

Big data modeling to predict platelet usage and minimize wastage in a tertiary care system

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Maintaining a robust blood product supply is an essential requirement to guarantee optimal patient care in modern health care systems. However, daily blood product use is difficult to anticipate. Platelet products are the most variable in daily usage, have short shelf lives, and are also the most expensive to produce, test, and store. Due to the combination of absolute need, uncertain daily demand, and short shelf life, platelet products are frequently wasted due to expiration. Our aim is to build and validate a statistical model to forecast future platelet demand and thereby reduce wastage. We have investigated platelet usage patterns at our institution, and specifically interrogated the relationship between platelet usage and aggregated hospitalwide patient data over a recent consecutive 29-mo period. Using a convex statistical formulation, we have found that platelet usage is highly dependent on weekday/weekend pattern, number of patients with various abnormal complete blood count measurements, and location-specific hospital census data. We incorporated these relationships in a mathematical model to guide collection and ordering strategy. This model minimizes waste due to expiration while avoiding shortages; the number of remaining platelet units at the end of any day stays above 10 in our model during the same period. Compared with historical expiration rates during the same period, our model reduces the expiration rate from 10.5 to 3.2%. Extrapolating our results to the \sim 2 million units of platelets transfused annually within the United States, if implemented successfully, our model can potentially save ~80 million dollars in health care costs.

prediction | blood | supervised learning

Blood products, including red blood cells, plasma, and platelets, are therapeutics essential to daily in-patient hospital care. Many modern medical interventions would be rendered impossible without these precious resources. Although it is a certainty that these products are needed in any hospital setting, it is uncertain how much and when each product is needed. Additionally, all these products are perishable, the shortest practical shelf life in many cases being only 3 d for platelets. Platelet transfusions are generally used to mitigate or prevent bleeding in a variety of patient populations, including actively bleeding patients (e.g., trauma, surgery) and patients with dysfunctional or insufficient platelets. This combination of uncertain quantity of demand, product perishability, and expense makes predicting hospital-wide platelet usage a highpriority endeavor for all hospitals and blood banks.

Key goals for blood product management in health care systems are to both (i) always have enough products available to keep up with patient demand even in cases of emergencies and (ii) minimize the number of products wasted due to expiration. The primary aim of medical professionals is always to optimize patient care, leading transfusion services and blood centers to err on the side of overstocking, resulting in the expiration and wastage of some platelet products.

Provided by volunteer donors, platelets are processed by community blood centers that manufacture, test, and distribute these blood components to their hospital customers. Platelets products donated by single donors (apheresis platelet) are the most challenging and costly to collect, store, test, and manage, from an inventory perspective (1-3). Difficulties include fewer apheresis platelet donors than whole blood donors, longer donation process than a whole-blood donation, less capacity for collection due to more complicated and costly equipment required, and a practical shelf life of only 3 d, due to 2 d sequestration in testing (1, 4, 5). Furthermore, because products are not available on the day of collection and because daily usage is quite variable, collection strategies must attempt to match current inventory levels with highly variable and unpredictable future usage.

Therefore, a common strategy employed by many blood centers is to aim for an expiration rate of 8 to 9%, with 2 million units transfused annually and the outdated rate for apheresis platelets in 2013 being 11.0% of total units produced (6), effectively producing a surplus buffer so as to always have enough platelet units on the shelf for unanticipated patient demand. However, in large tertiary care settings where close to 15,000 units of platelets are potentially transfused annually, this wastage may translate to close to \$1 million in expiration alone, depending on the geographically determined price of a unit of platelets. In addition to the cost of the product, there are also indirect costs such as staffing/nursing costs to receive and return the products, and opportunity costs of donors' hours of donation time. Although the wastage appears large, it is seen as a necessary precaution to have enough products on hand at all times to meet the highly variable and unpredictable patient demand.

