

Prevalence of Rhesus & Kell phenotypes among blood donors of Bangladesh

Ayesha Khatun, Sheikh Saiful Islam Shaheen, Atiar Rahman, Subarna Saha, Sushanta Kumar Basak

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Department of Transfusion Medicine,
BSMMU, Dhaka (AK, SSI, AR, SS, SKB)

For Correspondence:
Sheikh Saiful Islam Shaheen
Email: drskshaheen@yahoo.com

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Abstract

The Rhesus blood group system is one of the most polymorphic and highly immunogenic systems in humans. Because of its high and strong immunogenicity Rh D antigen testing is mandatory before issuing a compatible blood. There are five major antigens i.e. DCEe in the Rhesus (Rh) blood group system. On the other hand from the immunogenicity point of view Kell antigen is next to the Rh system. Both of them may cause severe hemolytic transfusion reaction and hemolytic disease of fetus and new born. Exposure of Rhesus negative individuals to Rhesus positive blood through transfusion or pregnancy is most likely to stimulate production of Rhesus antibodies. These antibodies may cause Hemolytic Disease of Fetus and Newborn (HDFN) and Delayed Hemolytic Transfusion Reaction (DHTR). Like Rhesus antibodies, Kell antibodies may also cause HDFN and DHTR. So far we know, there is not enough study regarding antigens C, c, E & e of Rh or K, k antigen of Kell blood group system regarding these antigens in the donors in Bangladesh, thereby exposing transfused patients to these antigens negative patients. To determine the phenotype prevalence of the Rh and Kell blood group systems in the blood donors in Bangladesh, a descriptive cross sectional study was done in the laboratory of Department of Transfusion Medicine, Bangabandhu Sheikh Mujib Medical University, during the period of 1st January 2020 to 31 December 2020. Rhesus Phenotype CCDee is highest (48.4%) & CCDEe & ccDEE both are lowest (0.4%). Most probable Rhesus Genotype CDe/CDe (R1R1) is highest (48.4%) and CDe/CDE (R1Rz) & cDE/cDE (R2R2) both are lowest (0.4%). Kell Genotype kk is highest (99.2%) and Kk is lowest (0.8%).

Introduction

Karl Landsteiner and Wiener discovered Rhesus (Rh) antigen in 1940, one of the most polymorphic and immunogenic systems is Rhesus (Rh) system. As it is highly immunogenic, testing for the RhD along with the A, B antigens has been made mandatory in pre-transfusion testing.

There are more than 52 antigens in the Rh blood group discovered at present, but the major ones are D,C,E, c and e. In the Kell blood group system, more than 24 antigens have been discovered, among them Kell (K) and k (Cellano) is the most important.

There is not enough report available regarding the prevalence of these antigens in the people of Bangladesh.

The phenotype of a blood group of an individual is the observable expression of the genes inherited by the person and reflects the

biologic activity of the genes. The presence or absence of antigens on red cells as determined by serological testing represents the phenotype.¹

The phenotype of clinically significant blood group antigens on the donor red blood cells (RBC) is required to be known at times when alloimmunization is particularly undesirable, as in young females, pregnant women, and patients who are expected to require repeated transfusions in life, such as thalassemia or sickle cell disease patients. When selecting blood for transfusion to such patients, it would be useful if we have access to already phenotyped RBCs of donor population so that particular antigen typed blood can be given to such patients to prevent alloimmunization.²

In sub-saharan countries, few practice this systematic search for antigens C, c, e, E and K in the donors and recipients, thereby exposing the transfused patient to high risk of alloimmunization.³

The typing of blood group antigens and determination of phenotype for blood and blood components are a prerequisite for an efficient and safe blood transfusion. Rh antibodies are usually immunoglobulin G, and they are produced early during life.⁴⁻⁶

The knowledge of the distribution of Rhesus antigen in a population is critical in managing a transfusion service in areas such as antenatal serology, paternity testing as well as selecting compatible blood and blood products. Even after Karl Landsteiner's discovery in 1900, transfusion reactions were still prevalent.⁷

Knowledge about the frequency of red cells antigens phenotypes in Ivorian population is important for the creation of a donor data bank and to minimize risks of alloimmunization. This requires the determination of the immunological characteristics of blood products and blood recipients by performing immunohematological analysis such as phenotyping in Rh and Kell blood group systems. Currently, there are thirty-three major blood group systems.⁸

