## **EFFECTIVENESS OF PLATELET TRANSFUSION IN DENGUE FEVER PATIENTS:** A SYSTEMATIC REVIEW

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### **ABSTRACT**

Dengue fever is a global health concern characterized by severe thrombocytopenia and an increased risk of bleeding. Platelet transfusion is often used to manage thrombocytopenia, but its effectiveness remains uncertain. This review aims to evaluate the efficacy of platelet transfusion in preventing bleeding and improving clinical outcomes among dengue fever patients. A systematic search was conducted across PubMed, ScienceDirect, and Google Scholar using keywords such as "platelet transfusion," "dengue fever," and "thrombocytopenia." Nine studies, including randomized controlled trials and observational studies, were analyzed for outcomes related to bleeding incidence, platelet recovery, and hospitalization duration. The findings indicate that prophylactic platelet transfusion does not significantly reduce bleeding incidence or improve recovery rates in dengue patients. It may be associated with delayed platelet recovery, prolonged hospitalization, and increased risk of adverse events.

**Keywords:** Dengue Fever, Platelet Transfusion, Thrombocytopenia, Bleeding, Hospitalization.

### INTRODUCTION

Millions of people worldwide suffer from dengue fever each year, which is brought on by the dengue virus spread by Aedes mosquitoes. Thrombocytopenia, a sign of a severe dengue infection, raises questions about potential bleeding complications. Although platelet transfusion has been routinely used to reduce the risk of bleeding, its effectiveness and safety are still up for debate. With an emphasis on overall clinical outcomes bleeding prevention, this review attempts to methodically assess the data regarding the efficacy of platelet transfusion in dengue fever patients.

Platelets play an important role in maintaining hemostasis by adhering to blood vessel sites and forming platelet plaques.1 Healthy individuals have blood platelet levels of 150x109 to 400x109 platelets per liter. Thrombocytopenia can cause bleeding symptoms ranging from petechiae to intracranial hemorrhage, pulmonary hemorrhage, and death (Andriastuti, 2024).

Refractory thrombocytopenia can be caused by immune and nonimmune factors. Non-immune causes are more likely to cause poor posttransfusion responses. Non-immune causes should be evaluated and addressed. The most common immune cause is the presence of antibodies to human leukocyte (HLA) and/or antigens human platelet antigens (HPA). These antibodies can be formed from exposure to transfusion, pregnancy, or transplantation. Immune causes refractory thrombocytopenia should be evaluated by identifying the presence of anti-HLA and anti-HPA antibodies. If these antibodies are identified, there are several strategies to identify compatible platelet units. One strategy is crossmatching. Donor platelets from crossmatching results can be given to patients who have anti-HLA and/or anti-HPA antibodies (Manoharan, 2016).

### LITERATURE REVIEW

Platelet transfusion in dengue hemorrhagic fever (DHF) is done to increase platelet levels in the blood prevent bleeding. and Platelet transfusion is usually given if the platelet levels patient's drop drastically to below the critical limit or if the patient experiences bleeding (Asmarani, 2021).The decision to give platelet transfusion

to DHF patients should be based on several factors below.

- 1. Platelet count. The platelet count drops quite significantly and is very low, which is around 20,000 pieces per microliter of blood (mcL).
- 2. Clinical signs of bleeding. DHF patients experience active bleeding, including bruising on bleeding the skin. gums, nosebleeds. or internal bleeding.
- 3. Other factors. The age and overall health condition of the patient can influence the decision to transfuse platelets (Kaur, 2014).

## **RESEARCH METHODS**

Google Scholar, ScienceDirect, and PubMed were all thoroughly searched using keywords such as "platelet transfusion," fever," and "thrombocytopenia." Nine studies, including randomized controlled trials and observational studies, were analyzed for outcomes related to bleeding incidence, platelet recovery, and hospitalization duration. **PICO** (Population = dengue fever patient, Intervention = transfusion platelet, Comparison = not given platelet transfusion, and Outcome = improve recovery rates) was the basis for the inclusion criteria. (Barker et al., 2023). Table 1 contains an overview of the research or data extraction.

