

TRANSFUSION MEDICINE ILLUSTRATED

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CLPC SEMINARS - SPRING 2021



OBJECTIVES

- **REVIEW VARIOUS CASE STUDIES INVOLVING TRANSFUSION MEDICINE**
- **DESCRIBE THE UNDERLYING PROBLEM FOR EACH CASE**
- **LIST THE SOLUTIONS OR THERAPIES USED TO CORRECT THE UNDERLYING PROBLEM IN EACH CASE PRESENTED**

TRANSFUSION JOURNAL

- **AABB'S SCHOLARLY, PEER-REVIEWED MONTHLY JOURNAL**
- **EACH ISSUE HAS ARTICLE UNDER "TRANSFUSION MEDICINE ILLUSTRATED"**
 - **CASE STUDY**
 - **HISTORICAL RECORD**
 - **PHOTOGRAPH OR STATISTICAL GRAPH**

THIS BLOOD IS NO GOOD!

- **PARSONS, J. ROSENBAUM, L. CROOKSTON, K. & KAMEL, H. (2014) INCIDENTAL ARTERIAL PHELOBOTOMY OF A WHOLE BLOOD DONOR. *TRANSFUSION*, 54, 1220.**



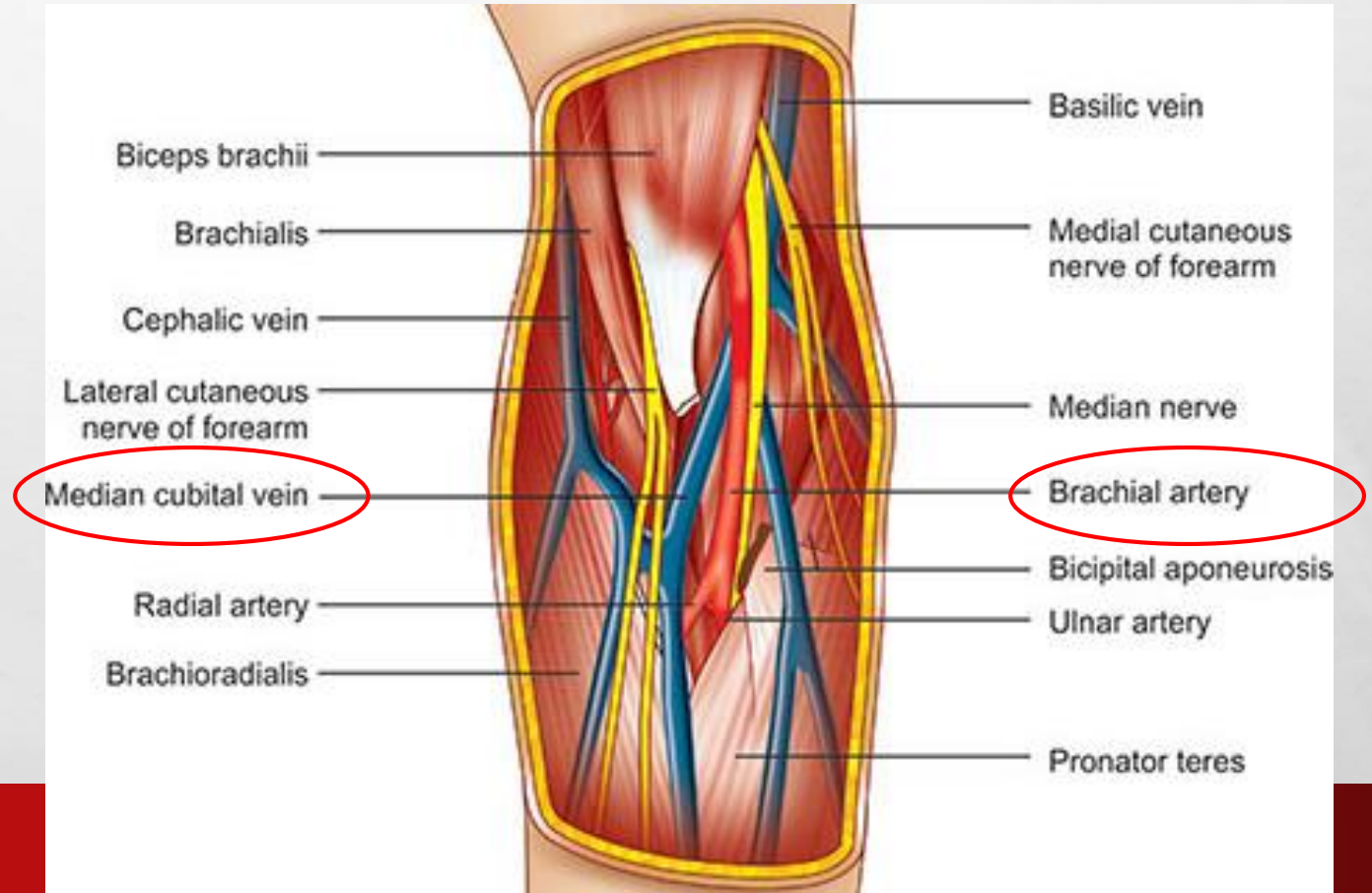
THIS BLOOD IS NO GOOD!

- **LEFT RBC UNIT COLLECTED FROM BRACHIAL ARTERY**
- **RIGHT RBC UNIT COLLECTED FROM MEDIAN CUBITAL VEIN**



THIS BLOOD IS NO GOOD!

- **MEDIAN CUBITAL VEIN VS. BRACHIAL ARTERY**
- **DON'T FORGET ABOUT NERVE DAMAGE AS WELL!**



THIS BLOOD IS NO GOOD!

- **MOST COMMON COMPLICATION FROM ARTERIAL PHLEBOTOMY ARE HEMATOMAS**
 - **33% ARTERIAL PUNCTURES VS. 0.35% IN GENERAL DONOR POPULATION**
- **RANGE OF ARTERIAL PHLEBOTOMY IS 1:9000 – 1:34,000**
 - **DEPENDS ON PHLEBOTOMISTS' EXPERIENCE AND DONOR'S UNIQUE ANATOMY**



THIS BLOOD IS NO GOOD!

- **ARTERIAL-DERIVED BLOOD IS ACCEPTABLE TO USE**
- **MOST BLOOD CENTERS HAVE PHLEBOTOMISTS REMOVE NEEDLE AND APPLY PRESSURE FOR 10 MINUTES**
- **DONORS WITH WAXING AND WANING HEMATOMAS SHOULD BE EVALUATED FOR PSEUDOANEURYSM BY ULTRASOUND**



GREEN WITH ENVY?

- **BERGLUND, D. LINDVALL, P. & LUBENOW, N. (2014)**
GREEN PLASMA. *TRANSFUSION*, 55, 245.



GREEN WITH ENVY?