Blood group prevalence plays a role in evolution, genetics research, blood transfusion and organ transplantation. Modern medicine is also working on relationship of blood group with environment.⁹

Study of Rhesus (Rh) blood group antigens, Phenotype and Rh antibodies are very useful in routine and advanced clinical practice in blood transfusion center. Moreover, it can be used for population genetic studies.^{10,11}

Eight percent of D negative patients exposed to D positive red cells may develop anti-D IgG antibodies that may persist for the rest of their lives, which can cause Hemolytic Transfusion Reaction (HTR) and Disease of Fetus and Newborn (HDFN).¹²

Methods

A Descriptive cross-sectional study was conducted from 1st January 2020 to 31st December 2020 in the Department of Transfusion Medicine, BSMMU. A structured questionnaire was used.

Blood donors coming to the Department of Transfusion Medicine, BSMMU fulfilling inclusion criteria were taken as sample by purposive convenient sampling technique. Inclusion criteria were fulfilled for being a blood donor. The participants consented to participate voluntarily in the present study. The data obtained from this study was analyzed for frequencies (absolute) and percentage.

Results

The study comprised of 256 donors. Among the 256 donors, 21-30 years age group donor is more (53.9%) and 41-50 years

age group donor is less (7.4%). Male donor is predominant (89.8%) and Female donor is (10.2%). (Table-I, Table-II)

In case of ABO & Rh (D) blood groups donors, O Rh(D)+ve donor is more (33.2%) & O Rh(D)-ve donor is lowest (0.8%). In this study, Rhesus Phenotype CCDee is highest (48.4%) & CCDEe & ccDEE both are lowest (0.4%). Most probable Rhesus Genotype CDe/CDe (R_1R_1) is highest (48.4%) and CDe/CDE (R_1R_2) & cDE/cDE (R_2R_2) both are lowest (0.4%). In this study, Kell Genotype kk is highest (99.2%) and Kk is lowest (0.8%). Among 6 A-ve blood group donors, all Rh genotypes are cde/cde (rr) (100%). Among 63 A+ve blood group donors, most Rh genotypes are CDe/CDe (R_1R_1) (52.4%) and lowest are cDe/cDe (R_0R_0) (1.6%). Among 17 AB+ve blood group donors, most Rh genotypes are CDe/CDe (R_1R_1) (41.2%) and lowest are CDe/Cde (R_1R_2) (11.8%). Among 10 B-ve blood group donors, most Rh genotypes are cde/cde (rr) (80.0%) and lowest are Cde/cde (r'r) (20.0%). Among 73 B+ve blood group donors, most Rh genotypes are CDe/CDe (R_1R_1) (50.7%) and lowest are cDe/cDe (R_0R_0) (1.4%) & CDe/CDE (R_1R_2) (1.4%). Among 02 O-ve blood group donors, all Rh genotypes are cde/cde (rr) (100%). Among 85 O+ve blood group donors, most Rh genotypes are CDe/CDe (R_1R_1) (55.3%) and lowest are cDe/cDe (R_0R_0) (1.2%) & cDE/cDE (R_2R_2) (1.2%). In this study, only 2 study samples are Kk genotype, 1 is O+ve blood group another is A+ve blood group. (Table-III)

Table-I

Age distribution of the study respondents (n=256)		
Age (years)	Frequency (n)	Percentage (%)
<20	19	7.4
21-30	138	53.9
31-40	80	31.3
41-50	19	7.4
Total	256	100.0
Mean \pm SD	29.2 \pm 6.9	
Range (min - max)	18 - 50	

Table-II

Sex distribution of the study respondents (n=256)		
Sex	Frequency (n)	Percentage (%)
Male	230	89.8
Female	26	10.2
Total	256	100.0
Male : Female	8.8:1	

Table-III		
Frequency of most probable Rhesus genotype in the studied respondents (n=256)		
Most Probable Rhesus genotype	Frequency (n)	Percentage (%)
Cde/cde (r'r)	2	0.8
cDe/cDe (R ₀ R ₀)	6	2.3
CDe/cde (R ₁ r)	68	26.6
CDe/CDe (R ₁ R ₁)	124	48.4
CDe/cDE (R ₁ R ₂)	26	10.2
CDe/CDE (R ₁ R ₂)	1	0.4
cDE/cde (R ₂ r)	12	4.7
cDE/cDE (R ₂ R ₂)	1	0.4
cde/cde (rr)	16	6.3
Total	256	100.0