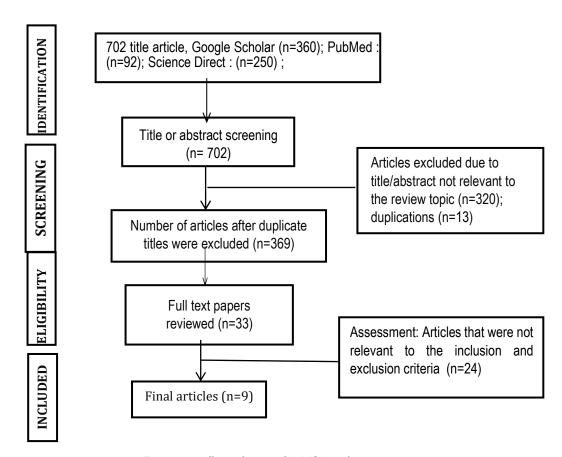


Figure 1. flowchart of PRISMA diagram

# Search Strategy

A comprehensive literature search was conducted using PubMed, ScienceDirect, and Google Scholar with keywords "platelet transfusion," "dengue fever," and "thrombocytopenia." The search spanned articles published up to 2024. The inclusion criteria were:

- 1. Articles published in English.
- 2. Studies focusing on dengue fever patients receiving platelet transfusion.
- Study designs including randomized controlled trials, cohort studies, or observational studies.
- 4. Reporting of relevant clinical outcomes, such as bleeding incidence, platelet recovery, and hospitalization duration.

# Study Selection and Data Extraction

From 702 identified studies, 369 were screened after duplicate removal. Following title and abstract screening, 33 full-text articles were assessed for eligibility, with nine meeting the inclusion criteria. Data extraction focused on study design, population characteristics, interventions, and reported outcomes.

## PRISMA Flow Diagram

- 1. Records identified through database searching: 702.
- 2. Records after duplicates removed: 369.
- 3. Full-text articles assessed for eligibility: 33.
- 4. Studies included in qualitative synthesis: 9.

Tabel 1

Title	Author	N	Design	Intervent	Tool	Outcom
articles	s and year of publica tion	14	Design	ion	1001	e
Platelet Transfusion and Tranexamic Acid in the Treatment of Bleeding in Dengue Fever	(Saddiq ue et al., 2022) Pakista n	10 0	Retrospecti ve review of patient data. Groups: Patients classified into four treatment groups (details on treatments classified but not specified in the excerpt).	• Interve ntions Studied : Platelet transfus ion and/or intraven ous tranexa mic acid	Data Collection: Review of clinical records from Jinnah Hospital, Lahore, Pakistan	Results; There was no significa nt differen ce in the median time betwee n groups from the start of treatme nt to the end of bleedin g (p value = 0.724). Among the side effects reporte d were pruritus and stomach ache. •The study found that in individu als with clinical bleedin g from dengue fever, platelet transfus

					ion and/or tranexa mic acid do not significa ntly improve standar d care treatme nt and may even have negativ e side effects.
Predictors and Clinical Outcomes of Poor Platelet Recovery in Dengue	(Archul eta et al., 2020) Singap ore	The study utilized a multicente r, prospective design	Monitorin g platelet recovery in dengue patients was part of the strategy, with an emphasis on the conseque nces of platelet transfusio n.	The study employed clinical assessment s and laboratory tests to evaluate platelet counts and other baseline characteris tics.	Platelet recover y and clinical outcom es associat ed with dengue severity, such as hospital ization duration and the occurre nce of severe dengue, were the main outcom es assessed . The study found that poor platelet recover y in