- **64-YEAR OLD WOMAN WITH BREAST CANCER**
- **EDTA SAMPLE COLLECTED PRIOR TO SURGERY FOR TYPE AND SCREEN (LEFT SAMPLE)**
 - **CENTRIFUGED SAMPLE DEMONSTRATED BRIGHT GREEN PLASMA**
- **REPEAT BLOOD COLLECTION 24 HOURS LATER SHOWED NORMAL YELLOW PLASMA COLOR (RIGHT SAMPLE)**



GREEN WITH ENVY?

- **TYPE AND SCREEN RESULTS WERE NORMAL**
 - **USED GEL-BASED MICROTYPING SYSTEM**
- **PATIENT'S SURGERY WAS MAPPING AND REMOVAL OF MALIGNANT TUMORS IN LYMPH NODES**
 - **PATENT BLUE V USED TO IDENTIFY LYMPH DRAINAGE PASSAGEWAYS**



GREEN WITH ENVY?

- **PATENT BLUE V USED IN LYMPHANGIOGRAPHY AND SENTINEL NODE BIOPSIES AS A DYE TO COLOR LYMPH VESSELS**
 - **BRIEFLY GIVES PLASMA GREEN COLOR BY ADDING BLUE TO NORMAL YELLOW PLASMA**
 - **CAN CAUSE SEVERE ALLERGIC REACTIONS**
 - **CAN INTERFERE WITH O₂ SATURATION LEVELS USING PULSE OXIMETRY**



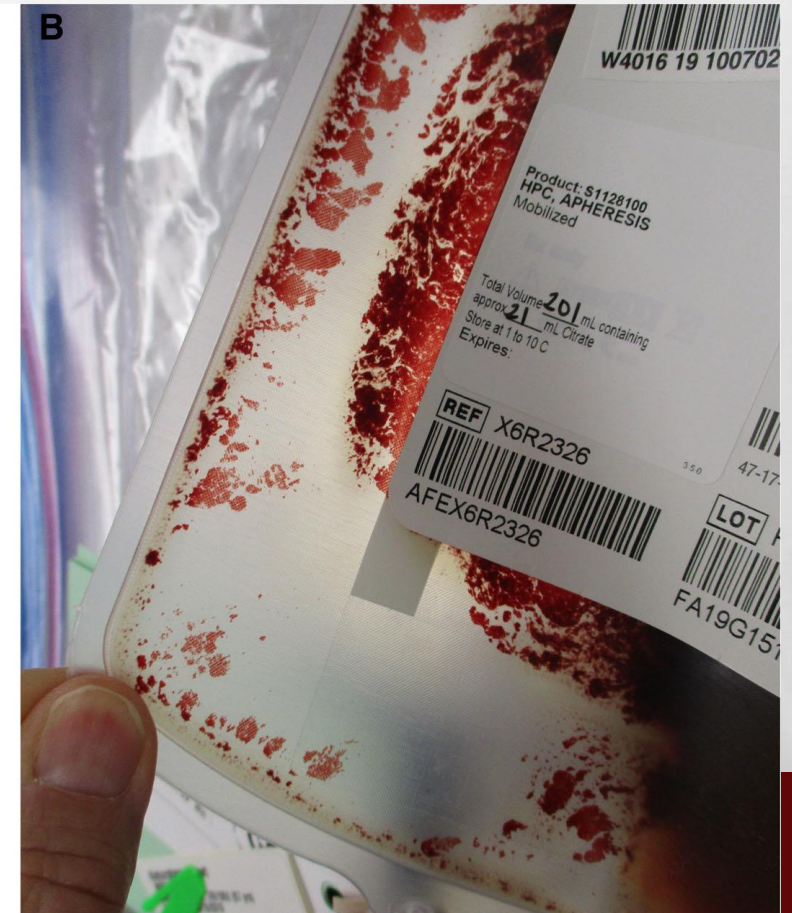
GREEN WITH ENVY?

- **GREEN PLASMA IS ALSO SEEN IN INCREASED CERULOPLASMIN LEVELS**
 - **ASSOCIATED WITH ESTROGEN-CONTAINING BIRTH CONTROL PILLS**



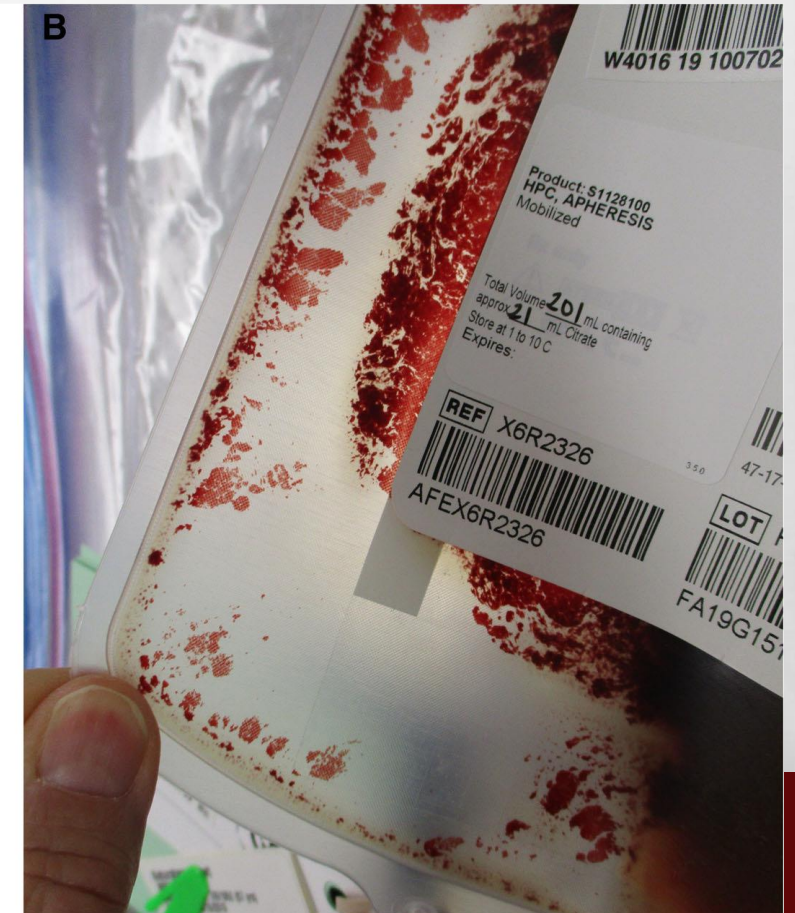
CLUMPING IN THE COLD

- **YUAN, S. ZHUANG, L. STINSON, S. BARTON, P. NOELLER, J. GARCIA, A. & WANG, S. (2020) RED CELL CLUMPING IN STEM CELL COLLECTION BAG. *TRANSFUSION*, 60, 673-674.**



