Combined Group I and III ABO Discrepancies in Multiple Myeloma with IgG-Lambda Type: A Case Report

Joonhong Park^a Dong Wook Jekarl^a Suk Young Park^b Soyoung Shin^a

Departments of ^aLaboratory Medicine and ^bInternal Medicine, College of Medicine, Catholic University of Korea, Seoul, Republic of Korea

Key Words

Multiple myeloma · ABO discrepancies · Rouleau formation · Loss of isoagglutinin

Abstract

Objective: To report a case with unusual ABO discrepancies caused by coexistence of the loss of anti-B isoagglutinin and rouleau formation. **Clinical Presentation and Intervention:** A 79-year-old female diagnosed as having multiple myeloma (MM) with monoclonal IgG- λ type showed rouleau formation in peripheral blood smear. The ABO and Rh blood type before the diagnosis of MM was A+, but the following ABO grouping was interpreted as AB+. The ABO genotype revealed the subtypes A102 and O101, which confirmed her ABO phenotype as A+. **Conclusion:** This was a case of combined group I and III ABO discrepancies mimicking blood group AB.

© 2016 S. Karger AG, Basel

modern laboratory technologies and reagents developed, ABO discrepancies still occur, which means the unexpected pattern of antigen on the cell and the opposite antibody in the serum. ABO discrepancies may be arbitrarily divided into 4 major categories: briefly, group I is associated with unexpected reactions in the reverse typing due to weakly reacting or missing antibodies, whereas group II is associated with unexpected reactions in the forward typing due to weakly reacting or missing antigens. Group III is caused by protein or plasma abnormalities and results from rouleau formation or pseudoagglutination. Finally, group IV is due to miscellaneous problems such as polyagglutination [1, 2]. The purpose of this report was to describe a case with unusual ABO discrepancies caused by coexistence of the loss of anti-B isoagglutinin and the rouleau formation in a patient diagnosed as multiple myeloma (MM) with monoclonal gammopathy of the IgG- λ type.

Introduction

The ABO blood group is the most important blood type system in human blood transfusion, which exists as 4 allelic forms of AB, A, B, and O. Notwithstanding the

Case Report

A 79-year-old female with type 2 diabetes mellitus was admitted to Daejeon St. Mary's Hospital for the evaluation of right-sided chest pain with no associated injuries. Contrast-enhanced computed tomography revealed multiple fractures at the right 4th to 10th ribs. In the peripheral blood smear, rouleau formation was promi-

Table 1. Laboratory findings of our patient and reported cases showing ABO discrepancy in literature review

Sex/ age	Diagnosis	ABO discrepancy (forward/reverse)	Discrepancy cause	Discrepancy category	ABO genotype	IgG, mg/dl	IgA, mg/dl	IgM, mg/dl	Ref.
F/79	MM with IgG (λ)	A/A → AB/AB	Loss of anti-B and rouleau formation	Groups I and III	ABO*A/O	6,468	90	20.4	This study
F/56	MM with IgG (κ)	A/AB	Loss of anti-B	Group I	n.d.	4,970	<29.9	<26.4	3
F/60	MM with λ	B/AB	Loss of anti-A	Group I	n.d.	382.8	<28.5	<29	3
M/52	MM with IgG (λ)	B/AB	Loss of anti-A	Group I	n.d.	5,243	<28.5	<29	3
F/78	MM with IgA (κ)	B/AB	Loss of anti-A	Group I	ABO*B/O	236	4,430	<20	4
F/13	Idiopathic agammaglobulinemia	B/AB	Loss of anti-A	Group I	n.d.	<33.3	<6.7	<4.2	5
F/30	Gallbladder abscess	O/B	Loss of anti-B	Group I	ABO*O/O	39	46	<5	6
F/55	Liver cancer	O/B	Loss of anti-B	Group I	ABO*O/O	63	65	12	6
M/20	Common variable immunodeficiency	O/B	Loss of anti-B	Group I	ABO*O/O	104	<13	<15	7
M/16	Acute osteomyelitis	O/B	Loss of anti-B	Group I	ABO*O/O	770	244	13.5	8

MM = Multiple myeloma; n.d. = not done.

nent. Laboratory findings showed white blood cell count of 8,500/mm³, hemoglobin of 6.7 g/dl, platelet count of 142,000/mm³, IgG of 6,468 mg/dl, IgA of 90 mg/dl, IgM of 20.4 mg/dl, κ light chain of 48.26 mg/l (serum, reference 3.3–19.4) and 138.36 mg/l (urine, 0–32.71), λ light chain of 1,259 mg/l (serum, 5.71–26.30) and 1,195.40 mg/l (urine, 0–4.99), blood urea nitrogen of 101.4 mg/dl, and creatinine of 3.62 mg/dl. Serum and urine immunofixation showed monoclonal gammopathy of the IgG-λ type reflecting an abnormal albumin-to-globulin ratio. Patient red blood cells (RBCs) were evaluated for reactivity of the forward typing with Novaclone anti-A and anti-B murine monoclonal reagents (Dominion Biologicals Ltd., Dartmouth, N.S., Canada) and of the reverse typing with Affirmagen A1 and B RBC cell reagents (Ortho Clinical Diagnostics, Pencoed, UK). The ABO and Rh blood type before the diagnosis of MM had been A+ in previous medical records, but the following ABO grouping showed A+ in red cells but not anti-B in serum by a conventional tube test. A repeated ABO test by the microtube column agglutination technique using the Diamed-ID LISS/Coombs gel card containing anti-IgG and anti-C3d within the gel matrix (DiaMed GmbH, Cressier, Switzerland) was interpreted as AB+ (the forward type: A cell 4+, B cell 2+; the reverse type: absence of anti-A and anti-B). They were negative for both Coombs test and irregular antibody screening. To resolve this ABO discrepancy, the regions containing coding sequences of exons 6 and 7 in the ABO gene were directly sequenced on an ABI 3130XL Genetic Analyzer (Applied Biosystems, Foster City, Calif., USA). The ABO genotype contained the subtypes *A102* and *O101*; thus, her ABO phenotype was confirmed to be A+. She received several units of A+ packed RBC transfusion without complication, undergoing the conservative treatment due to her old age.

