

doi: 10.13241/j.cnki.pmb.2020.22.041

限制性输血与开放性输血对急性上消化道出血患者凝血功能、血液流变学及预后的影响 *

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摘要 目的:探讨限制性输血与开放性输血对急性上消化道出血患者凝血功能、血液流变学及预后的影响。**方法:**选取 2018 年 1 月~2020 年 1 月期间我院收治的急性上消化道出血患者 80 例,根据随机数字表法分为对照组($n=40$)和研究组($n=40$),对照组患者输血方式采用开放性输血,研究组患者输血方式采用限制性输血,比较两组患者治疗 24 h 后、48 h 后、72 h 后的止血率。统计两组患者死亡率、疗效、再出血率和不良事件发生率。比较两组治疗前、治疗 72 h 后的 Blatchford 评分及凝血功能指标:凝血酶原时间(PT)、活化部分凝血活酶时间(APTT)以及血液流变学指标:全血黏度、血浆黏度、红细胞比容。**结果:**研究组治疗 24 h 后的止血率高于对照组 ($P<0.05$);两组治疗 48 h 后、治疗 72 h 后的止血率组间比较差异无统计学意义 ($P>0.05$)。两组治疗 72 h 后 Blatchford 评分均下降,且研究组低于对照组($P<0.05$)。两组治疗 72 h 后 PT、APTT 均升高,且研究组高于对照组($P<0.05$)。两组死亡率比较差异无统计学意义($P>0.05$);研究组不良事件总发生率、再出血率均低于对照组($P<0.05$)。研究组治疗后的临床总有效率高于对照组($P<0.05$)。两组治疗 72 h 后全血黏度、血浆黏度、红细胞比容均升高,且研究组高于对照组($P<0.05$)。**结论:**与开放性输血相比,急性上消化道出血患者采用限制性输血,可迅速止血,有效防止患者凝血功能紊乱及血液流变学异常,同时还可减少不良事件总发生率、再出血率,可进一步改善患者预后。

关键词:限制性输血;开放性输血;急性上消化道出血;凝血功能;血液流变学;预后

中图分类号:R573.2;R457.1 文献标识码:A 文章编号:1673-6273(2020)22-4382-05

The Effect of Restrictive and Open Blood Transfusion on Coagulation, Hemorheology and Prognosis in Patients with Acute Upper Gastrointestinal Hemorrhage*

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ABSTRACT Objective: To explore the effect of restrictive and open blood transfusion on coagulation, hemorheology and prognosis in patients with acute upper gastrointestinal hemorrhage. **Methods:** 80 patients with acute upper gastrointestinal bleeding who were admitted to our hospital from January 2018 to January 2020 were selected, patients were divided into control group ($n=40$) and study group ($n=40$) according to the method of random number table. The open blood transfusion was used in the control group, and the restrictive blood transfusion was used in the study group. The mortality rate, curative effect, rebleeding rate and adverse events of the two groups were counted. The hemostasis rate of the two groups was compared at 24 hours, 48 hours and 72 hours after treatment. The Blatchford score and coagulation indexes: prothrombin time (PT) and activated partial thromboplastin time (APTT), Hemorheology indexes: whole blood viscosity, plasma viscosity, hematocrit of the two groups were compared before and 72 hours after treatment. **Results:** The hemostasis rate of the study group was higher than that of the control group ($P<0.05$), and there were no significant differences in hemostasis rate between the two groups at 48h and 72h after treatment ($P>0.05$). The Blatchford score of the two groups decreased at 72 hours after treatment, and that of the study group was lower than that of the control group ($P<0.05$). PT and APTT increased in the two groups 72 hours after treatment, and the study group was higher than the control group ($P<0.05$). There was no significant difference in mortality between the two groups ($P>0.05$). The total incidence of adverse events and rebleeding rate in the study group were lower than those in the control group ($P<0.05$). The total clinical effective rate of the study group was higher than that of the control group ($P<0.05$). After 72 hours of treatment, the whole blood viscosity, plasma viscosity and RBC specific volume of the two groups were all increased, and the study group was higher than the control group ($P<0.05$). **Conclusion:** Compared with the open blood transfusion, the restrictive blood transfusion in the patients with acute upper gastrointestinal hemorrhage can quickly stop bleeding, effectively prevent the disorder

* 基金项目:国家自然科学基金项目(82369526);陕西省自然科学基金项目(2015831675J)

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(收稿日期:2020-04-26 接受日期:2020-05-22)

of coagulation function, reduce the total incidence of adverse events and rebleeding rate, and further improve the prognosis of the patients.