Most probable Rhesus Genotype CDe/CDe (R₁R₁) is highest (48.4%) and CDe/CDE (R₁R₂) & cDE/cDE (R₂R₂) both are lowest (0.4%). (Table-IV)

Table-V		
Frequency of Kell genotype in the studied respondents (n=256)		
Kell genotype	Frequency (n)	Percentage (%)
kk	254	99.2
Kk	2	0.8
Total	256	100.0

In this study, Kell Genotype kk is highest (99.2%) and Kk is lowest (0.8%). (Table-V)

Among 6 A-ve blood group donors, all Rh genotypes are cde/cde (rr) (100%). Among 63 A+ve blood group donors, most Rh genotypes are CDe/CDe (R₁R₁) (52.4%) and lowest are cDe/cDe (R₀R₀)(1.6%). Among 17 AB+ve blood group donors, most Rh genotypes are CDe/CDe (R₁R₁) (41.2%) and lowest are CDe/cde (R₁R₂) (11.8%). Among 10 B-ve blood group donors, most Rh genotypes are cde/cde (rr) (80.0%) and lowest are Cde/cde (r'r) (20.0%). Among 73 B+ve blood group donors, most Rh genotypes are CDe/CDe (R₁R₁) (50.7%) and lowest are cDe/cDe (R₀R₀)(1.4%) & CDe/CDE (R₁R₂)(1.4%). Among 02 O-ve blood group donors, all Rh genotypes are cde/cde (rr) (100%). Among 85 O+ve blood group donors, most Rh genotypes are CDe/CDe (R₁R₁) (55.3%) and lowest are cDe/cDe (R₀R₀)(1.2%) & cDE/cDE (R₂R₂) (1.2%). (Table-VI)

Table-IV								
Most Probable Rhesus Genotype according to ABO & Rh(D) blood group								
Most Probable Rhesus Genotype	ABO & Rh(D) Blood Group							Total (n=256)
	A-ve (n=6)	A+ve (n=63)	AB +ve (n=17)	B-ve (n=10)	B+ve (n=73)	O-ve (n=2)	O+ve (n=85)	
Cde/cde (r'r)	0(0.0)	0(0.0)	0(0.0)	2(20.0)	0(0.0)	0(0.0)	0(0.0)	2(0.8)
cDe/cDe (R ₀ R ₀)	0(0.0)	1(1.6)	3(17.6)	0(0.0)	1(1.4)	0(0.0)	1(1.2)	6(2.3)
CDe/cde (R ₁ r)	0(0.0)	18(28.6)	5(29.4)	0(0.0)	24(32.9)	0(0.0)	20(23.5)	68(26.6)
CDe/CDe (R ₁ R ₁)	0(0.0)	34(54.0)	7(41.2)	0(0.0)	37(50.7)	0(0.0)	47(55.3)	124(48.4)
CDe/cDE (R ₁ R ₂)	0(0.0)	7(11.1)	2(11.8)	0(0.0)	5(6.8)	0(0.0)	12(14.1)	26(10.2)
CDe/CDE (R ₁ R ₂)	0(0.0)	0(0.0)	0(0.0)	0(0.0)	1(1.4)	0(0.0)	0(0.0)	1(0.4)
cDE/cde (R ₂ r)	0(0.0)	3(4.8)	0(0.0)	0(0.0)	5(6.8)	0(0.0)	4(4.7)	12(4.7)
cDE/cDE (R ₂ R ₂)	0(0.0)	0(0.0)	0(0.0)	0(0.0)	0(0.0)	0(0.0)	1(1.2)	1(0.4)
cde/cde (rr)	6(100.0)	0(0.0)	0(0.0)	8(80.0)	0(0.0)	2(100.0)	0(0.0)	16(6.3)
Total	6(100)	63(100)	17(100)	10(100)	73(100)	2(100)	85(100)	256(100)