Hospitals and blood banks have not yet developed a feasible method for predicting fluctuations in product usage. We aim to build and validate a reliable statistical model for blood banking

Significance

In modern hospital systems where complicated, severely ill patient populations are the norm, there is currently no reliable way to forecast the use of perishable medical resources to enable a smart and economic way to deliver optimal patient care. We here demonstrate a statistical model using hospital patient data to quantitatively forecast, days in advance, the need for platelet transfusions. This approach can be leveraged to significantly decrease platelet wastage, and, if adopted nationwide, would save approximately 80 million dollars per year. We believe our approach can be generalized to all other aspects of patient care involving timely delivery of perishable medical resources.

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that would have a highly significant impact in improving quality, safety, access to care, and cost of care. Measures to increase efficiency and cost-effectiveness in health care are becoming increasingly important (7, 8), and an accurate predictive algorithm for blood product usage could greatly contribute to value-based care in transfusion medicine.

Due to the critical need for platelets nationwide (9), and the particularly short shelf life of platelet products, there is an unmet need for a system that can forecast platelet usage in a hospital setting. In the care of each patient, the decision to transfuse platelets is not random. Previous studies have investigated patient parameters in specific clinical scenarios as an indicator for platelet usage (10-15). Many of these have focused on small patient populations, and manual chart reviews. Among other results, one study indicated that day-of-the-week status could help in predicting platelet usage (14).

Here, we forecast hospital-wide platelet needs a priori by leveraging the "big data" available within modern hospital electronic medical records (EMR). We hypothesized that looking into hospital-wide patient data associated with clinical transfusion decisions would allow us to accurately predict platelet usage 3 d in advance, allowing ample time for blood centers to alter collection and importing strategies. Our institution, Stanford Blood Center (SBC), is the community blood center that provides all blood products used at both our associated hospitals, Stanford Health Care (SHC) and Lucille Packard Children's Hospital (LPCH). We have developed a statistical model at our institution, using shared clinical data, that forecasts platelet demand 3 d in advance, and reduces combined wastage from expiration by the blood center and transfusion service. Using recent historical data comprising 29 consecutive months, our model reduces the expiration rate by two-thirds (from 10.5 to 3.2%) over the same period. In addition to reducing expiration due to wastage, this model prohibits platelet shortages; there are at least 10 units remaining on the shelf at the end of each day, thereby never compromising patient care. With an annual transfusion rate of ~13,000 units at our institution, this could potentially reduce our wastage by 950 units annually without compromising patient care. Extrapolating to total platelet usage of 2 million units in the United States per year, if implemented successfully, this approach could save 80 million dollars in annual health care costs.

Materials and Methods

Inventory Flow. The majority of blood products collected at SBC is distributed to the Transfusion Service located in the SHC adult hospital (SHC-TS). SHC-TS acts as a central transfusion hub from which blood products are distributed to various units and locations within SHC, as well as to LPCH. All of the data for transfusable products, to both SHC and LPCH, are managed and recorded by SHC-TS. Additionally, essentially all of the blood products issued for transfusion through SHC-TS are supplied by SBC.

SHC EMR Data Acquisition. We obtained Stanford IRB approval to perform chart review (Protocol ID 350350) of SHC EMR. Through the Stanford Translational Research Integrated Database Environment, we obtained SHC data within the date range January 1, 2013 through June 1, 2015. The dataset includes complete blood count (CBC) and hospital census by location. We computationally cleaned the data and aggregated them into a useable relational database for further analyses. In combination with day-of-week status and number of transfused products from the previous week, these variables account for 43 covariates (Tables S1 and S2 show the details for covariates other than day-of-week status). From LPCH, CBC and hospital census data were unavailable for research purposes during the same time period.

Transfusion and Expiration Data Acquisition. All of the transfusion data and a portion of the expiration data within the specified date range were obtained from the SafeTrace Tx (Haemonetics) database system at SHC-TS. These data represented transfused products for both SHC and LPCH. For each product, we obtained the donation identification number, issue date and time, expiration date and time, discard date and time, ABO blood group status, and Rh status. Due to the inventory management procedure in place between SBC and SHC-TS, a large fraction of the expired platelet products is shipped back to SBC. Therefore, total platelet wastage due to expiration was determined by combining the number platelet units that expired at SBC and SHC-TS. Expired platelet products at SBC were obtained from the SafeTrace (Haemonetics) system at SBC.