Key words: Restrictive blood transfusion; Open blood transfusion; Acute upper gastrointestinal hemorrhage; Coagulation; Hemorheology; Prognosis

Chinese Library Classification(CLC): R573.2; R457.1 Document code: A

Article ID: 1673-6273(2020)22-4382-05

前言

急性上消化道出血是指屈氏韧带以上的食管、胃、十二指肠等部位病变引起的急性出血，该病临床主要表现为呕血、黑便、贫血等症状，若未能及时予以治疗，可发生失血性循环衰竭，危及患者性命^[1-3]。现临床针对该病的治疗主要为输血治疗，即根据患者出血量的多少，给予等量全血以纠正机体贫血状态^[4-5]。开放性输血是临床常见的输血方式，但近年来的实践表明不少急性上消化道出血患者经输血治疗后易出现再次出血的情况^[6-7]。限制性输血作为一种新兴的输血方式，在较低的血红蛋白(Hb)开始输血，补充的上限也有所不同，可有效节约血液资源，但有关其在急性上消化道出血的疗效尚存在一定的争议^[8-9]。鉴于此，本研究通过探讨不同输血方式对急性上消化道出血患者凝血功能、止血效果及预后的影响，以期为临床急性上消化道出血的治疗提供数据支持，整理报道如下。

1 资料与方法

1.1 一般资料

选取 2018 年 1 月 ~2020 年 1 月期间我院收治的急性上消化道出血患者 80 例，此次研究已通过我院伦理学委员会批准进行。患者纳入标准：(1)均为首次发病；(2)经胃镜检查确诊；(3)知情本研究且签署了同意书。排除标准：(1)合并严重心肝肾等脏器功能障碍者；(2)合并精神障碍无法正常沟通交流者；(3)近期有外科手术史者；(4)合并凝血功能障碍者；(5)妊娠或哺乳期手术者；(6)符合手术指征需立刻给予手术治疗者。上述患者根据随机数字表法分为对照组(n=40)和研究组(n=40)，其中对照组男 28 例，女 12 例，年龄 24~58 岁，平均(39.67±4.52)岁；体质量指数 21~26 kg/m²，平均(23.67±0.74)kg/m²。研究组男 26 例，女 14 例，年龄 25~61 岁，平均(40.19±5.27)岁；体质量指数 22~26 kg/m²，平均(23.91±0.83)kg/m²。两组一般资料比较差异无统计学差异($P>0.05$)。

1.2 方法

所有患者入院后均给予相关常规基础治疗，包括留置胃管、洗胃、禁食等、药物止血、护胃等。在此过程中，收缩压控制在 80~90 mmHg，并开放静脉通路进行扩容。随后对照组给予

开放性输血方式，即当患者 Hb<90 g/L 时，输入全部失血量的红细胞悬液，输血目标为 Hb 上升至 90~110 g/L。研究组行限制性输血方式，即当患者 Hb<70 g/L 开始输血，无需输入全部失血量的红细胞悬液，为失血量的 30%~50% 即可，输血目标为 Hb 上升至 70~90 g/L。

1.3 观察指标

(1)记录两组患者治疗 24 h 后、48 h 后、72 h 后的止血率。(2)统计两组患者死亡率、再出血率和不良事件发生率，其中再出血指征为患者止血成功后再出现呕血、血容量不足、新鲜黑便情况。不良事件包括发热、输血反应、细菌感染、肺水肿等。(3)于治疗前、治疗 72 h 后采用 Blatchford 评分系统评价患者病情程度，Blatchford 评分系统包括收缩压、血尿素氮、血红蛋白、黑便、脉率、晕厥、肝脏疾病、心力衰竭等项目组成，评分越高病情程度越严重，其中高危： ≥ 6 分，低危： <6 分^[10]。(4)于治疗前、治疗 72 h 后抽取患者肘静脉血 4 mL，以 3000 r/min 的速度离心 15 分钟，血清分离后在 2h 内进行检测，应用 CA1500 全自动血凝仪(日本希森美康公司)对患者的凝血功能指标凝血酶原时间(PT)、活化部分凝血活酶时间(APTT)进行检测。采用普利生 LBY-N7500B 型全自动血液流变仪检测全血黏度、血浆黏度、红细胞比容。(5)记录两组临床总有效率。疗效判定标准：痊愈：治疗 24 h 后出血停止，症状及体征基本消失，体温恢复正常；好转：治疗 48 h 后出血停止，症状及体征基本消失，体温恢复正常；无效：治疗 72 h 后出血未停止，症状及体征未见改善。总有效率 = 痊愈率 + 好转率。

