

## Thawing Time Difference between Fresh Frozen Plasma Using Ziplock Plastic and non-Ziplock Plastic in Blood Transfusion Unit Dr. Hasan Sadikin General Hospital Bandung, Indonesia

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### Abstract

**Background:** The quality of fresh frozen plasma (FFP) in a clinical setting depends on thawing time. Thawing using a water bath is often used in blood transfusion units because it is easy to perform, affordable, and easy to look for. Protective plastics (Ziplock and non-Ziplock) are used to reduce the risk of contamination. This study aimed to determine whether there is a difference in thawing time between FFP using Ziplock plastic and non-Ziplock plastic.

**Methods:** This experimental design was conducted in the Blood Bank Unit of Dr. Hasan Sadikin General Hospital Bandung, Indonesia from June–August 2021. Samples were divided into two groups, namely thawing using Ziplock and non-Ziplock. Each group consisted of volume 160–200 mL and 201–240 mL.

**Results:** The total samples were 24 FFP bags. In the group of the bag 160–200 mL, the median thawing time using Ziplock plastic was 8 minutes (8–16 minutes), non-Ziplock was 15 minutes (8–16 minutes) ( $p=0.111$ ), whereas in a group of bags with volume 201–240 mL, the median thawing time using Ziplock was 15 minutes (8–28 minutes), non-Ziplock was 20 minutes (14–30 minutes) ( $p=0.332$ ). Although there was a time difference in both groups, the difference was non-significant.

**Conclusions:** The thawing time between the small bag with a volume of 160–200 mL and the larger volume of 201–240 mL shows no difference. Ziplock plastic can be used to reduce the risk of contamination.

**Keywords:** FFP, non-Ziplock plastic, thawing time, water bath, Ziplock plastic

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### Introduction

Fresh frozen plasma (FFP) is prepared from single units of whole blood or plasma collected from apheresis techniques. It is frozen at  $-18$  to  $-30$  °C within eight hours of collection and, when appropriately stored, is usable for one year from the collection date.<sup>1,2</sup> Fresh frozen plasma contains fibrinogen (400 to 900 mg/unit), albumin, protein C, protein S, antithrombin, and tissue factor pathway inhibitors.<sup>2,3</sup> The FFP transfusion is used to replace some coagulation factors and is indicated in patients who experience bleeding with a history of routine use of warfarin who will undergo invasive procedures, thrombotic

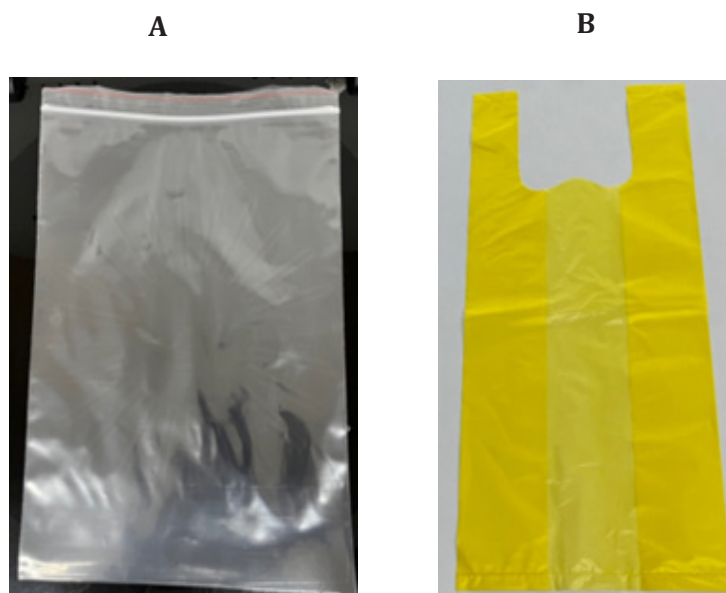
thrombocytopenia purpura (TTP), hemolytic uremic syndrome (HUS), deficiency of coagulation factors, massive transfusion in hypocoagulable and hypovolemic patients. After FFP transfusion, coagulation factors will increase by as much as 20%.<sup>1,4,5</sup>

Bleeding cases requiring FFP transfusion, the administration of FFP depends on the rate at which FFP is thawed.<sup>6,7</sup> Before a frozen bag of plasma can be used for transfusion, it should be warmed to a desired transfusable temperature.<sup>8</sup> Several tools can be used to defrost FFP, namely a dry oven (incubator with temperature control) which requires a fast defrosting time of 10 minutes and is able to reduce the risk of contamination by bacteria.<sup>9</sup>