Statistical Modeling. Our goal is to minimize platelet wastage while ensuring we have enough platelet units to meet patient demand every day. Since the objective is not prediction but an optimal ordering strategy, standard supervised learning is not ideal for this problem. Hence we developed a constrained optimization strategy. To achieve our goal, accurate prediction for the next day is not sufficient, as the acquired platelets spend the first 2 d in testing and will only be available for transfusion on the third day. To adequately account for the demand in 3 d, we instead built a prediction model for the platelet usage for the next 3 d, with an objective to minimize the waste. Moreover, on each day, we aim to collect a number of platelet units such that (i) we will have sufficient platelets to cover the predicted demand and (ii) we will have enough supply on the shelf in cases of unpredictable emergency situations.

To formalize our model, we first introduce the following notation. Let y_i be the number of platelet units used on day i, x_i be the number of fresh platelet units arriving on day *i*, $r_i(1)$ be the number of platelet units that expired on day i + 1, $r_i(2)$ be the number of platelet units that expired on day i + 2, z_i be the vector of 43 covariates measured on day *i*, and w_i be the number of platelet units wasted on day i. For our model, it was important to forecast the platelet demand at least 3 d into the future. Collection occurs on day i + 1, and platelets will be available for use on days i + 3 through i + 5. Therefore, on day i (i.e., 1 d before collection and 3 d before product availability), a forecast would be helpful, since there is still actionable time to maneuver and adapt certain donor recruitment strategies.

Using the 43 covariates available from the hospital data, we predict the platelet usage for the next 3 d to be

$$t_i = \boldsymbol{z}_i^T \boldsymbol{\beta},$$

where β is the vector of coefficients for each of the covariates. We solve the following optimization problem,

$$(\beta) = \sum_{i=1}^{n} w_i + \lambda \|\beta\|_1,$$
 [1]

subject to

3 d's total need
$$t_i = x_i^T \beta_i$$
 [2]

new arrival (collecting strategy) $x_{i+3} = t_i - r_i(1) - r_i(2) - x_{i+1} - x_{i+2}$, [3]

waste
$$w_i \ge [r_{i-1}(1) - y_i]_+$$
 [4]

actual remaining $r_i(1) = [r_{i-1}(2) + r_{i-1}(1) - y_i - w_i]_+$ [5]

$$r_i(2) = \left[x_i - \left[y_i + w_i - r_{i-1}(2) - r_{i-1}(1) \right]_+ \right]_{\perp}$$
[6]

fresh units remaining
$$r_i(2) \ge c_0$$
. [7]

Here the subscript + indicates a positive value, such that $[x]_{\perp}$ is equal to x if x > 0 and zero otherwise. Eq. 1 is the objective we want to minimize, which is the number of total wasted units with a sparsity penalty on the underlying coefficients, similar to the lasso method (16). Eq. 2 is the predicted total units for the next 3 d using the 43 predictors detailed above. Eq. 3 gives a collection strategy for platelets that ensures we collect enough platelet units to cover the third day's predicted usage. Eqs. 4-6 detail the calculations for the wasted units and leftover units. For Eq. 4, the original mathematical definition of the waste is an equality, but we have relaxed it to an inequality to make the problem convex. In Eq. 7, we place a constraint that the remaining freshly received platelets on the shelf should not be less than a certain threshold c_0 . For a major hospital with ~40 daily platelet transfusions, we set the threshold at 30. The index is shifted 3 d into the future because the units we decide to collect on day *i* will not be available for transfusion until day i + 3.

The above optimization problem can be expressed as a convex optimization problem with linear constraints. We solve the above problem using the R language library lpSolve (17). The penalty parameter λ is chosen through eightfold cross-validation.