1.4 统计学方法

采用 SPSS25.0 进行数据分析，计数资料以率表示，行卡方检验，计量资料以均值± 标准差的形式表示，行 t 检验。检验标准设置为 $\alpha=0.05$ 。

2 结果

2.1 两组止血率比较

研究组治疗 24 h 后的止血率高于对照组($P<0.05$)；两组治疗 48 h 后、治疗 72 h 后的止血率组间比较差异无统计学意义($P>0.05$)；详见表 1。

表 1 两组止血率比较 [例(%)]

Table 1 Comparison of hemostasis rate between the two groups [n(%)]

Groups	24 hours after treatment	48 hours after treatment	72 hours after treatment
Control group(n=40)	4(10.00)	21(52.50)	33(82.50)
Study group(n=40)	14(35.00)	25(62.50)	37(92.50)
χ^2	7.168	0.818	1.829
P	0.007	0.366	0.176

2.2 两组 Blatchford 评分比较

对照组治疗前 Blatchford 评分为(6.19± 0.47)分,治疗 72 h 后 Blatchford 评分为(3.27± 0.38)分;研究组治疗前 Blatchford 评分为(6.25± 0.41)分,治疗 72 h 后 Blatchford 评分为(1.96± 0.37)分;两组治疗 72 h 后 Blatchford 评分均下降(t 对照组

=30.555, $P=0.000$; t 研究组 =49.869, $P=0.000$),且研究组低于对照组($t=12.861$, $P=0.000$)。

2.3 两组凝血功能指标比较

两组治疗前 PT、APTT 比较无差异($P>0.05$);两组治疗 72 h 后 PT、APTT 均升高,且研究组高于对照组($P<0.05$);详见表2。

表 2 两组凝血功能指标比较($\bar{x} \pm s$)

Table 2 Comparison of coagulation indexes between the two groups($\bar{x} \pm s$)

Groups	PT(s)		APTT(s)	
	Before treatment	72 hours after treatment	Before treatment	72 hours after treatment
Control group(n=40)	11.90± 1.13	14.03± 1.34*	28.29± 3.47	31.83± 3.24*
Study group(n=40)	11.72± 1.26	16.52± 1.47*	27.91± 4.52	35.32± 3.51*
t	0.673	7.917	0.422	4.621
P	0.503	0.000	0.674	0.000

Note: compared with before treatment, * $P<0.05$.

2.4 两组预后指标比较

两组死亡率比较差异无统计学意义($P>0.05$);研究组不良

事件总发生率、再出血率均低于对照组($P<0.05$);详见表 3。

表 3 两组预后指标比较 例(%)

Table 3 Comparison of prognostic indexes between the two groups n(%)

Groups	Adverse events					Mortality rate	Rebleeding rate
	Fever	Transfusion reaction	Bacterial infection	Pulmonary edema	Total incidence rate		
Control group(n=40)	2(5.00)	6(15.00)	4(10.00)	3(7.50)	15(37.50)	2(5.00)	18(45.00)
Study group(n=40)	1(2.50)	1(2.50)	1(2.50)	1(2.50)	4(10.00)	1(2.50)	7(17.50)
χ^2	8.352	0.346					7.040
P	0.004	0.556					0.008

2.5 两组总有效率比较

研究组治疗后的临床总有效率高于对照组($P<0.05$);详见

表 4 两组总有效率比较 例(%)

Table 4 Comparison of total effective rate between the two groups n(%)

Groups	Recovery	Better	Invalid	Total efficiency
Control group(n=40)	9(22.50)	19(47.50)	12(30.00)	28(70.00)
Study group(n=40)	14(35.00)	23(57.50)	3(7.50)	37(92.50)
χ^2				6.646
P				0.010

2.6 两组血液流变学指标比较

两组治疗前全血黏度、血浆黏度、红细胞比容比较无差异($P>0.05$);两组治疗 72 h 后全血黏度、血浆黏度、红细胞比容均升高,且研究组高于对照组($P<0.05$);详见表 2。

压下降、凝血功能紊乱,进而出现休克或猝死等不良事件^[13-15]。据以往报道统计^[16],急性上消化道大出血的死亡率可高达 10%,60 岁以上老人出血死亡率可高达 30%~50%。为了有效纠正贫血以及失血,输血疗法是临床治疗急性上消化道出血的有效急救方法^[17],但现阶段,临床对于输血治疗的具体时机及方法,仍存在一定的争议。以往情况下,临床认为急性失血时 Hb 低于 90 g/L 的患者进行及时的输血就能够起到治疗的效果^[18,19],但大量的输血又会导致体积扩容不足,产生一定的不良反应,同时不必要的输血会增加各种疾病的传播,反而影响输血