The second tool is a microwave oven; the advantage of this tool is that it has the fastest defrosting ability compared to other methods, which is  $\pm 2-3$  minutes; the disadvantages are that it is expensive and has limited capacity.<sup>9</sup> The third tool is a water bath, the advantage of this tool is that it is easy to use by placing the FFP into a water bath whose temperature has been set between 30–37 °C for 20–30 minutes and the price is relatively affordable and easy to obtain, the disadvantage is that there is a risk of contamination when defrosting it without protective plastic.<sup>10</sup> Among the three FFP liquefaction tools, the water bath is the most frequently used.<sup>11,12</sup> A good quality FFP bag must meet several requirements such as being resistant to cold and hot temperatures, mechanically resistant, not releasing harmful substances into blood components, and having no color (transparent). Materials that meet these requirements are polyvinyl chloride (PVC).<sup>2,12</sup> The use of plastic bags is highly recommended for thawing FFP in water bath to avoid contamination by bacteria. Defrosting FFP in the water bath without using protective plastic will increase the risk of septicemia as reported in a study conducted in Germany,<sup>10</sup> which found cases of septicemia caused by *Pseudomonas aeruginosa* due to thawing of FFP in water bath that were not coated with plastic. Further research shows that a liquid volume of 0.025 mL on thawing without using plastic will increase the infection rate.<sup>10</sup>

Therefore, the plastic bags used in the FFP liquefaction process must be resistant to chemicals, and water vapor, have a high melting point (115–135 °C), be easy to obtain, and be inexpensive.<sup>12,13</sup> The best plastic material for this purpose is polyethylene, which consists of high-density polyethylene (HDPE) and low-density polyethylene (LDPE).<sup>14</sup>

Factors that can affect the thawing time of FFP are volume, number of bags of FFP being defrosted simultaneously, thawing temperature, and the type of protective plastic. According to research in London, the larger the volume of disbursed FFP, the longer it will take to melt.<sup>15,16</sup> Other studies stated that the melting time of FFP without the use of plastic at 37 °C takes 17 minutes, and the melting process using plastic at the same temperature takes 28 minutes. Although faster, thawing of FFP without the use of plastic can pose a risk of septicemia.<sup>17-19</sup> Dry ovens and microwave ovens are rarely available and more expensive than water bath. Therefore, the use of a water bath can help hospitals or blood care units to choose which method has the fastest thawing time without risking contamination. This study aimed to analyze the difference in the thawing time of FFP using Ziplock and non-Ziplock plastics. Ziplock plastic is a resealable sliding storage, container plastics, and non-Ziplock plastic in this study is a yellow plastic used to store or dispose medical items. These plastics can be seen in Figure 1.



**Figure 1 Protective Plastics**

Note: A= Ziplock plastic, B= Non-Ziplock plastic

**Table 1 Data Characteristics of FFP Samples**

Sample Characteristics (Blood Type)	Ziplock n	Non-Ziplock n
Blood type		
A+	3	2
B+	1	3
AB+	0	1
O+	8	6

### Methods

This study used an experimental method with a sample size of 24 FFP bags determined based on the Federer formula.<sup>20</sup> The study was conducted at the Blood Services Unit of Dr. Hasan Sadikin General Hospital Bandung, Indonesia from June to August 2021.

This study used an FFP bag that was used commercially and met the predetermined volume groups, namely 160–200 mL and 201–240 mL. Apart from volume, the FFP groups were divided into two groups of defrosting, using Ziplock and non-Ziplock plastic bags. Before thawing, water bath was warmed until 37 °C for 10 minutes in the Ziplock and non-Ziplock groups. The chi-square test was carried out to determine the characteristics of the data, the normality test of the data using Shapiro-Wilk followed by a different test using Mann Whitney for the analysis of time differences with p-value <0.05 was considered significant. This study protocol was approved by the Ethical Committee of Dr. Hasan Sadikin General Hospital Bandung, Indonesia, with number LB.02.01/X.6.5/253/2021.

### Results

During the study, 24 samples were obtained, with a description of the characteristics listed in Table 1. This study found that the highest demand for FFP was in the O+ blood group, both from the Ziplock and non-Ziplock groups, and the least demand was in the AB+ blood group, both from the Ziplock and non-Ziplock groups (Table 1).

Volume in the Ziplock group was less than in the non-Ziplock group, but the difference was not significant (Table 2). The data normality test was carried out using Shapiro Wilk and the data was not normally distributed, followed by the Mann-Whitney test to analyse the difference in thawing time between FFP using Ziplock and non-Ziplock plastic (Table 3). From the analysis results, there was a non-significant time difference between thawing using Ziplock and non-Ziplock plastics (Table 3).