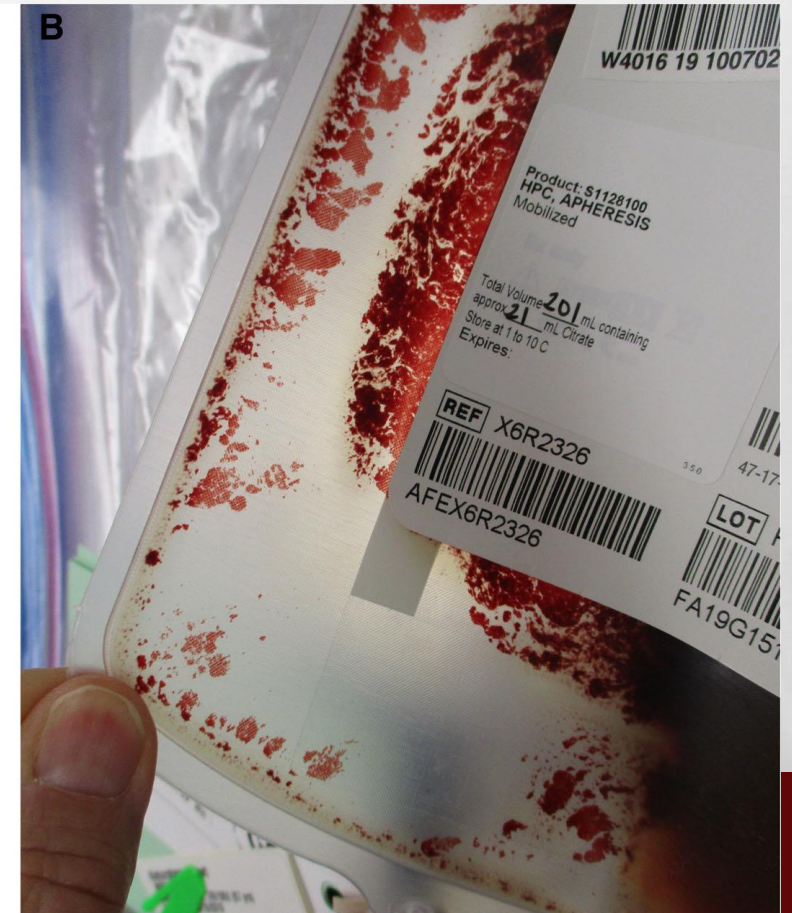
CLUMPING IN THE COLD

- **67-YEAR OLD MALE WITH DIFFUSE LARGE B-CELL LYMPHOMA (DLBCL)**
- **AUTOLOGOUS PERIPHERAL BLOOD STEM CELL (PBSC) COLLECTION WAS PERFORMED**
 - **60 ML/MIN FLOW RATE WITH TOTAL VOLUME OF 12.6 L PROCESSED**



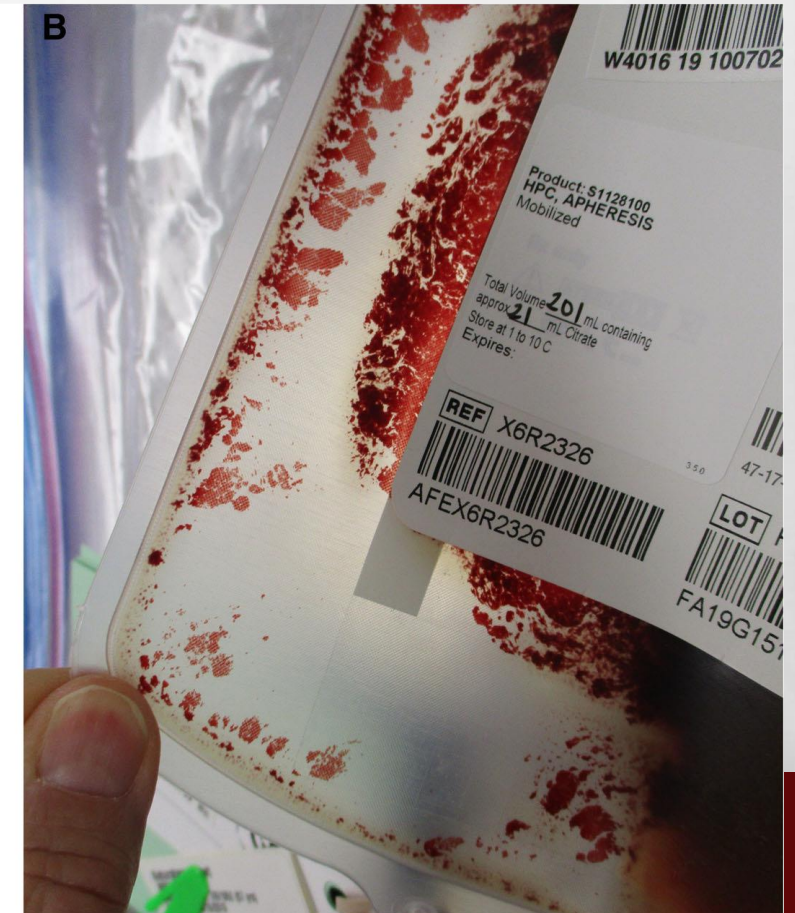
CLUMPING IN THE COLD

- **PBSC COLLECTION WAS UNREMARKABLE AND NO ABNORMALITIES OF PRODUCT WERE NOTED DURING OR AFTER COLLECTION**
- **FINAL PRODUCT WAS STORED AT 4°C FOR 20 MINUTES AWAITING PICKUP FROM LAB PERSONNEL**
- **UPON PICKUP, AGGLUTINATION WAS OBSERVED IN PRODUCT**



CLUMPING IN THE COLD

- **NO HISTORY OF COLD AGGLUTININ DISEASE OR HEMOLYTIC ANEMIA**
- **ANTIBODY SCREEN SHOWED 2+ PANAGGLUTINATION AT ROOM TEMP**
 - **NO AGGLUTINATION AT AHG**
- **DAT POSITIVE FOR C3 ONLY**
- **COLD AGGLUTININ SUSPECTED**