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Results

Platelet Transfusion and Expiration Patterns. Within the specified date range, we obtained more than 120,000 transfusion records. Of these 120,000 transfusions, ~30,000 were platelet transfusions, with platelet transfusions occurring on every day over the 29-mo period. As shown in Fig. 1*A*, daily platelet usage was highly variable, showing a transfused daily average of 35.4 units and a standard of deviation of 9.2. To gain a better understanding of the daily usage patterns of platelets, we investigated whether daily usage varied across different months or day of week (Fig. 1 *B* and *C*). As expected, platelet usage was statistically significantly higher on the weekdays compared with weekends (mean 38.4 vs. 27.8, *P* value = 2.2e-16). These results are summarized by day of week in Table S3. There was a significant difference in daily usage rates across different months, as determined by one-way ANOVA (*F* = 4.441, *P* = 1.57e-06).

We additionally explored the rate of daily platelet wastage due to expiration. Similar to the daily variation in transfused products, we also found that platelet expiration varied daily (Fig. 2*A*). On average, 3.7 units of platelets were expired per day, with a SD of 6.7. We next interrogated daily expirations by day of week and month (Fig. 2 *B* and *C*). There were statistically significant differences in wastage when comparing weekday to weekend (weekend mean = 2.56, weekday mean = 4.09, P = 5.4e-04) as well as among the different months as determined by one-way ANOVA (F = 4.9, P = 2.16e-07).

During this 29-mo period, ~3,200 units expired either at SHC-TS or SBC. This translates into an expiration rate of 10.3% during this 29-mo period, containing both the training and validation sets (discussed in *Minimizing Platelet Expiration*). For the time period corresponding to the validation set, the expiration rate was 10.5%.

Predictors of Platelet Usage. To accurately predict platelet usage and minimize wastage, we first investigated which covariates within the data had strong correlations with platelet usage. We have CBC data for the SHC patients for every day during the 29-mo period, constituting ~70% of the daily transfusions. The missing CBC data for the approximately remaining 30% of the transfusions mainly resided within LPCH, the pediatric hospital, because we did not have access

to its EMR. Given that we had high-quality data for the majority of transfusions, we proceeded with an aggregative strategy for analyses.

For each given day i, for each CBC value, we averaged the daily number of patients with an abnormal value (defined in Tables S1 and S2) over the previous 7 d. From the hospital census data, we used the number of patients in each specified unit location for day i. These metrics were also used as covariates in our model. Furthermore, since day-of-the-week had a clear effect on platelet usage (Fig. 1), we used the day-of-the-week status of day i as an additional covariate. Moreover, we discovered that past platelet usage had a clear pattern of autocorrelation, with sustained periods of high or low demands. Thus, we also averaged the daily number of platelet units transfused over the previous 7 d, and included that metric as a covariate.

CBC measurements such as platelet count (PLT) and mean corpuscular hemoglobin concentration (MCHC) are influential predictors. The number of patients in the H1 unit—in which resided Neurosurgery, Trauma, and Vascular patients—also has a strong positive correlation with high platelet usage. The 36 covariates are listed in Tables S1 and S2, which includes, for each covariate, its mean, SD, range, the definition we used as abnormal for each CBC value, and the approximate medical services associated with each unit location. The predictors include the daily average number of platelet units transfused in the past week, daily average number of people with each abnormal CBC measurement over the previous 7 d, and the number of patients in different units in the hospital. In addition to these 36 covariates, we also used the day-of-week status, which gave us a total of 43 covariates.

We solved the optimization problem detailed in *Statistical Modeling* and obtained the following values: (*i*) β , a vector of length 43 containing the weights for the 43 covariates used in the model, and (*ii*) x_i , the number of fresh platelet units coming on day *i*.

The ℓ_1 penalty used in the minimization (Eq. 1) enforces sparsity in the 43 coefficients. Therefore, only 35 of the 43 weights were estimated as nonzero. In Table S4, we list the top 16 coefficients from the CBC and census data, in addition to dayof-week status and previous week's platelet usage, with the highest absolute weights.