3 讨论

急性上消化道出血主要由肝硬化引起的食管胃底静脉曲张破裂、消化性溃疡、糜烂性出血性胃炎以及恶性肿瘤等引起^[11,12]。该病发病急骤,病情进展迅速,短期内即可引起患者血

疗效,降低患者预后效果^[20]。随着医疗研究的深入,限制性输血理念被提出,现已受到不少患者及其医师的喜爱,逐渐广泛应

用于临床治疗中^[21,22]。

表 5 两组血液流变学指标比较($\bar{x} \pm s$)
Table 5 Comparison of hemorheology indexes between the two groups ($\bar{x} \pm s$)

Groups	Whole blood viscosity(mPa/s)		Plasma viscosity(mPa/s)		Hematocrit(%)	
	Before treatment	72 hours after treatment	Before treatment	72 hours after treatment	Before treatment	72 hours after treatment
Control group(n=40)	3.28±0.35	4.06±0.28*	1.51±0.25	2.03±0.24*	36.16±6.23	42.56±5.65*
Study group(n=40)	3.32±0.24	4.53±0.25*	1.57±0.19	2.69±0.37*	36.09±6.74	49.98±7.17*
t	0.596	7.919	1.208	9.465	0.048	5.141
P	0.553	0.000	0.321	0.000	0.962	0.000

Note: Compared with before treatment, *P<0.05.

本次研究结果显示,研究组治疗后的临床总有效率以及止血率高于对照组,提示急性上消化道出血患者采用限制性输血,早期止血率效果优于对照组,这主要是因为限制性输血在患者 Hb<70 g/L 时才开始输血,可最大化激活患者的自我修复调节能力^[23,24]。研究组治疗 72 h 后的 Blatchford 评分低于对照组,可能是因为限制性输血的早期止血效果较好,且当患者止血成功后,医生可对患者具体出血情况进行再评估,实时更新治疗方案,有利于患者的病情改善^[25]。既往研究显示^[26],人体在早期活动性出血尚未彻底止血前,大量且快速的输入血液不仅不能保证重要器官的灌流,反而引起血压升高,并进一步抑制体内凝血因子的活性。从本研究的结果来看,急性上消化道出血患者采用限制性输血,可有效防止患者凝血功能紊乱。限制性输血通过相对少量且渐进的输血形式,在为机体及时的恢复组织器官供血、供氧的同时,还能有效减轻组织器官的负担和压力,使患者循环功能得到明显改善,选择性的收缩皮肤及肌肉血管,从而使血液重新分布^[27]。此外,研究组不良事件总发生率、再出血率均低于对照组,主要是因为限制性输血可以对出血或抢救治疗有更好的控制,进而减少再出血发生率。同时,限制性输血输入血量较少输血时间短,可减少外来血液的过多输入,降低外来血液对机体系统的过度影响,继而减少不良事件总发生率^[28]。急性上消化道出血时,首先引起体内体液分布发生紊乱和变化,当出血量超过一定量时,可引起毛细血管静脉端压力下降,组织间液和细胞外液顺压力向血管内转移,导致血液发生稀释,引起机体红细胞比容下降。而血浆粘度主要由所含的大分子物质纤维蛋白原和球蛋白的量所决定,急性上消化道出血时球蛋白升高,纤维蛋白原明显下降,综合平衡导致血浆粘度不升反降。全血粘度则受血浆粘度、红细胞比容所影响,随着它们的水平下降而下降^[29]。本研究中两组治疗 72 h 后全血粘度、血浆粘度、红细胞比容均升高,且研究组高于对照组,可见急性上消化道出血患者采用限制性输血可有效防止血液流变学异常,可能是因为限制性输血可通过提高少量鲜血增加血液粘度,降低红细胞脆性,改变红细能力,有效恢复机体供养环境,改善机体血液流变学状况^[30]。另两组死亡率方面比较未见差异,可能是两种输血方式均安全可靠,也或许是本次研究样本量偏少,导致结果存在一定的偏倚,后续报道将扩大样本量以获取更为准确的结果。

综上所述,与开放性输血相比,急性上消化道出血患者采用限制性输血,可迅速止血,有效防止患者凝血功能紊乱、血液流变学异常,同时还可减少不良事件总发生率、再出血率,其效果令人满意,可进一步改善患者预后。

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