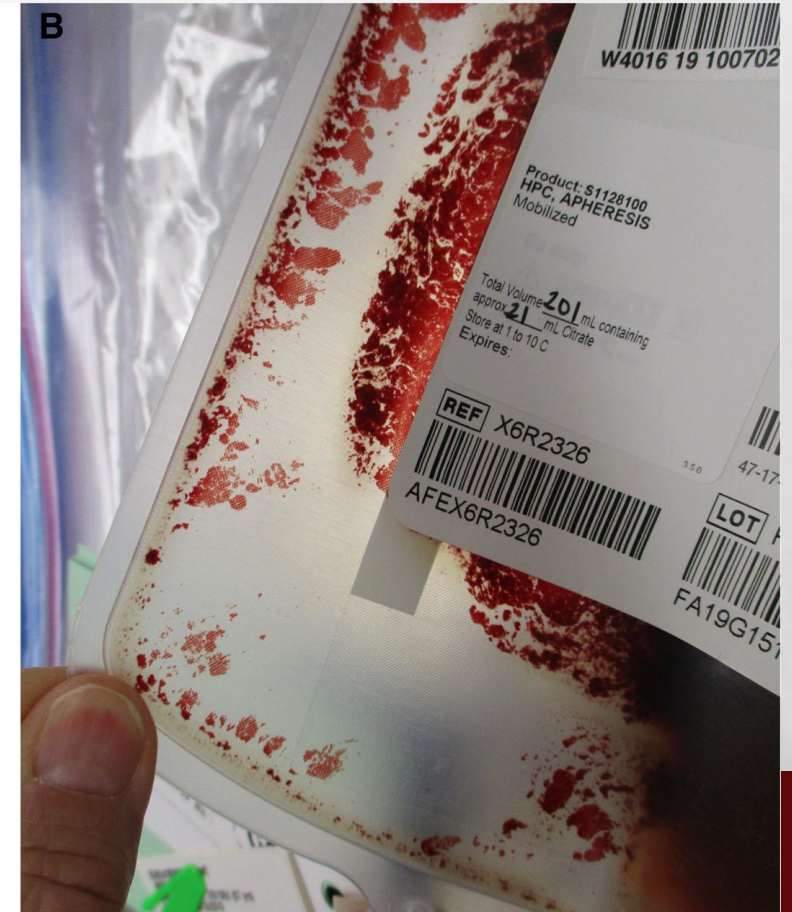
CLUMPING IN THE COLD

- **NEW BLOOD SAMPLE TESTED REVEALING:**
 - **COLD AGGLUTININS WITH HIGH TITERS**
 - **BROAD THERMAL AMPLITUDE**
 - **RELATIVE ANTI-I SPECIFICITY**
 - **WEAK RBC AUTOAGGLUTINATION AT 30°C**

	4°C	22°C	30°C	37°C
Group O RBCs (I)	256	8	1	0
Group O Cord RBCs (i)	2048	128	1	0
Patient's RBCs	256	8	1	0

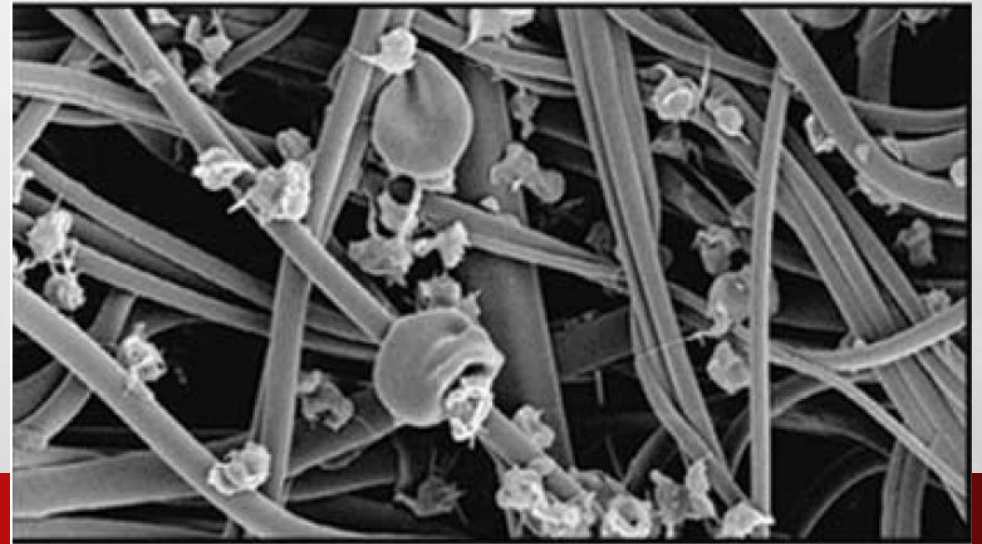
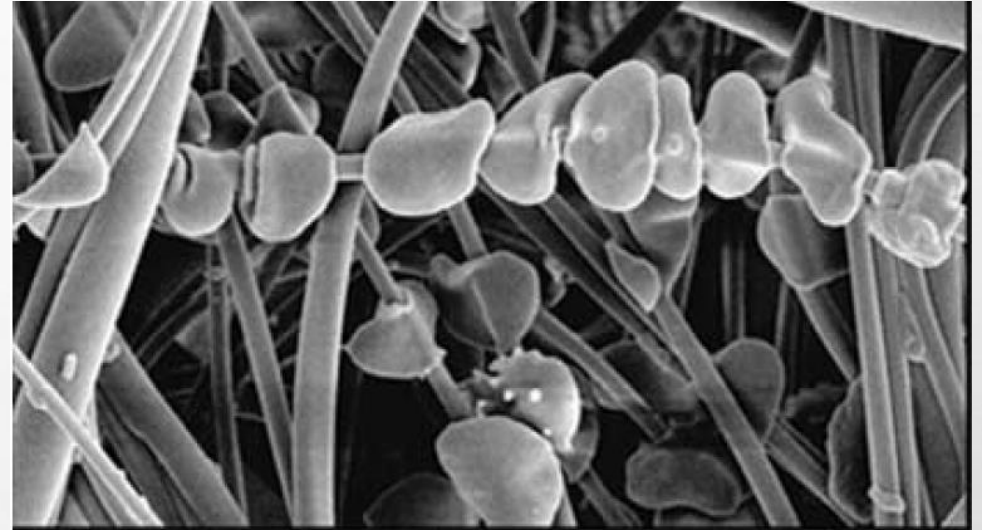
CLUMPING IN THE COLD

- **RBC CLUMPING CAN LEAD TO INSUFFICIENT PBSC QUALITY PRODUCTS**
- **APPROPRIATE PRECAUTIONS CAN BE TAKEN IF COLD AGGLUTININS ARE SUSPECTED AHEAD OF TIME**