Table-VI

Kell	ABO&Rh(D) Blood Group							Total (n=256)
	A-ve (n=6)	A+ve (n=63)	AB+ve (n=17)	B-ve (n=10)	B+ve (n=73)	O-ve (n=2)	O+ve (n=85)	
Genotype								
k k	6(100.0)	62(98.4)	17(100.0)	10(100.0)	73(100.0)	2(100.0)	84(98.8)	254(99.2)
K k	0(0.0)	1(1.6)	0(0.0)	0(0.0)	0(0.0)	0(0.0)	1(1.2)	2(0.8)
Total	6(100.0)	63(100.0)	17(100.0)	10(100.0)	73(100.0)	2(100.0)	85(100.0)	256(100.0)

In this study, only 2 study samples are Kk genotype, 1 is O+ve blood group another is A+ve blood group

Discussion

Among the 256 donors, 21-30 years age group donor is more (53.9%) and 41-50 years age group donor is less (7.4%). Male donor is predominant (89.8%) and Female donor is (10.2%). In case of ABO & Rh(D) blood groups donors, O Rh(D)+ve donor is more (33.2%) & O Rh(D)-ve donor is lowest (0.8%). In this study, Rhesus Phenotype CCDee is highest (48.4%) & CCDEe & ccDEE both are lowest (0.4%). Most probable Rhesus Genotype CDe/CDe (R_1R_1) is highest (48.4%) and CDe/CDE (R_1R_2) & cDE/cDE (R_2R_2) both are lowest (0.4%). Similar results were found in the study of Sangeeta P, Sonal J et al where CCDee frequency was 42.2% and probable genotype was R_1R_1 (CDe/CDe).¹³ Some other studies also agreed with this study.^{14,15,16}

In this study, Kell Genotype kk is highest (99.2%) and Kk is lowest (0.8%). A study in North Indian blood donors and some other studies agreed with this study.^{13,17,18,19}

Among 6 A-ve blood group donors, all Rh genotypes are cde/cde (rr) (100%). Among 63 A+ve blood group donors, most Rh genotypes are CDe/CDe (R_1R_1) (52.4%) and lowest are cDe/cDe (R_0R_0) (1.6%). Among 17 AB+ve blood group donors, most Rh genotypes are CDe/CDe (R_1R_1) (41.2%) and lowest are CDe/Cde (R_1R_2) (11.8%). Among 10 B-ve blood group donors, most Rh genotypes are cde/cde (rr) (80.0%) and lowest are Cde/cde (r'r) (20.0%). Among 73 B+ve blood group donors, most Rh genotypes are CDe/CDe (R_1R_1) (50.7%) and lowest are cDe/cDe (R_0R_0) (1.4%) & CDe/CDE (R_1R_2) (1.4%). Among 02 O-ve blood group donors, all Rh genotypes are cde/cde (rr) (100%). Among 85 O+ve blood group donors, most Rh genotypes are CDe/CDe (R_1R_1) (55.3%) and lowest are cDe/cDe (R_0R_0) (1.2%) & cDE/cDE (R_2R_2) (1.2%). In this study, only 2 study samples are Kk genotype, 1 is O+ve blood group another is A+ve blood group. These findings were in accordance with the findings of previous studies by Hafid Z, Anass Y, Jean U et al.²⁰

Conclusion

In case of Rhesus blood group system, most probable Rhesus Genotype CDe/CDe (R_1R_1) is highest (48.4%). These population are vulnerable to develop alloantibody against other red cell antigen. These alloantibody are clinically significant and can cause hemolytic disease of fetus and newborn & hemolytic transfusion reaction. In case of Kell blood group system, Kell Genotype kk (Cellano) is highest (99.2%) and Kk (Kell) is lowest (0.8%). Kell antigen is very strong but frequency is low. In this case, alloimmunization causes severe hemolytic disease of fetus and newborn & hemolytic transfusion reaction. It is very difficult to identify the cause. This study will help to make a donor pool and supply blood according to Rh phenotype & Kell phenotype.

Ethical issues

This research was approved by the Institutional Review Board (IRB) of BSMMU. The participants were informed about the risks, benefits and possible outcomes of this work and the information were given in an easy and local language (Bengali). Informed written consent was taken from each donor.

Conflict of Interest

Authors declare no conflict of interest.

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