### Discussion

From the data characteristics in Table 1, it is found that the most blood type is O+ and

**Table 2 Comparison of Volumes Using Zip Lock and non-Zip Lock Plastics**

	Ziplock Plastic (n=12)	Non-Ziplock Plastic (n=12)	p-value
	Median (Min.–Max.)	Median (Min.–Max.)	
Volume (mL)	201(180–220)	208 (183–239)	0.524

**Table 3 Comparison of FFP Thawing Time Using Ziplock Plastic and non-Ziplock Plastic**

FFP Bag Volume	Thawing Time (Minute)		Median Difference (95% CI)	p-value
	Ziplock Plastic (n=12)	Non-Ziplock Plastic (n=12)		
	Median (Min.–Max.)	Median (Min.–Max.)		
160–200 mL	8 (8–16)	15 (8–16)	-7 (-8–0)	0.11
201–240 mL	15 (8–28)	20 (14–30)	-5 (-15–8)	0.33

the least is AB+. This is in accordance with research in India,<sup>11</sup> that out of a total of 10,000 donors, blood type O+ is the most common (37.1%) followed by B+ (32.3%), A+ (22.8%), and AB+ (7.7%). The distribution of this blood group corresponds to the percentage in the world, which is the most O+ blood group.<sup>21,22</sup>

The result of this study showed that there was no significant difference in thawing time using Ziplock and non-Ziplock plastic. The difference in thawing time is not significant because thawing time is mainly influenced by the volume of FFP, not the type of plastic.<sup>23</sup> In our study, the more the volume, the longer the thawing time. The theoretical basis for this statement is that the more volume, the greater the heat transfer required for thawing. This is in accordance with another research which carried out the thawing process using a microwave assisted thawing apparatus in a food system model with a thick and homogeneous consistency. There was no significant change in volume during the freezing and thawing process. The model was divided into two treatment groups, namely samples with a thickness of 3.6 cm and 5.5 cm. From this study, it was found that volume was the factor that most significantly affecting the thawing time, samples with a thickness of 5.5 cm took three times longer than samples with a thickness of 3.6 cm. From these research, it was found that volume is the factor that most influences thawing time.<sup>22,23</sup>

Ziplock plastic base material is LDPE with a thickness of  $\pm 0.5$  mm, thinner than non-Ziplock plastic with a thicker HDPE base material (10–30 mm). In addition, the Ziplock plastic has a tight plastic tip, so that no air comes in and out during defrosting process, and the temperature during defrosting is stable. In contrast to non-Ziplock plastic, which does not have a tight end, causing air to enter and exit during defrosting so that the temperature during defrosting is unstable, this also causes non-Ziplock plastic not to be completely submerged during defrosting, in contrast to Ziplock plastic which is completely submerged.<sup>22,23</sup>

Although not significantly different, defrosting using Ziplock plastic can be an option in daily practice since the melting temperature is stable and can reduce the risk of contamination because LDPE plastic is resistant to high temperatures up to 90 °C.

There are several limitations in this study, namely that temperature measurements were not performed at the end of defrosting process; no hemostasis tests were carried out after the

defrosting process; no culture examination was performed after defrosting to determine whether contamination occurred.

In conclusion, there is no significant difference in the duration of FFP thawing using Ziplock and non-Ziplock plastic. However, Ziplock plastic can be an option to reduce the risk of contamination.

## References

1. Uhl L. Clinical use of plasma components. UpToDate [Internet]. [cited 2022 June 10]. Available from: <https://www.uptodate.com/contents/clinical-use-of-plasma-components>.
2. O'Shaughnessy DF, Atterbury C, Maggs PB, Murphy M, Thomas D, Yates S, et al. Guidelines for the use of fresh-frozen plasma, cryoprecipitate and cryosupernatant. *Br J Haematol*. 2004;126(1):11–28.
3. Khawar H, Kelley W, Stevens JB, Guzman N. Fresh frozen plasma (FFP). Treasure Island (FL): StatPearls Publishing; 2021. [cited 2021 April 11]. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK513347/>.
4. Veera RRL, Schneider D, Pickens PV. Fresh frozen plasma (FFP) usage and appropriateness in adult medical inpatients: a retrospective audit. *Blood*. 2012;120(21):4375.
5. Cardigan R, Green L. Thawed and liquid plasma—what do we know? *Vox Sang*. 2015;109(1):1–10.
6. Meledeo MA, Peltier GC, McIntosh CS, Corley JB, Bynum JA, Cap AP. Field-expedient thawing of fresh-frozen plasma. *Transfusion*. 2020;60(Suppl 3):S87–95.
7. Churchill WH, Schmidt B, Lindsey J, Greenberg M, Boudrow S, Brugnara C. Thawing fresh frozen plasma in a microwave oven: a comparison with thawing in a 37 °C waterbath. *Am J Clin Path*. 1992;97(2):227–32.
8. Khandpur RS. Plasma thawing equipment. In: Khandpur RS, editor. *Compendium of biomedical instrumentation*. 1<sup>st</sup> Ed. Hoboken: John Wiley & Son; 2019. p.1537–9.
9. Retno D, Zainal AR, Machsoos BD, Hermanto DH, Wardhani SO. Perbedaan kualitas fresh frozen plasma yang dicairkan dengan metode konvensional dan dengan metode alat FFP thawer. *J Peny Dalam*. 2012;13(1):21–7.
10. Pinki S, Mohan G, Rafi A, Innah S, Thomas