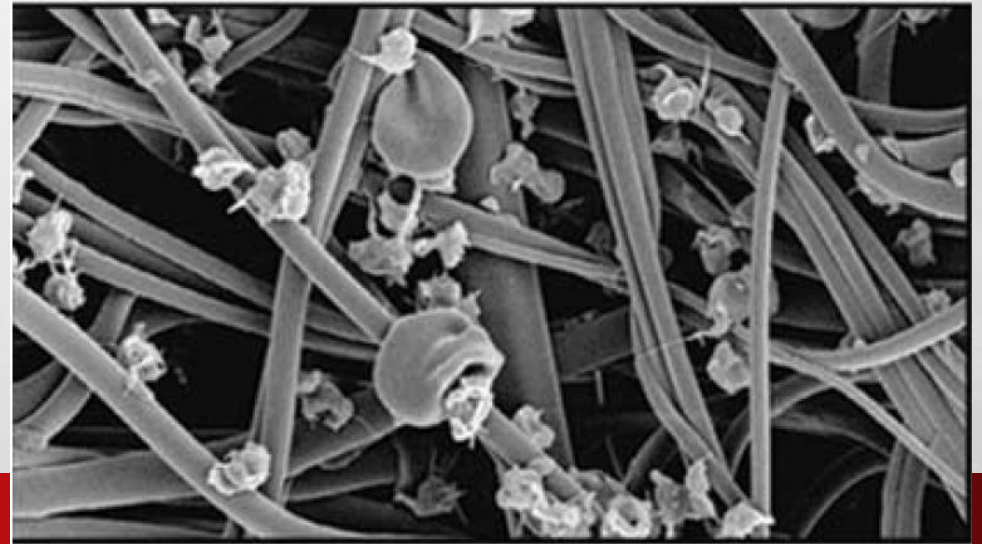
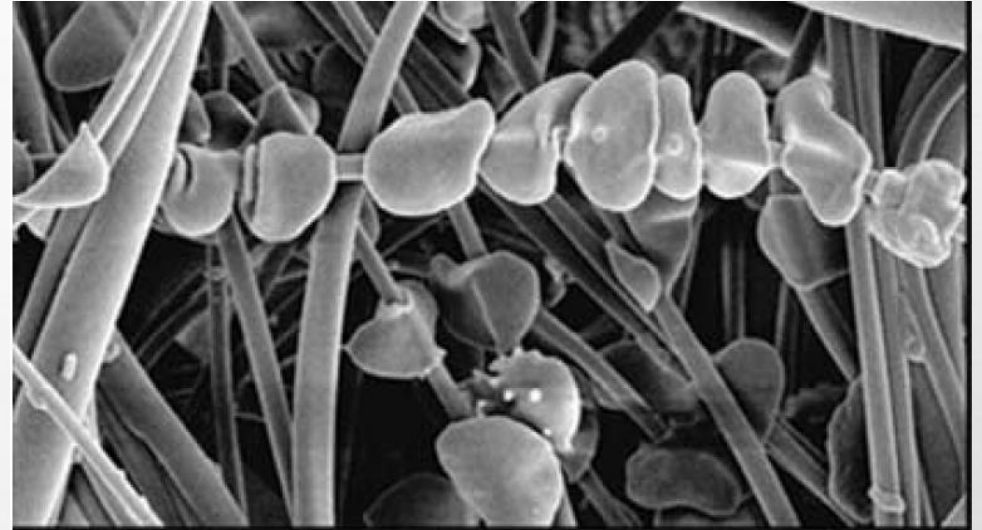
A STICKY SITUATION

- **GORLIN, J. & GUDINO, M. (2001) A STICKY SITUATION: WBC REDUCTION OF RBCS FROM DONORS WITH SICKLE CELL TRAIT. *TRANSFUSION*, 41, 1192.**



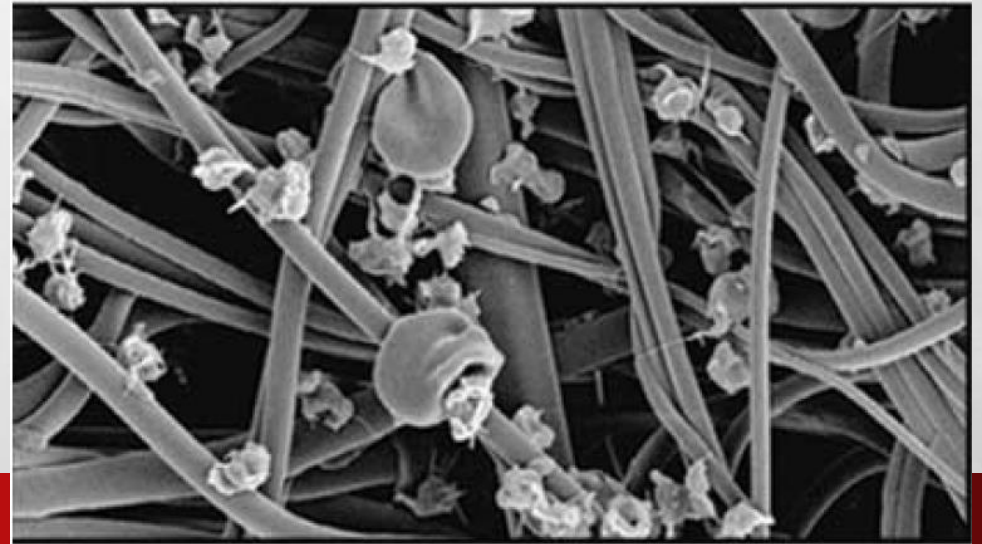
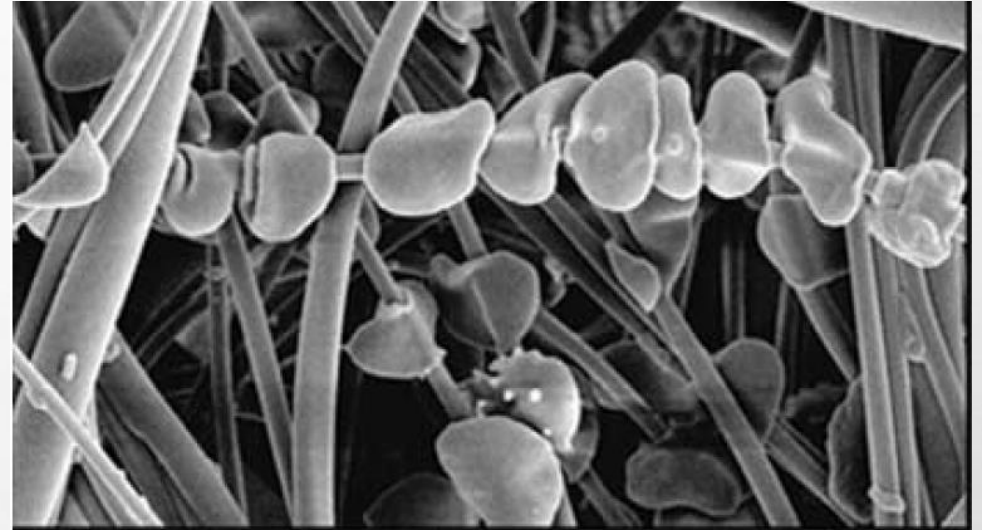
A STICKY SITUATION

- **HALF OF UNITS FROM DONORS WITH SICKLE CELL TRAIT FAIL TO COMPLETE WBC FILTRATION**
 - **APPARENT “CLOGGING” OF FILTER**
 - **UNIT IS DISCARDED**
- **THOSE UNITS THAT COMPLETE FILTRATION DO NOT MEET STANDARDS FOR WBC REDUCTION**