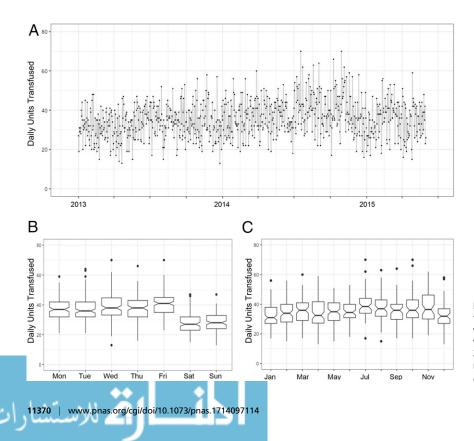
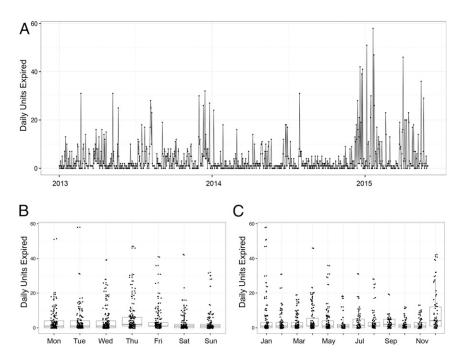


Fig. 1. Daily platelet usage patterns. (A) Daily usage from January 1, 2013 through May 31, 2015 is highly variable. (B) Mean daily platelet usage is higher on weekdays than weekend (38.4 vs. 27.8, P value = 2.2e-16). (C) Mean daily platelet usage varies with month as determined by one-way ANOVA (F = 4.44, P = 1.57e-06).

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We see from Table S4 that the most prominent predictors are the average platelet usage for the last week (4.21) and the number of patients in the H1 unit, housing Neurosurgery, Trauma, and Vascular patients. It is also worth noticing that there is a clear weekday/weekend effect. The platelet usage for the next 3 d is substantially lower if it covers the weekend (Friday: -3.14), and starts increasing on weekdays (Sunday: 1.95). Surprisingly, the number of patients with low PLT was assigned a negative weight; we would expect that more patients with abnormally low PLT **Fig. 2.** Daily platelet expiration wastage patterns by day of week and month. (*A*) Similar to daily transfusion rates, daily expiration rates from January 1, 2013 through May 31, 2015 is also highly variable. (*B*) Mean daily platelet expiration wastage is higher on weekdays than weekend (4.09 vs. 2.56, *P* value = 5.4e-04). (*C*) Mean daily platelet expiration wastage varies with month as determined by one-way ANOVA (*F* = 4.90, *P* = 2.16e-07).

would require more transfusions. However, as shown in Table S4, one of the strongest predictors of future platelet usage is the platelet usage during the past 7 d. Furthermore, as shown in Fig. 3, there is a correlation between PLT and red blood cell count (RBC). Therefore, the negative weight of low PLT on future platelet demand is the high correlation of the low PLT measurement with the RBC measurement as well as the platelet usage during the past week, both of which have already been assigned positive weights. In fact, when we look at the marginal correlations

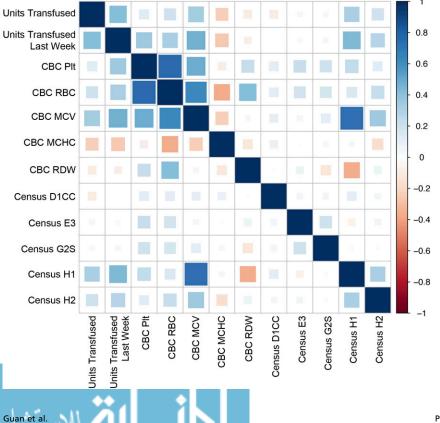
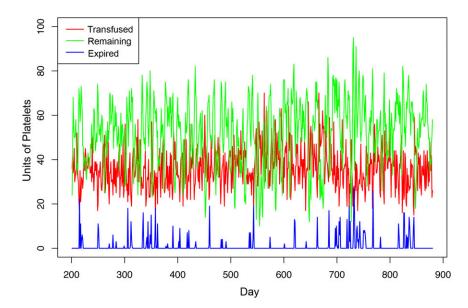


Fig. 3. Marginal correlation plot between the platelet usage and the selected predictors, with correlations coded from +1 (dark blue) to -1 (dark red), as indicated by the heat map legend on the right. Abnormal CBC values are aggregated as the average daily number of patients with specific abnormal CBC (MCHC; MCV, mean corpuscular volume; Plt, platelet count; RBC; RDW, red cell distribution width). Census data are reported as the number of patients in the indicated unit for the day.