- T. Rapid dry plasma thawing system: An alternative to conventional thawing baths. *Asian J Transfus Sci.* 2017;11(2):147-50.
11. Heger A, Pock K, Römisch J. Thawing of pooled, solvent/detergent-treated plasma octaplasLG®: validation studies using different thawing devices. *Transfus Med Hemother.* 2017;44(2):94-8.
  12. Sojiphan K, Iverson A, Kim S. Materials selection analysis: bag for viable blood storage [Internet]. 2004. [cited 2021 April 24]. Available from: [https://www.researchgate.net/publication/291766516\\_Materials\\_Selection\\_Analysis\\_Bag\\_for\\_Viable\\_Blood\\_Storage](https://www.researchgate.net/publication/291766516_Materials_Selection_Analysis_Bag_for_Viable_Blood_Storage).
  13. Zahra NM, Siswanto S, Widiyanti P. The role of chitosan on polyvinyl chloride (PVC)-glycerol biocomposites for blood bag application. *JBBBE.* 2018;37:94-106.
  14. Platton S, Elegbe O, Bower L, Cardigan R, Lancut J, McCullagh J, et al. Thawing times and hemostatic assessment of fresh frozen plasma thawed at 37°C and 45°C using water-bath methods. *Transfusion.* 2019;59(11):3478-84.
  15. Chavan SK. Determination of rate and analysis of reasons for discarding blood and blood components in a blood bank of tertiary care hospital: a retrospective study. *Int J Res Med Sci.* 2017;5(3):1111-5.
  16. Green L, Bolton-Maggs P, Beattie C, Cardigan R, Kallis Y, Stanworth S, et al. British Society of Haematology Guidelines on the spectrum of fresh frozen plasma and cryoprecipitate products: their handling and use in various patient groups in the absence of major bleeding. *Br J Haematol.* 2018;181(1):54-67.
  17. Plotz RD, Ciotola RT. Thawing of fresh-frozen plasma at 45 °C versus 37 °C: comparison using satellite packs of the same donor units. *Am J Clin Pathol.* 1988;89(3):381-4.
  18. Dhantole L, Dubey A, Sonker A. A study on factors influencing the hemostatic potential of fresh frozen plasma. *Asian J Transfus Sci.* 2019;13(1):23-9.
  19. Castillo B, Dasgupta A, Klein K, Tint H, Wahed A. *Transfusion medicine for pathologists: a comprehensive review for board preparation, certification, and clinical practice.* 1<sup>st</sup> Ed. Amsterdam: Elsevier; 2018.
  20. Ihwah A, Deoranto P, Wijana S, Dewi IA. Comparative study between Federer and Gomez method for number of replication in complete randomized design using simulation: study of Areca Palm (*Areca catechu*) as organic waste for producing handicraft paper. *IOP Conf Ser Earth Environ Sci.* 2018;131(1):012049.
  21. Agrawal A, Tiwari AK, Mehta N, Bhattacharya P, Wankhede R, Tulsiani S, et al. ABO and Rh (D) group distribution and gene frequency; the first multicentric study in India. *Asian J Transfus Sci.* 2014;8(2):121-5.
  22. Simpson V. World population by percentage of blood types-WorldAtlas [Internet]. 2020. [Cited 2022 May 1]. Available from: <https://www.worldatlas.com/articles/what-are-the-different-blood-types.html>.
  23. Virtanen AJ, Goedeken DL, Tong CH. Microwave assisted thawing of model frozen foods using feed-back temperature control and surface cooling. *J Food Sci.* 2006;62(1):150-4.