Effect of

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on Clot

Dengue

Transfusion

Strength in

Fever with

(Sunda

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Bhaska

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						experie nced bleedin g problem s, they had no effect on the strength of the clot. Even after getting platelet transfus ions, most patients still experie nced bleedin g.
Safety and costs of blood transfusion practices in dengue cases in Brazil	(Macha do et al., 2019) Brazil	36	Prospective observation al study	he study focused on the use of blood compone nts in the managem ent of dengue cases, following WHO/PA HO recomme ndations. First Stage: Included all dengue cases hospitaliz ed from January to Decembe	Structure     d     questionn     aire for     data     collectio     n.  Analysis of medical records from hospitalize d patients with confirmed dengue diagnoses.	• 52 (16.1%) of the 323 patients receive d transfus ions of blood compon ents. • 25 (48%) receive d transfus ions accordi ng to WHO recomm endatio ns (with criteria) , while 27 (52%)

r 2010 receive during a d major transfus epidemic. ions 2. Second without Stage: criteria. Conducte d from Among March 271 2016 to patients Decembe not r 2017 transfus during a ed, 12 nonshould epidemic have year. receive d blood compon ents accordi ng to WHO guidelin es. The study highligh ts discrepa ncies in the applicat ion of WHO recomm endatio ns for blood transfus ion in dengue cases, indicati ng a need for improve d adheren ce to guidelin es to ensure

appropri

						ate clinical manage ment and resourc e utilizati on in the treatme nt of dengue patients
Prophylac tic platelet transfusio n plus supportiv e care versus supportiv e care alone in adults with dengue and thromboc ytopenia: a multicent re, randomis ed controlle d trial	(Lye et al., 2017) Singap ore	372	The study was a multicent re, randomis ed controlle d trial (RCT) with an intention -to-treat analysis. (PFTC)	The intervent ion involved administ ering prophyla ctic platelet transfusi ons (4 units of pooled platelets each day) to patients with a platelet count of 20,000 per µL or lower, in addition to supportive care.	The study utilized web-based randomisati on for participant allocation and SAS version 9.3 for statistical analyses.	The primary outcomes included the occurrence of severe bleeding and the incidence of warning signs of severe dengue by day 7. The results indicated no significant difference in these outcomes between the transfusion group and the control group.
A study protocol for a randomis ed controlle d trial evaluatin g clinical effects of	(Ypma et al., 2016) Nether lands	618	Randomis ed, single- blinded, multicent re controlle d trial	Comparis on of pathogen -reduced platelet concentr ates treated by the Mirasol	Daily monitoring of bleeding symptoms; adjudication by blinded adjudicators and an automated algorithm	Primary endpoint is the percentage of patients experienci ng WHO grade ≥2 bleeding complicati

platelet transfusio n products: the Pathogen Reduction Evaluatio n and Predictiv e Analytical Rating Score (PREPARe S) trial.			system versus standard plasma- stored platelet concentr ates		ons.
Effective ness of Pooled Platelet Transfusi on in Concorda nt and Discordan t Groups among Dengue Patients	(Bhat et al., 2016) India	The study utilized a randomiz ed control trial design to evaluate the effective ness of pooled platelet transfusions	Patients received pooled platelet transfusi ons, with a focus on comparin g outcome s between ABO-compatible and ABO-incompat ible transfusi ons	Platelet counts were measured using standard laboratory techniques, and Corrected Count Increment (CCI) was calculated to assess the effectivenes s of the transfusions	The study found that the median post-transfusion Platelet Count Increment (PPI) and Corrected Count Increment (CCI) were significantl y higher among responders compared to non-responders . Higher post-transfusion platelet count increments were noted in recipients of ABO-compatible pooled platelet transfusion s. The study