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(Fig. 3), we can see that low PLT measurements have a weak positive correlation with platelet usage.

Minimizing Platelet Expiration. For subsequent analyses involving platelet usage prediction, we used a dynamic training and prediction model. We first split the data into training and validation sets. For the training set, we used all data from the first consecutive 200 d. Using the base model established from the training set, we proceeded to predict platelet demand and calculate appropriate collection volumes every day from day 201 onward. To account for potential changes in patient populations and practices during the period, we retrained our model every 7 d. For example, we start with the first 200 d to build our model and make predictions for number of units to collect per day from day 201 to day 207. We then retrained our model by incorporating the data from days 201 through 207, and then used this updated model to make daily predictions for number of units to collect from day 208 to day 214. This process continues until day 880, the end of our study period. Therefore, our validation set is considered to be days 201 through 880, since each daily prediction for that timeframe was based on a model that did not previously include that day.

By solving the optimization problem as described in *Statistical Modeling* and following our testing procedure described above, we obtained a collection strategy to collect x_{i+3} units on each day to ensure enough supply for the third day in the future. Adopting this strategy and testing it on the validation set, we find that we can reduce wastage to 3.2%, compared with the historical waste of 10.5% over the same time period as the validation set. Annualized, this corresponds to a reduction in the number of wasted platelet products from ~1,400 per year to just 430 per year.

Most importantly, although we significantly reduced wastage from expiration, we ensured that there is never a day where platelet inventory falls short and that there is always enough supply on the shelf every day. We determined the end-of-day remaining units and the wasted units on our modeled collection strategy on the validation set. Fig. 4 illustrates the actual daily units transfused during the validation period, overlaid with our modeled end-of-day remaining units and wasted units. Using our modeled collection strategy, there was not a single day when we were short of inventory, when patients' platelet demand outpaced platelet supply. Additionally, at the end of each day, there was always a healthy remainder of platelet units on the shelf, with the lowest being 10 units remaining on hand.

In Fig. 5, we further illustrate the cumulative units wasted over a 2-y period in comparison with the cumulative number of units

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Fig. 4. Daily platelets transfused (red line), modeled remaining (green line), and modeled wastage due to expiration (blue line) during the validation period (from day 201 to day 880), with $c_0 = 30$. With the model during the validation set, there was never a day wherein there were not enough platelets on the shelf. The number of units remaining at the end of each day stayed above 10, and the expiration rate during this period was reduced to 3.2%.

transfused. There is a substantial reduction in wasted platelet units, using our modeled method compared with the historical wastage during the same time period. We also see that the number of wasted units is less variable over time under our modeled strategy.

Discussion

Blood products remain essential to modern medicine, and it is crucial to maintain enough supply within each hospital's transfusion service to adequately meet patient demand, which has been mostly unpredictable. Due to their short shelf life, platelet products are most severely affected by fluctuations in usage. Combined with a high cost of production, storage, and testing, appropriate inventory management of platelet products is even more crucial in

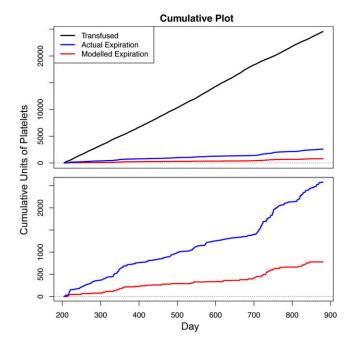


Fig. 5. Cumulative plot for historical number of platelet units transfused (black line), historical waste due to expiration (blue line), and projected waste due to expiration under our model (red line), during the validation period from day 201 to day 880. The total number of units historically transfused and wasted during this period was 24,700 and 2,600, respectively. Using our model, the cumulative waste was reduced to 780, representing a reduction in expiration rate from 10.5 to 3.2%.

