT>30s	572 trauma activation patients with New Injury Severity Score>8, single center	US	82%	NISS 25 to 43 depending on absence/presence of TBI	Median SBP 105 to 134mmHg depending on absence/presence of SBP	23% INR>1.3; 32% PTT>30s; 37% ACT>128s; 19% MA<55mm; 19% Angle<65°; 10% Hyperfibrinolysis	NA	field or ED	Isolated TBI associated with delayed clot formation	TBI associated with delay in clot initiation, fibrin cross-linking, hypofibrinogenemia, deficiencies in the intrinsic pathway, no abnormalities in fibrinolysis
Leeper et al. ⁷²	2019	2015-2019	hyperfibrinolysis LY30 >=3.0%	285 children, single center	US	75%	17	4%	19% with hyperfibrinolysis	Hyperfibrinolysis 32%	ED	NA	NA

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Cohen et al. ⁷³	2019	2012-2013	INR > 1.2, EXTEM A5 <=35mm, EXTEM LI30 < 97%	40 patients: US and Coalition military personnel identified as having war injuries resulting in activation of DCR	Craig Air Force Theater Hospital – Bagram, Kandahar NATO Hospital in the Afghaniestan Theater	IED 80%; rocket propelled grenade 8%	22	100%	70%	NA	ED	NA	NA
Thorn et al. ⁷⁴	2019	2012-2016	INR > 1.5 or PTT > 60s	15370 adults with ISS>15, multicenter	Germany	93%	25	Median SBP TIC 116mmHg non-TIC 130mmHg	11%	NA	ED	NA	Modified COAST score: on-scene > 45 min or pre-hospital time > 90 min; low prehospital SBP; low ED temperature; pre-hospital thorax drain; abdominal AIS > 3.
Reed et al. ⁷⁵	2019	2012-2017	Age-adjusted abnormal values of INR, PTT and fibrinogen	156 children (<14yrs), 1524 adults, 303 older adults (>=65years), single center 545 pediatric polytrauma patients, single center	US	75%	17	Median Base Excess -3 mEq/l	Abnormal INR 37%; Abnormal PTT 6%; Abnormal fibrinogen 24%	Abnormal INR 14%; Abnormal PTT 52%; Abnormal fibrinogen 15%	ED	NA	Penetrating injury, ISS, Hematocrit, age<14yrs or >=65yrs
Nair et al. ⁷⁶	2020	2006-2015	INR>1.2	polytrauma patients, single center	US	91%	TIC ISS 25; non-	NA	16%	26%	ED	NA	Non-blunt trauma (penetrating or

Study	Publication year	Study period	TIC definition	Population	Country	Blunt trauma	ISS	Hypotension/Shock	TIC Incidence	TIC Case-fatality	Timing of TIC	Injury pattern	Risk Factors for TIC
Driessen et al. ⁷⁷	2020	2009-2016	Quick's value < 70% (INR > 1.4) or PTT ≥ 40 or platelet ≤ 100,000/mm ³	733 children, multicenter	Germany	92 to 97%	TIC ISS 9 27 to 36	NA	14%	42 to 49%	ED	TBI	unknown), ISS
James et al. ⁷⁸	2020	2010-2016	Age-adjusted abnormal values of INR, PT, PTT and platelets	155 children with isolated, blunt TBI, single center	Singapore	100%	16	14%	21%	24%	ED	Isolated TBI	AIS head, hypotension at triage

Quick's value of 70% equates to an INR of 1.4⁶⁶; AIS: Abbreviated Injury Scale; APACHE II: Acute Physiology and Chronic Health Evaluation Score II; BMI: body mass index; ED: emergency department; FDP: fibrin degradation products; GCS: Glasgow coma scale; ISS: injury severity score; NA: not available; PT: prothrombin time; PTT: activated partial thromboplastin time; SAH: subarachnoid hemorrhage; TBI: traumatic brain injury; TRU: trauma resuscitation unit; TT: thrombin time.

JAAM DIC criteria: (1) SIRS ≥ 3 points, (2) platelet < 80,000/mm³ or (3) platelet < 120,000/mm³, (4) INR ≥ 1.2, (5) fibrinogen and fibrin degradation products (FDP) ≥ 25 µg/mL or (6) FDP ≥ 10 µg/mL.