						concluded that ABO-compatible pooled platelet transfusion s are more effective in increasing platelet counts in dengue patients compared to ABO-incompatible transfusion s, highlightin g the importance of blood group compatibili ty in transfusion practices for better clinical outcomes.
Potential Harm of Prophylac tic Platelet Transfusi on in Adult Dengue Patients	(Lee et al., 2016) Vietna mes	788	The study utilized a retrospec tive cohort design, analyzing clinical outcomes between patients who received prophylac tic platelet transfusions and those who did not.	The intervent ion was prophyla ctic platelet transfusi on administ ered to patients with critically low platelet counts.	Data collection involved clinical records, which included demographic information, clinical features, laboratory results, and treatment outcomes. Statistical analyses were performed using logistic regression	The outcomes measured included: Incidence of clinical bleeding Platelet increment the day after transfusion Time for platelet count to exceed 50,000/mm  Length of hospital stay

					and propensity score matching	• Intensive care unit (ICU) admission • Mortality Conclusion The study concluded that prophylactic platelet transfusion in adult dengue patients with low platelet counts may not be beneficial and could potentially lead to adverse outcomes. The findings suggest a need for careful consideration of transfusion practices in this patient population.
Lack of Efficacy of Prophylac tic Platelet Transfusi on for Severe Thromboc ytopenia in Adults with Acute Uncompli	(Lye et al., 2015) Singap ore	256	This was a retrospec tive cohort study conducte d in the	The intervent ion involved prophyla ctic platelet transfusi on, which was defined as platelet transfusi	The study utilized medical chart reviews to extract demographic characteristi cs, clinical and laboratory data, as well as treatment and outcome	Patients who got transfusion s and those who did not were compared for subsequent bleeding, platelet increase, and platelet recovery.

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			frequently seen in
			acute
			dengue
			infections,
			the study found that
			it was not
			a reliable
			indicator
			of serious
			bleeding.

# RESEARCH RESULT **Study Characteristics**

The nine included studies represented a mix of randomized controlled trials (RCTs) and retrospective cohort analyses. Sample sizes ranged from 74 to 7,500 patients, and the studies were conducted in diverse settings, including Singapore, India, Brazil, and Pakistan. The study populations primarily included hospitalized dengue patients with thrombocytopenia, defined as platelet counts below 20,000/mm<sup>3</sup> (Assir et al., 2013). **Key Findings** 

1. Prophylactic Platelet Transfusion: Several studies (e.g., Lye et al., 2016) found no statistically significant reduction in clinical bleeding rates among patients receiving prophylactic platelet transfusion compared to those managed with supportive care alone.

ln some cases, transfused patients experienced delayed platelet recovery compared to non-transfused counterparts.

2. Clinical Bleeding and Recovery:

Transfusion often delayed platelet recovery by 1-2 days and extended hospitalization duration. For instance, patients in the transfused group in Lye et al. (2016) required an average of one additional day to achieve platelet counts above 50,000/mm<sup>3</sup>.

3. Adverse Events: Adverse events such as mucosal bleeding and fluid overload were more frequently observed in patients receiving transfusions.

Risks associated with transfusion included prolonged hospital stays and potential complications like infections or allergic reactions.

4. Subgroup Analysis:

Outcomes varied based on baseline severity and initial platelet levels. Sundar et al. (2019)demonstrated that patients with lower baseline platelet counts (<10,000/mm<sup>3</sup>) had marginally better responses

to transfusion in terms of platelet increment but still experienced extended hospital stays(Kaur & Kaur, 2014).

The findings indicate that prophylactic platelet transfusion does not significantly reduce bleeding incidence or improve recovery rates in dengue patients. It may be associated with delayed recovery, platelet prolonged hospitalization, and increased risk of adverse events.

### DISCUSSION

The findings indicate that prophylactic platelet transfusion does not offer a clear clinical benefit managing dengue-associated thrombocytopenia. Instead, it may contribute to slower endogenous platelet production, potentially due to suppression of thrombopoietindriven recovery. Furthermore, the additional risks associated with transfusion, such as fluid overload and increased resource utilization, highlight the need for restrictive transfusion practices. (Newland et al., 2019).