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light of their therapeutic indications and efficacy. We demonstrate a statistical model that can adequately forecast patient platelet demand to guide inventory management and decrease wastage due to expiration, while avoiding platelet shortages.

At our institution, where $\sim 13,000$ apheresis platelet products are transfused annually, the daily usage variation is high, with the daily average being 35.4 units per day, and a 25.9% coefficient of variation. These initial findings demonstrate the difficulty of accurately estimating platelet usage. Approximation cannot be based solely on historical mean daily usage. The current strategy employed at our institution is approximating by daily median usage, resulting in an historical expiration rate of 10.3% during this studied time period. More importantly, historical transfusion rates cannot account for changes in patient population, physician practices, or other factors that can dynamically influence platelet usage.

Based on these considerations, we investigated the relationship between platelet usage and in-patient hospital data, including census, CBC, day of week, and number of platelet units transfused during the previous week. We interrogated a total of 43 covariates and found 35 covariates to be significant for predicting platelet usage, using l_1 to enforce sparsity. The strongest predictors of future platelet usage are the number of platelets transfused the previous week, the number of platelets transfused the previous week, the number of patients within our hospital's sublocation H1 (which includes the Trauma, Neurosurgery, and Vascular Care patients), and the day of the week.

The data furthermore indicate that this relationship can be leveraged to predict future usage. From this, we developed a model that would guide current platelet collection strategy based on a predicted future platelet demand, thus allowing some time for optimizing donor recruitment strategies. The model prohibits a shortage of platelets for any day while minimizing the number of expired products. We used a dynamic and predictive model that could account for potential changes in patient populations and transfusion practices. Using our model, the expiration rate was reduced to 3.2% compared with the historical 10.5% over the same period. At our institution, this difference would result in ~950 units saved annually. If successfully implemented and extrapolated nationwide where there are ~ 2 million units of platelets transfused, with an approximate 11% expiration rate, our approach can potentially save 80 million dollars in health care costs. Furthermore, and most importantly, this reduction in wastage is accomplished while still maintaining a robust platelet inventory at the end of each day, in the event of unpredictable surges in patient demand (e.g., trauma).

Our model can be further refined by obtaining patient data from LPCH (our institution's pediatric hospital) to accompany the transfusion data. There are technical difficulties in obtaining

- Tynngård N (2009) Preparation, storage and quality control of platelet concentrates. Transfus Apher Sci 41:97–104.
- van der Meer PF (2013) Platelet concentrates, from whole blood or collected by apheresis? Transfus Apher Sci 48:129–131.
- 3. Vassallo RR, Murphy S (2006) A critical comparison of platelet preparation methods. *Curr Opin Hematol* 13:323–330.
- 4. Garraud O, et al. (2016) Improving platelet transfusion safety: Biomedical and technical considerations. *Blood Transfus* 14:109–122.
- Cardigan R, Williamson LM (2003) The quality of platelets after storage for 7 days. Transfus Med 13:173–187.
- 6. Advanced Transfusion and Cellular Therapies Worldwide (2016) *Donor Hemovigilance AABB Blood Survey*. Available at www.aabb.org/research/hemovigilance/bloodsurvey/ Pages/default.aspx. Accessed March 1, 2017.
- Jaspan D (2007) Erythropoietic therapy: Cost efficiency and reimbursement. Am J Health Syst Pharm 64:S19–S29.
- Lopez-Plaza I, Weissfeld J, Triulzi DJ (1999) The cost-effectiveness of reducing donor exposures with single-donor versus pooled random-donor platelets. *Transfusion* 39:925–932.
- Estcourt LJ (2014) Why has demand for platelet components increased? A review. *Transfus Med* 24:260–268.
 Vinholt PJ, Alnor A, Nybo M, Hvas AM (2016) Prediction of bleeding and prophylactic
- platelet transfusions in cancer patients with thrombocytopenia. *Platelets* 27:547–554. 11. Covin R, et al. (2003) Factors affecting transfusion of fresh frozen plasma, platelets,
- and red blood cells during elective coronary artery bypass graft surgery. Arch Pathol Lab Med 127:415–423.