Discussion

This was a case of combined group I and III ABO discrepancy caused by coexistence of the loss of anti-B isoagglutinin and the rouleau formation in MM with monoclonal gammopathy of the IgG-λ type. Several Korean

cases had been reported with MM [3, 4] or with other diseases [5–8] about group I ABO discrepancies due to the loss of isoagglutinins only (table 1). An ABO incompatibility reaction such as acute hemolytic reaction or fatal complication can occur in our patient if she receives the wrong type of AB+ RBC transfusion instead of true type A+.

M protein produced by MM cells is a common cause of group III ABO discrepancies resulting from elevated levels of globulin. Abnormally elevated paraproteins cause the rouleau formation which is stacks or aggregations of RBCs due to reduction in 'zeta potential' (charge on RBC membrane) [2]. It could be mistaken for agglutination by medical technicians in the forward grouping. The washing technique rids the RBC membranes of the paraproteins and frees the RBCs in the case of rouleau formation in the reverse grouping. In true agglutination, red cells will continue to clump after washing with normal saline [3].

Meanwhile, group I ABO discrepancies can also occur when patients have depressed antibody production or cannot produce the ABO antibodies. Weak agglutination reactions may be obtained with reagent RBC cells and are a result of weak expression of anti-A and anti-B in the serum. Group I ABO discrepancy should be suspected because RBC and serum grouping reactions are normally very strong. IgM is the predominant isotype found in blood group A or B individuals, although small quantities of IgG antibody can be detected. In blood group O serum IgG is the major isotype for anti-A and anti-B [2]. Therefore, the severe deficiency of IgM would be responsible for group I ABO discrepancies in MM with blood group A or B [2].

Recently, Kaur et al. [9] reported that a few (0.06%, 28/44,425) blood group discrepancies were observed in 44,425 blood groups. Among them, 20 subgroups of A and B were a main sample-related problem of ABO discrepancies. In order to resolve the ABO discrepancy and to provide compatible blood for transfusion, it is necessary to obtain relevant historical information from the patient and to provide medical and technical expertise as well as education for the blood bank technologist in transfusion medicine-related testing, practice, risks, and related areas [2]. ABO genotyping can also be done with a high level of accuracy for the resolution of ABO serological discrepancies [10].

Conclusion

This was a case of combined group I and III ABO discrepancies mimicking blood group AB in MM with monoclonal IgG- λ type. ABO genotyping was very useful to resolve this unusual ABO discrepancy.

Disclosure Statement

There is no conflict of interest.

References

- 1 Harmening DM: The ABO blood group system; in Harmening DM (ed): *Modern Blood Banking and Transfusion Practices*, ed 6. Philadelphia, Davis Co, 2012, pp 136–148.
- 2 Roback JD, Combs MR, Grossman BJ, et al (eds): *Technical Manual*, ed 17. Bethesda, American Association of Blood Banks (AABB), 2011. <http://www.aabb.org/>.
- 3 Ha JS, Kim EJ, Chun HJ, et al: Three cases of multiple myeloma showing ABO discrepancy. *Korean J Blood Transfus* 1998;9:289–293.
- 4 Kim SY, Oh SH, Park KS, et al: ABO discrepancy in an elderly patient with IgA kappa-type multiple myeloma. *Ann Hematol* 2010;89:747–748.
- 5 Park TS, Oh SH, Choi JC, et al: A case of agammaglobulinemia detected by ABO discrepancy in a 13-year-old girl. *Korean J Lab Med* 2002;22:364–366.
- 6 Suh IB, Chang EA, Kim HJ, et al: Two cases of ABO discrepancy due to hypogammaglobulinemia. *Korean J Blood Transfus* 2003;14:240–245.
- 7 Oh SH, Kang CI, Kim J, et al: ABO discrepancy in a young Korean serviceman with common variable immunodeficiency. *Ann Hematol* 2010;89:629–630.
- 8 Jung CL, Cha MK, Jun BH, et al: A case of IgM deficiency with B cell deficiency detected by ABO discrepancy in a patient with acute osteomyelitis. *Ann Lab Med* 2013;33:208–211.
- 9 Kaur G, Kaur P, Basu S, et al: Blood group discrepancies at a tertiary care centre – analysis and resolution. *Int J Lab Hematol* 2014;36:481–487.
- 10 Anstee DJ: Red cell genotyping and the future of pretransfusion testing. *Blood* 2009;114:248–256.