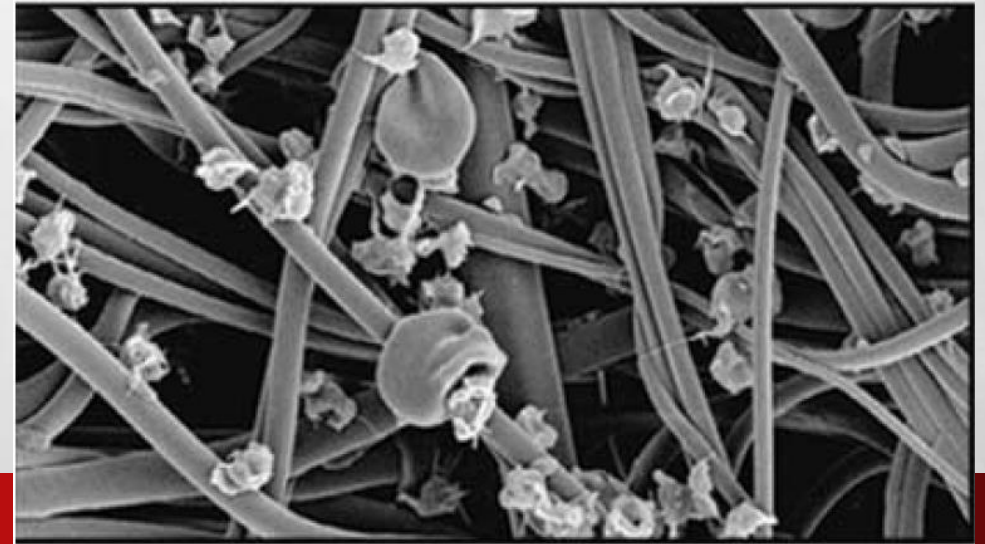
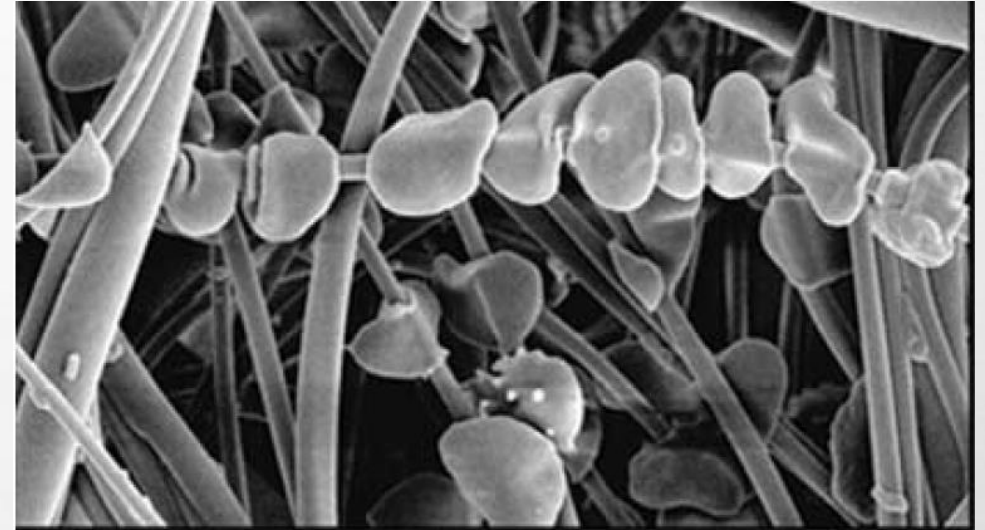
A STICKY SITUATION

- **TOP PICTURE – DONOR WITH SICKLE CELL TRAIT**
- **BOTTOM PICTURE – DONOR WITHOUT SICKLE CELL TRAIT (CONTROL)**
- **BOTH WHOLE BLOOD UNITS FILTERED WITHIN 2 HOURS OF COLLECTION**
 - **CONTROL UNIT COMPLETED FILTRATION IN 12 MINUTES**
 - **SICKLE CELL TRAIT UNIT CLOGGED FILTER IN 7 MINUTES WITH ONLY 15% RBC RECOVERY**



A STICKY SITUATION

- **BOTH FILTERS WASHED WITH BUFFER AND CHEMICALLY FIXED**
- **IMAGED AT 5KV ON FIELD EMISSION SCANNING ELECTRON MICROSCOPE**
- **EXTENSIVE RBC ADHESION TO WBC REDUCTION FIBERS IN SICKLE TRAIT UNIT COMPARED TO CONTROL UNIT**



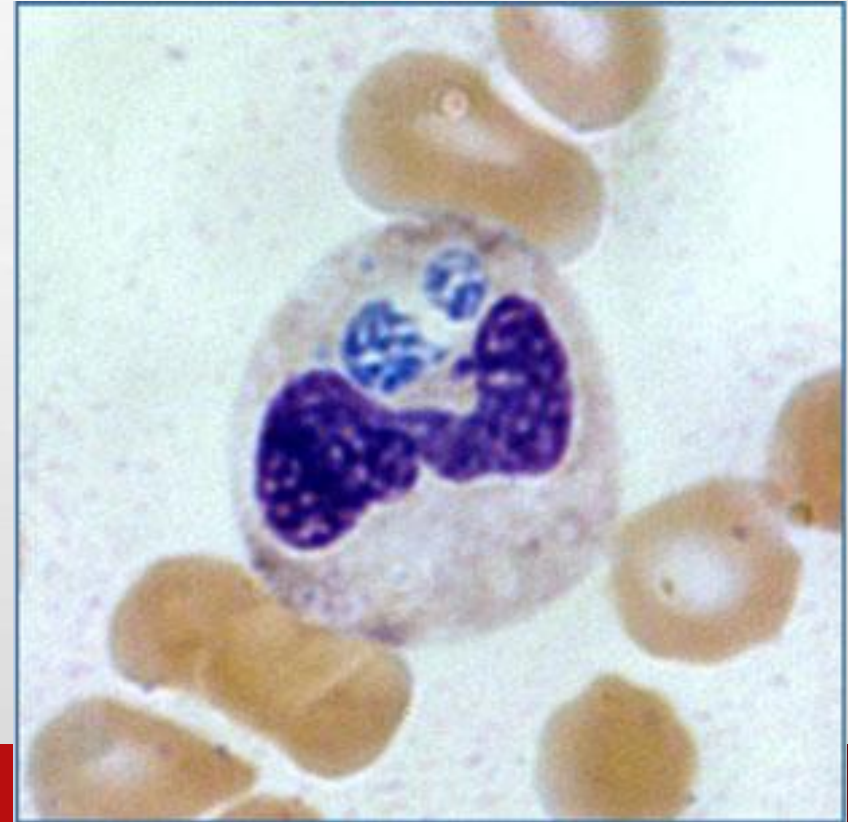
INTRACELLULAR ASSASSIN

- **GOEL, R. VASOVIC, L. WESTBLADE, L. HARRIS, R. D'AMBROSIA, D. EMMONS, F. LO, D. HSU, Y. JENKINS, S. & CUSHING, M. (2018) THE POWER OF A TRAINED EYE: FATAL TRANSFUSION-TRANSMITTED ANAPLASMOSIS AND A CASE STUDY IN INVESTIGATIVE WORK. *TRANSFUSION*, 59, 444-445.**