these data. One of the more obstinate obstacles in acquiring LPCH patient data during the specified date range is that the LPCH EMR changed from one system to another. We are uncertain how this change would affect the quality and consistency of our data. Furthermore, the number of LPCH platelet transfusions account for a minority compared with SHC. From these considerations, we decided to not include a data source that could be unreliable and would only account for a minority of transfusions. Additionally, our model uses aggregated data to minimize the lack of LPCH patient data.

Although our model is currently specific to platelets transfusion and wastage, this strategy can be extended to other blood products without difficulty. We focused on platelets initially because, if operationalized, wastage due to expiration of this product would see the most significant impact. Additionally, with anticipated regulatory guidance on the horizon relating to platelet storage (18– 23), there can be additional stress on the platelet inventory of the entire country. With this potential regulatory change being finalized, it becomes even more crucial to manage platelet inventory effectively to minimize wastage and maintain optimal patient care.

Furthermore, the strategy we used can be generalized to be applied to other health care-related products that have the following criteria: (i) always have to be on hand, (ii) have high costs, and (iii) are perishable (e.g., Epineprhine Auto-Injectors). If generally applied, this method represents a logical and automated approach to changing the way health care is delivered on a system-wide level. It can contribute to significant cost savings in the health care setting without ever compromising patient care. We are encouraged by the potential that these findings represent.

The first step to minimizing wastage while maintaining top-ofthe-line patient care is to understand the entire supply chain, from donor collection to patient usage. The relationship among SBC, SHC, and LPCH allows for a more fully transparent view of data to enable this type of vein-to-vein analysis. Our findings promise the potential to customize, in real time, donor recruitment strategies based on anticipated future patient demand. We plan to implement this data-driven process at our institution in the near future, using the findings herein as a starting point. We will also distribute an open source software package so that other hospitals and blood centers can implement this process.

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- Del Vecchio A, et al. (2001) Platelet transfusions in the neonatal intensive care unit: Factors predicting which patients will require multiple transfusions. *Transfusion* 41:803–808.
- Findlay JY, Rettke SR (2000) Poor prediction of blood transfusion requirements in adult liver transplantations from preoperative variables. J Clin Anesth 12:319–323.
- Critchfield GC, et al. (1985) Automatic prediction of platelet utilization by time series analysis in a large tertiary care hospital. *Am J Clin Pathol* 84:627–631.
- Kuliczkowski W, et al. (2015) Predicting bleeding risk by platelet function testing in patients undergoing heart surgery. *Clin Cardiol* 38:679–683.
- 16. Tibshirani R (1996) Regression shrinkage and selection via the lasso. J R Stat Soc 58:267-288.
- R Core Team (2013) R: A language and environment for statistical computing. R Foundation for Statistical Computing, Vienna, Austria. Available at www.R-project. org/, Accessed September 1, 2016.
- 18. Solheim BG (2008) Pathogen reduction of blood components. Transfus Apher Sci 39:75–82.
- Dodd RY (2003) Bacterial contamination and transfusion safety: Experience in the United States. Transfus Clin Biol 10:6–9.
- Bell CE, et al. (2003) Cost-effectiveness of transfusion of platelet components prepared with pathogen inactivation treatment in the United States. *Clin Ther* 25:2464–2486.
- Epstein JS, Vostal JG (2003) FDA approach to evaluation of pathogen reduction technology. *Transfusion* 43:1347–1350.
- Hess JR, Pagano MB, Barbeau JD, Johannson PI (2016) Will pathogen reduction of blood components harm more people than it helps in developed countries? *Transfusion* 56:1236–1241.
- Marks PW, Epstein JS, Borio LL (2016) Maintaining a safe blood supply in an era of emerging pathogens. J Infect Dis 213:1676–1677.

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