Clinical Implications

- 1. Restrictive Transfusion Policies: Platelet transfusion should be reserved for patients with active clinical bleeding or critical thrombocytopenia with imminent bleeding risk. Current evidence does not support routine prophylactic use.
- 2. Alternative Strategies:

Enhanced monitoring and supportive care. including hydration adequate management of comorbidities, may provide safer and more effective outcomes for most dengue patients.

Limitations

1. Study Design Variability:

Differences in methodologies, including criteria for transfusion and patient management protocols, may limit the generalizability of findings.

2. Lack of Longitudinal Data:

Few studies evaluated longoutcomes beyond hospitalization, such recurrence of thrombocytopenia or delayed complications.

3. Population-Specific Factors:

Variations in healthcare infrastructure and transfusion protocols across study regions could influence outcomes.

## **CONCLUSION**

The evidence from this systematic review suggests that platelet transfusion should not be used prophylactically in dengue fever patients with thrombocytopenia in the absence of active bleeding. Clinicians should adopt evidence-based, restrictive transfusion policies to minimize potential risks and optimize resource utilization. Further high-quality randomized controlled trials are essential to establish definitive

guidelines for platelet transfusion in dengue management.

### **REFERENCES**

- Archuleta, S., Chia, P. Y., Wei, Y., Syed-Omar, S. F., Low, J. G., Oh, H. M., Fisher, D., Ponnampalavanar, S. S. L., Wijaya, L., Kamarulzaman, A., Lum, L. C. S., Tambyah, P. A., Leo, Y. S., & Lye, D. C. (2020). Predictors And Clinical Outcomes Of Poor Platelet Recovery In Adult Dengue With Thrombocytopenia: Multicenter, Prospective Study. Infectious Clinical 383-389. Diseases, 71(2), Https://Doi.Org/10.1093/Cid /Ciz850
- Andriastuti, M., Chozie, N. A., & Rahmani, S. Laporan Kasus Berbasis Bukti **Efektivitas** Transfusi Trombosit Hasil Uji Silang Serasi (Crossmatch) Pada Kondisi Trombositopenia Refrakter.
- E. Asmarani, (2021). Penatalaksanaan Trombositopenia Pada Pasien Dengue Hemorrhagic Fever (Doctoral Dissertation, Stikes Insan Cendekia Medika Jombang).
- Assir, M. Z. K., Kamran, U., Ahmad, H. I., Bashir, S., Mansoor, H., Anees, S. Bin, & Akram, J. Effectiveness (2013).Of Platelet Transfusion In Dengue Randomized Fever: Α Controlled Trial. Transfusion Medicine And Hemotherapy, 362-368. Https://Doi.Org/10.1159/000 354837
- Barker, T. H., Stone, J. C., Sears, K., Klugar, M., Tufanaru, C., Leonardi-Bee, J., Aromataris, E., & Munn, Z. (2023). The Revised Jbi Critical Appraisal