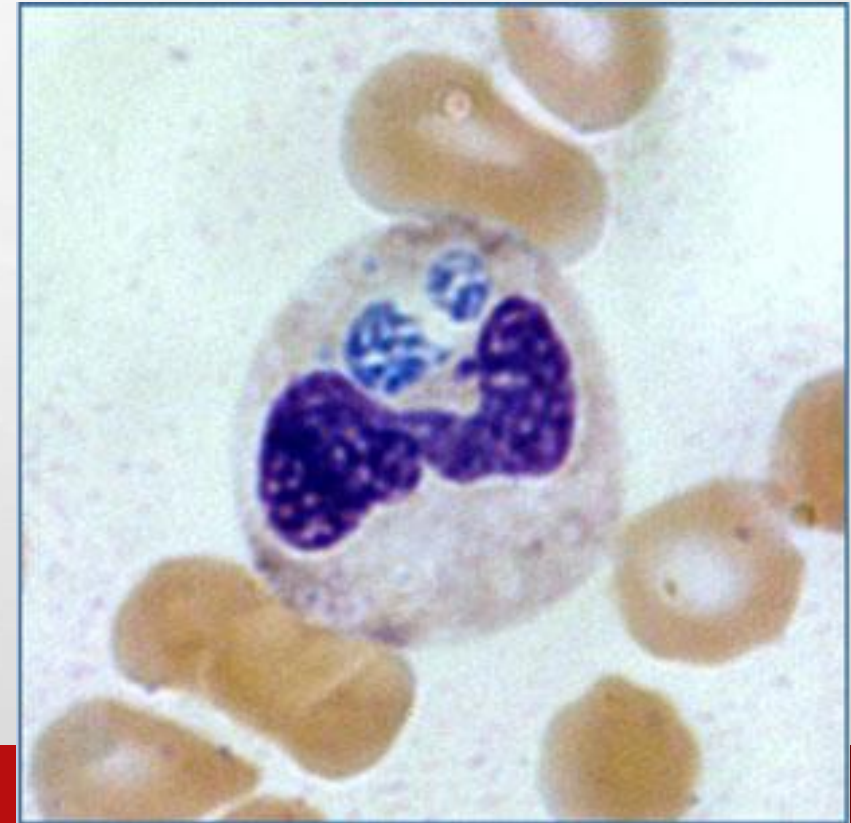
INTRACELLULAR ASSASSIN

- **78-YEAR OLD MALE WITH HISTORY OF CORONARY ARTERY DISEASE, DIABETES, AND CHRONIC KIDNEY DISEASE**
- **ADMITTED FOR ELECTIVE UROONCOLOGIC PROCEDURE**
- **PATIENT TRANSFUSED WITH MULTIPLE UNITS PREOPERATIVELY AND POSTOPERATIVELY FOR BLEEDING AND CORRECT SYMPTOMATIC ANEMIA**
- **SPIKED FEVER ON DAY +20 AND STARTED ON BROAD-SPECTRUM ANTIBIOTICS**



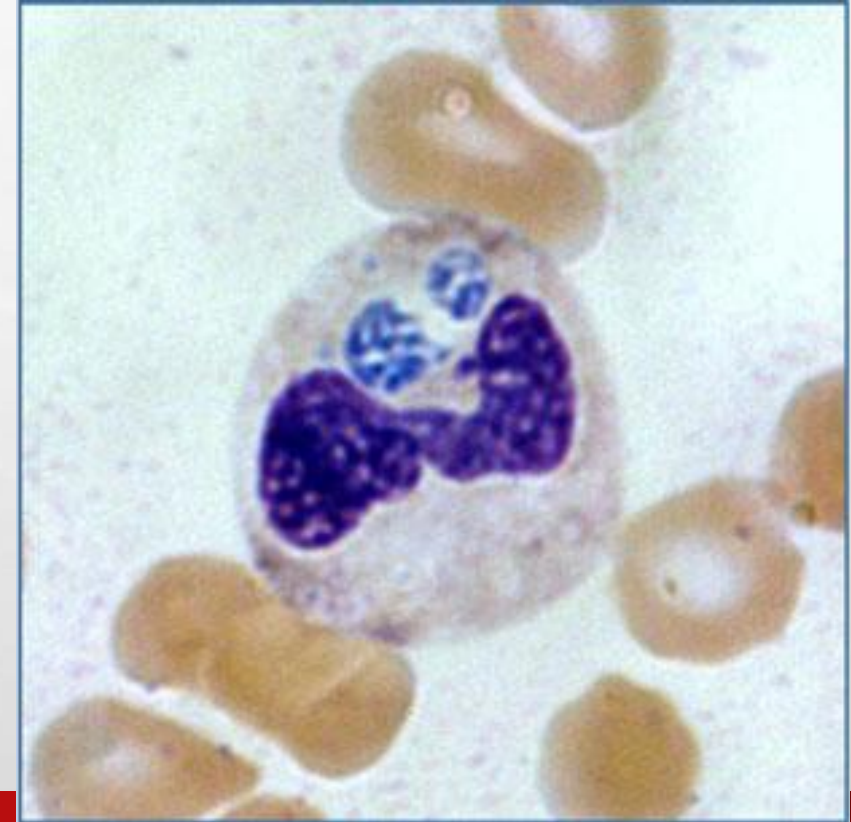
INTRACELLULAR ASSASSIN

- **BLOOD, URINE, THROAT CULTURES ALL NEGATIVE**
- **CONTINUED HIGH FEVER WITH HYPOXIA, HYPOTENSION, AND INCREASED TROPONIN**
- **TRANSFERRED TO ICU FOR CULTURE-NEGATIVE SEPSIS**
- **DAY 4 OF FEVER: HEMATOLOGY TECH OBSERVED GRANULOCYTIC INCLUSIONS ON PERIPHERAL SMEAR**
 - **CONSISTENT WITH *ANAPLASMA PHAGOCYTOPHILUM* MORULAE**