References

1. Sauaia A, Moore FA, Moore EE, et al. Epidemiology of trauma deaths: a reassessment. *J Trauma*. 1995;38(2):185-193.
2. Shackford SR, Mackerzie RC, Holbrook TL, et al. The epidemiology of traumatic death. A population-based analysis. *Arch Surg*. 1993;128(5):571-575.
3. Stewart RM, Myers JG, Dent DL, et al. Seven hundred fifty-three consecutive deaths in a level I trauma center: the argument for injury prevention. *J Trauma*. 2003;54(1):66-70; discussion 70-61.
4. Tien HC, Spencer F, Tremblay LN, Rizoli SB, Brenneman FD. Preventable deaths from hemorrhage at a level I Canadian trauma center. *J Trauma*. 2007;62(1):142-146.
5. Evans JA, van Wessem KJ, McDougall D, Lee KA, Lyons T, Balogh ZJ. Epidemiology of traumatic deaths: comprehensive population-based assessment. *World J Surg*. 2010;34(1):158-163.
6. Consunji RJ, Serrato Marinas JP, Aspuria Maddumba JR, Dela Paz DA, Jr. A profile of deaths among trauma patients in a university hospital: the Philippine experience. *J Inj Violence Res*. 2011;3(2):85-89.
7. Soreide K, Kruger AJ, Vardal AL, Ellingsen CL, Soreide E, Lossius HM. Epidemiology and contemporary patterns of trauma deaths: changing place, similar pace, older face. *World J Surg*. 2007;31(11):2092-2103.
8. Kahl JE, Calvo RY, Sise MJ, Sise CB, Thorndike JF, Shackford SR. The changing nature of death on the trauma service. *J Trauma Acute Care Surg*. 2013;75(2):195-201.
9. Kleber C, Giesecke MT, Tsokos M, et al. Overall distribution of trauma-related deaths in Berlin 2010: advancement or stagnation of German trauma management? *World J Surg*. 2012;36(9):2125-2130.
10. Trajano AD, Pereira BM, Fraga GP. Epidemiology of in-hospital trauma deaths in a Brazilian university hospital. *BMC Emerg Med*. 2014;14:22.
11. Davis JS, Satahoo SS, Butler FK, et al. An analysis of prehospital deaths: Who can we save? *J Trauma Acute Care Surg*. 2014;77(2):213-218.
12. Roberts DJ, Harzan C, Kirkpatrick AW, et al. One thousand consecutive in-hospital deaths following severe injury: Has the etiology of traumatic inpatient death changed in Canada? *Canadian journal of surgery Journal canadien de chirurgie*. 2018;61(3):150-152.
13. Arslan ED, Kaya E, Sonmez M, et al. Assessment of traumatic deaths in a level one trauma center in Ankara, Turkey. *Eur J Trauma Emerg Surg*. 2015;41(3):319-323.
14. Oyeniyi BT, Fox EE, Scerbo M, Tomasek JS, Wade CE, Holcomb JB. Trends in 1029 trauma deaths at a level I trauma center: Impact of a bleeding control bundle of care. *Injury*. 2017;48(1):5-12.
15. Drake SA, Holcomb JB, Yang Y, et al. Establishing a Regional Trauma Preventable/Potentially Preventable Death Rate. *Ann Surg*. 2018.
16. Jochems D, Leenen LPH, Hietbrink F, Houwert RM, van Wessem KJP. Increased reduction in exsanguination rates leaves brain injury as the only major cause of death in blunt trauma. *Injury*. 2018;49(9):1661-1667.
17. Calicut RA, Kornblith LZ, Conroy AS, et al. The why and how our trauma patients die: A prospective Multicenter Western Trauma Association study. *J Trauma Acute Care Surg*. 2019;86(5):864-870.
18. Raum MR, Bouillon B, Rixen D, et al. The Prognostic Value of Prothrombin Time in Predicting Survival after Major Trauma: a Prospective Analysis of 1,351 Patients from the German Trauma Registry. *European Journal of Trauma*. 2001;27(3):110-116.
19. Keller MS, Fendya DG, Weber TR. Glasgow Coma Scale predicts coagulopathy in pediatric trauma patients. *Semin Pediatr Surg*. 2001;10(1):12-16.
20. Brohi K, Singh J, Heron M, Coats T. Acute traumatic coagulopathy. *J Trauma Acute Care Surg*. 2003;54(6):1127-1130.
21. MacLeod JB, Lynn M, McKenney MG, Cohn SM, Murtha M. Early coagulopathy predicts mortality in trauma. *J Trauma Acute Care Surg*. 2003;55(1):39-44.
22. Carrick MM, Tyroch AH, Youens CA, Handley T. Subsequent development of thrombocytopenia and coagulopathy in moderate and severe head injury: support for serial laboratory examination. *J Trauma*. 2005;58(4):725-729; discussion 729-730.