- Tool For The Assessment Of Risk Of Bias For Randomized Controlled Trials. Jbi Evidence Synthesis, 21(3), 494-506. Https://Doi.Org/10.11124/Jb ies-22-00430
- Bhat, A., Chowdappa, V., & Masamatti, S. (2016).S. Effectiveness Of **Pooled** Platelet Transfusion In Concordant And Discordant Groups Dengue Among Patients. Journal Of Clinical And Diagnostic Research, Ec21-Ec24. 10(7), Https://Doi.Org/10.7860/Jcd r/2016/19278.8213
- Chaudhary, R., Khetan, D., Sinha, S., Sinha, P., Sonker, A., Pandey, P., ... & Ray, V. (2006). Transfusion Support Dengue Patients In A Hospital Based Blood Transfusion Service In North India. Transfusion And Apheresis Science, 35(3), 239-244.
- P., & Kaur, G. (2014). Kaur, Transfusion Support In Patients With Dengue Fever. International Journal Of Applied And Basic Medical Research, 4(3), 8. Https://Doi.Org/10.4103/222 9-516x.140708
- Lee, T. H., Wong, J. G. X., Leo, Y. S., Thein, T. L., Ng, E. L., Lee, L. K., & Lye, D. C. (2016). Potential Harm Of **Platelet** Prophylactic Transfusion In Adult Dengue Patients. Plos Neglected Tropical Diseases, 10(3), 1-10. Https://Doi.Org/10.1371/Jou rnal.Pntd.0004576
- Lye, D. C., Archuleta, S., Syed-Omar, S. F., Low, J. G., Oh, H. M., Wei, Y., Fisher, D., Ponnampalavanar, S. S. L., Wijaya, L., Lee, L. K., Ooi, E. E., Kamarulzaman, A., Lum, L. C., Tambyah, P. A., & Leo, Y.

- S. (2017).Prophylactic Platelet Transfusion Plus Versus Supportive Care Supportive Care Alone In With Adults Dengue And Thrombocytopenia: Multicentre, Open-Label, Randomised, Superiority Trial. The Lancet, 389(10079), 1611-1618.
- Https://Doi.Org/10.1016/S01 40-6736(17)30269-6
- Lye, D. C., Lee, V. J., Sun, Y., & Leo, Y. S. (2009). Lack Of Efficacy Of Prophylactic Platelet Transfusion For Severe Thrombocytopenia In Adults With Acute Uncomplicated Dengue Infection. Clinical Infectious Diseases, 48(9), 1262-1265. Https://Doi.Org/10.1086/597
- Machado, A. A. V., Negrao, F. J., Croda, J., De Medeiros, E. S., & Pires, M. A. D. S. (2019). Safety And Costs Of Blood Transfusion **Practices** Dengue Cases In Brazil. Plos 14(7), 1-14. Https://Doi.Org/10.1371/Jou rnal.Pone.0219287
- Manoharan, A., Janarthanam, V., Srivastava, R., Sowmya, S., & N. (2016).Gandhi, Effectiveness Of Platelet Transfusion In Dengue Fever. Int J Res Rev. 3, 5-9.
- Makroo, R. N., Raina, V., Kumar, P., & Kanth, R. K. (2007). Role Of Platelet Transfusion In The Management Of Dengue Patients In A Tertiary Care Hospital. Asian Journal Of

- Transfusion Science, 1(1), 4-7.
- Newland, Bentley, Α., Jakubowska, A., Liebman, H., Lorens, J., Peck-Radosavljevic, M., Taieb, V., Takami, A., Tateishi, R., & Younossi, Z. M. (2019). A Systematic Literature Review On The Use Of Platelet Transfusions In Patients With Thrombocytopenia. Hematology (United
  - Kingdom), 24(1), 679-719. Https://Doi.Org/10.1080/160 78454.2019.1662200
- Prashantha, B., Varun, S., Sharat, D., Murali Mohan, B. V., Ranganatha, R., Shivaprasad, Naveen, Μ. (2014).Prophyactic Platelet Transfusion In Stable Dengue Fever Patients: Is It Really Necessary?. Indian Journal Of Hematology And Blood Transfusion, 30, 126-129.
- Ypma, P. F., Van Der Meer, P. F., Heddle, N. M., Van Hilten, J. A., Stijnen, T., Middelburg, R. A., Hervig, T., Van Der Bom, J. G., Brand, A., & Kerkhoffs, J. L. H. (2016). A Study Protocol For A Randomised Controlled Trial Evaluating Clinical Effects Of Platelet Transfusion Products: The Pathogen Reduction Evaluation And Predictive Analytical Rating Score (Prepares) Trial. Bmj Open, 1-12. Https://Doi.Org/10.1136/Bmj open-2015-010156