23. Affonseca CA, Carvalho LF, Guerra SD, Ferreira AR, Goulart EM. Coagulation disorder in children and adolescents with moderate to severe traumatic brain injury. *J Pediatr (Rio J)*. 2007;83(3):274-282.
24. Rugeri L, Levriat A, David JS, et al. Diagnosis of early coagulation abnormalities in trauma patients by rotation thrombelastography. *J Thromb Haemost*. 2007;5(2):289-295.
25. Maegele M, Lefering R, Yucel N, et al. Early coagulopathy in multiple injury: an analysis from the German Trauma Registry on 8724 patients. *Injury*. 2007;38(3):298-304.
26. Niles SE, McLaughlin DF, Perkins JG, et al. Increased mortality associated with the early coagulopathy of trauma in combat casualties. *J Trauma*. 2008;64(6):1459-1463; discussion 1463-1455.
27. Kashuk JL, Moore EE, Johnson JL, et al. Postinjury life threatening coagulopathy: is 1:1 fresh frozen plasma-packed red blood cells the answer? *J Trauma*. 2008;65(2):261-270; discussion 270-261.
28. Talving P, Benfield R, Hadjizacharia P, Inaba K, Chan LS, Demetriades D. Coagulopathy in severe traumatic brain injury: a prospective study. *J Trauma*. 2009;66(1):55-61; discussion 61-52.
29. Frith D, Goslings JC, Gaarder C, et al. Definition and drivers of acute traumatic coagulopathy: clinical and experimental investigations. *J Thromb Haemost*. 2010;8(9):1919-1925.
30. Wafaisade A, Lefering R, Tjardes T, et al. Acute coagulopathy in isolated blunt traumatic brain injury. *Neurocrit Care*. 2010;12(2):211-219.
31. Wafaisade A, Wutzler S, Lefering R, et al. Drivers of acute coagulopathy after severe trauma: a multivariate analysis of 1987 patients. *Emerg Med J*. 2010;27(12):934-939.
32. Dirks J, Jørgensen H, Jensen CH, Ostrowski SR, Johansson PI. Blood product ratio in acute traumatic coagulopathy--effect on mortality in a Scandinavian level I trauma centre. *Scand J Trauma Resusc Emerg Med*. 2010;18:65.
33. Jeger V, Urwyler N, Zimmermann H, Exadaktylos AK. Trauma-induced coagulopathy in severely injured patients: knowledge lost in translation? *Emerg Med J*. 2010;27(7):551-552.
34. Tauber H, Innerhofer P, Breitkopf R, et al. Prevalence and impact of abnormal ROTEM® assays in severe blunt trauma: results of the 'Diagnosis and Treatment of Trauma-Induced Coagulopathy (DIA-TRE-TIC) study'. *Br J Anaesth*. 2011;107(3):378-387.
35. Johansson PI, Sørensen AM, Perner A, et al. Disseminated intravascular coagulation or acute coagulopathy of trauma shock early after trauma? An observational study. *Crit Care*. 2011;15(6):R272.
36. Greuters S, van den Berg A, Franschman G, et al. Acute and delayed mild coagulopathy are related to outcome in patients with isolated traumatic brain injury. *Crit Care*. 2011;15(1):R2.
37. Patregnani JT, Borgman MA, Maegele M, Wade CE, Blackburne LH, Spinella PC. Coagulopathy and shock on admission is associated with mortality for children with traumatic injuries at combat support hospitals. *Pediatr Crit Care Med*. 2012;13(3):273-277.
38. Hendrickson JE, Shaz BH, Pereira G, et al. Coagulopathy is prevalent and associated with adverse outcomes in transfused pediatric trauma patients. *J Pediatr*. 2012;160(2):204-209.e203.
39. Mitra B, Cameron PA, Mori A, Fitzgerald M. Acute coagulopathy and early deaths post major trauma. *Injury*. 2012;43(1):22-25.
40. Peiniger S, Nienaber U, Lefering R, et al. Glasgow Coma Scale as a predictor for hemocoagulative disorders after blunt pediatric traumatic brain injury. *Pediatr Crit Care Med*. 2012;13(4):455-460.
41. Genét GF, Johansson PI, Meyer MA, et al. Trauma-induced coagulopathy: standard coagulation tests, biomarkers of coagulopathy, and endothelial damage in patients with traumatic brain injury. *J Neurotrauma*. 2013;30(4):301-306.
42. Chhabra G, Sharma S, Subramanian A, Agrawal D, Sinha S, Mukhopadhyay AK. Coagulopathy as prognostic marker in acute traumatic brain injury. *J Emerg Trauma Shock*. 2013;6(3):180-185.
43. Xu SX, Wang L, Zhou GJ, Zhang M, Gan JX. Risk factors and clinical significance of trauma-induced coagulopathy in ICU patients with severe trauma. *Eur J Emerg Med*. 2013;20(4):286-290.

44. Alexiou GA, Lianos G, Fotakopoulos G, Michos E, Pachatouridis D, Voulgaris S. Admission glucose and coagulopathy occurrence in patients with traumatic brain injury. *Brain Inj*. 2014;28(4):438-441.
45. Neal MD, Brown JB, Moore EE, et al. Prehospital use of nonsteroidal anti-inflammatory drugs (NSAIDs) is associated with a reduced incidence of trauma-induced coagulopathy. *Ann Surg*. 2014;260(2):378-382.
46. MacLeod JB, Winkler AM, McCoy CC, Hillyer CD, Shaz BH. Early trauma induced coagulopathy (ETIC): prevalence across the injury spectrum. *Injury*. 2014;45(5):910-915.
47. Moore HB, Moore EE, Gonzalez E, et al. Hyperfibrinolysis, physiologic fibrinolysis, and fibrinolysis shutdown: the spectrum of postinjury fibrinolysis and relevance to antifibrinolytic therapy. *J Trauma Acute Care Surg*. 2014;77(6):811-817; discussion 817.
48. Tonglet ML, Minon JM, Seidel L, Poplavsky JL, Vergnion M. Prehospital identification of trauma patients with early acute coagulopathy and massive bleeding: results of a prospective non-interventional clinical trial evaluating the Trauma Induced Coagulopathy Clinical Score (TICCS). *Crit Care*. 2014;18(6):648.
49. Kutcher ME, Howard BM, Sperry JL, et al. Evolving beyond the vicious triad: Differential mediation of traumatic coagulopathy by injury, shock, and resuscitation. *J Trauma Acute Care Surg*. 2015;78(3):516-523.
50. de Oliveira Manoel AL, Neto AC, Veigas PV, Rizoli S. Traumatic brain injury associated coagulopathy. *Neurocrit Care*. 2015;22(1):34-44.
51. Peltan ID, Vande Vusse LK, Maier RV, Watkins TR. An International Normalized Ratio-Based Definition of Acute Traumatic Coagulopathy Is Associated With Mortality, Venous Thromboembolism, and Multiple Organ Failure After Injury. *Crit Care Med*. 2015;43(7):1429-1438.
52. Khan S, Davenport R, Raza I, et al. Damage control resuscitation using blood component therapy in standard doses has a limited effect on coagulopathy during trauma hemorrhage. *Intensive Care Med*. 2015;41(2):239-247.
53. Hagemo JS, Christiaans SC, Stanworth SJ, et al. Detection of acute traumatic coagulopathy and massive transfusion requirements by means of rotational thromboelastometry: an international prospective validation study. *Crit Care*. 2015;19(1):97.
54. Holcomb JB, Tilley BC, Baraniuk S, et al. Transfusion of plasma, platelets, and red blood cells in a 1:1:1 vs a 1:1:2 ratio and mortality in patients with severe trauma: the PROPPR randomized clinical trial. *JAMA*. 2015;313(5):471-482.
55. Strumwasser A, Speer AL, Inaba K, et al. The impact of acute coagulopathy on mortality in pediatric trauma patients. *J Trauma Acute Care Surg*. 2016;81(2):312-318.
56. Peltan ID, Rowhani-Rahbar A, Vande Vusse LK, et al. Development and validation of a prehospital prediction model for acute traumatic coagulopathy. *Crit Care*. 2016;20(1):371-371.
57. Moore HB, Moore EE, Liras IN, et al. Acute Fibrinolysis Shutdown after Injury Occurs Frequently and Increases Mortality: A Multicenter Evaluation of 2,540 Severely Injured Patients. *J Am Coll Surg*. 2016;222(4):347-355.
58. David JS, Voiglio EJ, Cesario E, et al. Prehospital parameters can help to predict coagulopathy and massive transfusion in trauma patients. *Vox Sang*. 2017;112(6):557-566.
59. Koami H, Sakamoto Y, Yamada KC, et al. What factor within the Japanese Association for Acute Medicine (JAAM) disseminated intravascular coagulation (DIC) criteria is most strongly correlated with trauma induced DIC? A retrospective study using thromboelastometry in a single center in Japan. *Eur J Trauma Emerg Surg*. 2017;43(4):431-438.
60. Mistral T, Boué Y, Bosson JL, et al. Performance of point-of-care international normalized ratio measurement to diagnose trauma-induced coagulopathy. *Scand J Trauma Resusc Emerg Med*. 2017;25(1):59.
61. Mador B, Nascimento B, Hollands S, Rizoli S. Blood transfusion and coagulopathy in geriatric trauma patients. *Scand J Trauma Resusc Emerg Med*. 2017;25(1):33-33.
62. Liras IN, Caplan HW, Stensballe J, Wade CE, Cox CS, Cotton BA. Prevalence and Impact of Admission Acute Traumatic Coagulopathy on Treatment Intensity, Resource Use, and Mortality: An Evaluation of 956 Severely Injured Children and Adolescents. *J Am Coll Surg*. 2017;224(4):625-632.
63. Moore EE, Chin TL, Chapman MC, et al. Plasma first in the field for postinjury hemorrhagic shock. *Shock*. 2014;41 Suppl 1:35-38.
64. Dwivedi AK, Sharma A, Sinha VD. Comparative Study of Derangement of Coagulation Profile between Adult and Pediatric Population in Moderate to Severe Traumatic Brain Injury: A Prospective Study in a Tertiary Care Trauma Center. *Asian J Neurosurg*. 2018;13(4):1123-1127.

65. Sperry JL, Guyette FX, Brown JB, et al. Prehospital Plasma during Air Medical Transport in Trauma Patients at Risk for Hemorrhagic Shock. *N Engl J Med*. 2018;379(4):315-326.
66. Fröhlich M, Mutschler M, Caspers M, et al. Trauma-induced coagulopathy upon emergency room arrival: still a significant problem despite increased awareness and management? *Eur J Trauma Emerg Surg*. 2019;45(1):115-124.
67. Baksaas-Aasen K, Van Dieren S, Balvers K, et al. Data-driven Development of ROTEM and TEG Algorithms for the Management of Trauma Hemorrhage: A Prospective Observational Multicenter Study. *Ann Surg*. 2019;270(6):1178-1185.
68. Sumiowski JJ, Christie SA, Kornblith LZ, et al. Discrepancies between conventional and viscoelastic assays in identifying trauma-induced coagulopathy. *Am J Surg*. 2019;217(6):1037-1041.
69. Ishii K, Kinoshita T, Kiridume K, et al. Impact of initial coagulation and fibrinolytic markers on mortality in patients with severe blunt trauma: a multicentre retrospective observational study. *Scand J Trauma Resusc Emerg Med*. 2019;27(1):25.
70. Albert V, Arulsevi S, Agrawal D, Pati HP, Pandey RM. Early posttraumatic changes in coagulation and fibrinolysis systems in isolated severe traumatic brain injury patients and its influence on immediate outcome. *Hematol Oncol Stem Cell Ther*. 2019;12(1):32-43.
71. Samuels JM, Moore EE, Silliman CC, et al. Severe traumatic brain injury is associated with a unique coagulopathy phenotype. *J Trauma Acute Care Surg*. 2018.
72. Leeper CM, Strotmeyer SJ, Neal MD, Gaines BA. Window of Opportunity to Mitigate Trauma-induced Coagulopathy: Fibrinolysis Shutdown not Prevalent Until 1 Hour Post-injury. *Ann Surg*. 2019;270(3):528-534.
73. Cohen J, Scorer T, Wright Z, et al. A prospective evaluation of thromboelastometry (ROTEM) to identify acute traumatic coagulopathy and predict massive transfusion in military trauma patients in Afghanistan. *Transfusion*. 2019;59(S2):1601-1607.
74. Thorn S, Lefering R, Maegele M, Gruen RL, Mitra B. Early prediction of acute traumatic coagulopathy: a validation of the COAST score using the German Trauma Registry. *Eur J Trauma Emerg Surg*. 2019.
75. Reed CR, Williamson H, Vatsaas C, et al. Higher mortality in pediatric and adult trauma patients with traumatic coagulopathy, using age-adjusted diagnostic criteria. *Surgery*. 2019;165(6):1108-1115.
76. Nair A, Flori H, Cohen MJ. Characterization of organ dysfunction and mortality in pediatric patients with trauma with acute traumatic coagulopathy. *Trauma surgery & acute care open*. 2020;5(1):e000382.
77. Driessen A, Wafaisade A, Lefering R, et al. Mechanism, frequency, transfusion and outcome of severe trauma in coagulopathic paediatric patients. *Eur J Trauma Emerg Surg*. 2020:1-9.
78. James V, Chong SL, Shetty SS, Ong GY. Early coagulopathy in children with isolated blunt head injury is associated with mortality and poor neurological outcomes. *J Neurosurg Pediatr*. 2020:1-7.