INTRACELLULAR ASSASSIN

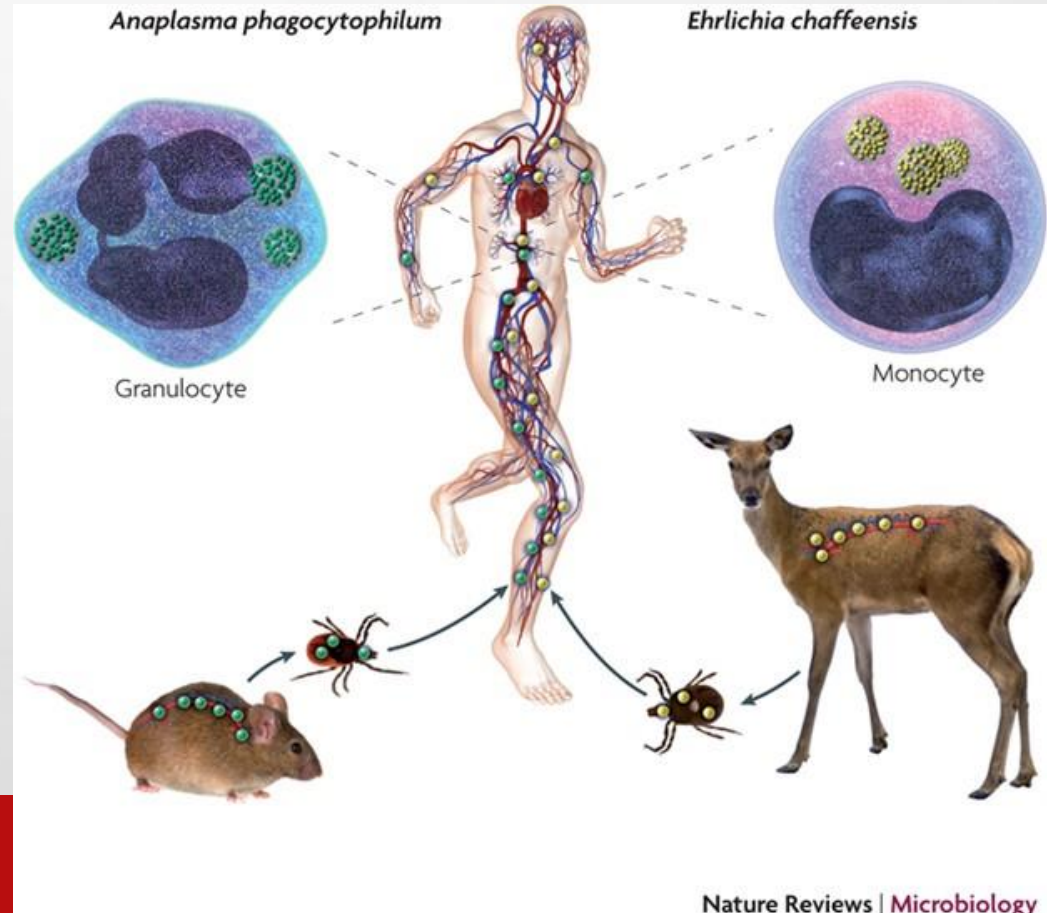
- ID CONSULTANT INITIATED DOXYCYCLINE IMMEDIATELY
 - PATIENT DIED OF SEPTIC SHOCK ON DAY +25
- BLOOD BANK REVEALED SEGMENT FROM LEUKOREduced RBC UNIT TRANSFUSED ON DAY +11 WAS POSITIVE FOR *A. PHAGOCYTOPHILUM*
- PATIENT'S SPECIMENS BEGINNING DAY +7 AFTER TRANSFUSION WERE ALSO POSITIVE
 - MORULAE FIRST SEEN ON PERIPHERAL SMEAR DAY +12 POST-TRANSFUSION



INTRACELLULAR ASSASSIN

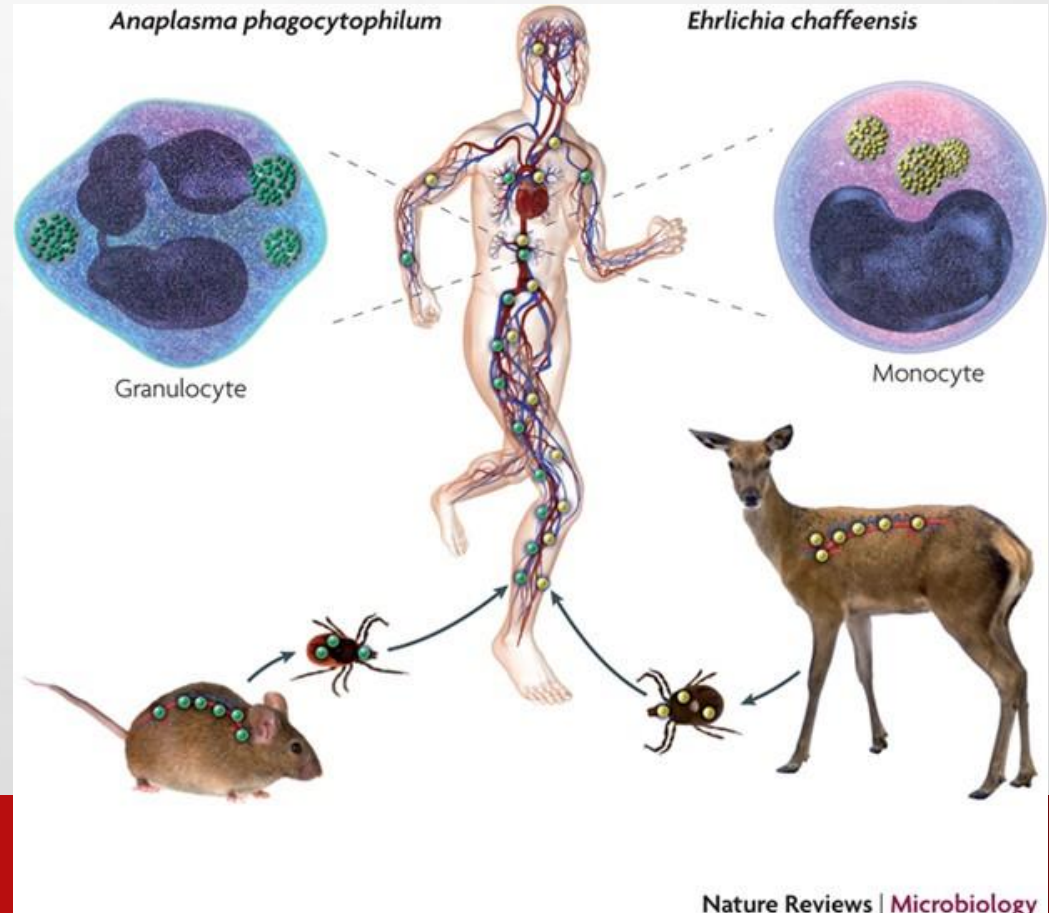
- ***A. PHAGOCYTOPHILUM***

- **OBLIGATE INTRACELLULAR BACTERIUM**
- **WILD MICE ARE HOSTS WHICH TRANSMIT TO HUMANS VIA TICK BITES**
- **LACK GENES FOR BIOSYNTHESIS OF THE LIPOPOLYSACCHARIDES/PEPTIDOGLYCANS THAT ACTIVATE HOST LEUKOCYTES**
- **ACQUIRE HOST CHOLESTEROL FROM THE LOW-DENSITY-LIPOPROTEIN UPTAKE PATHWAY**



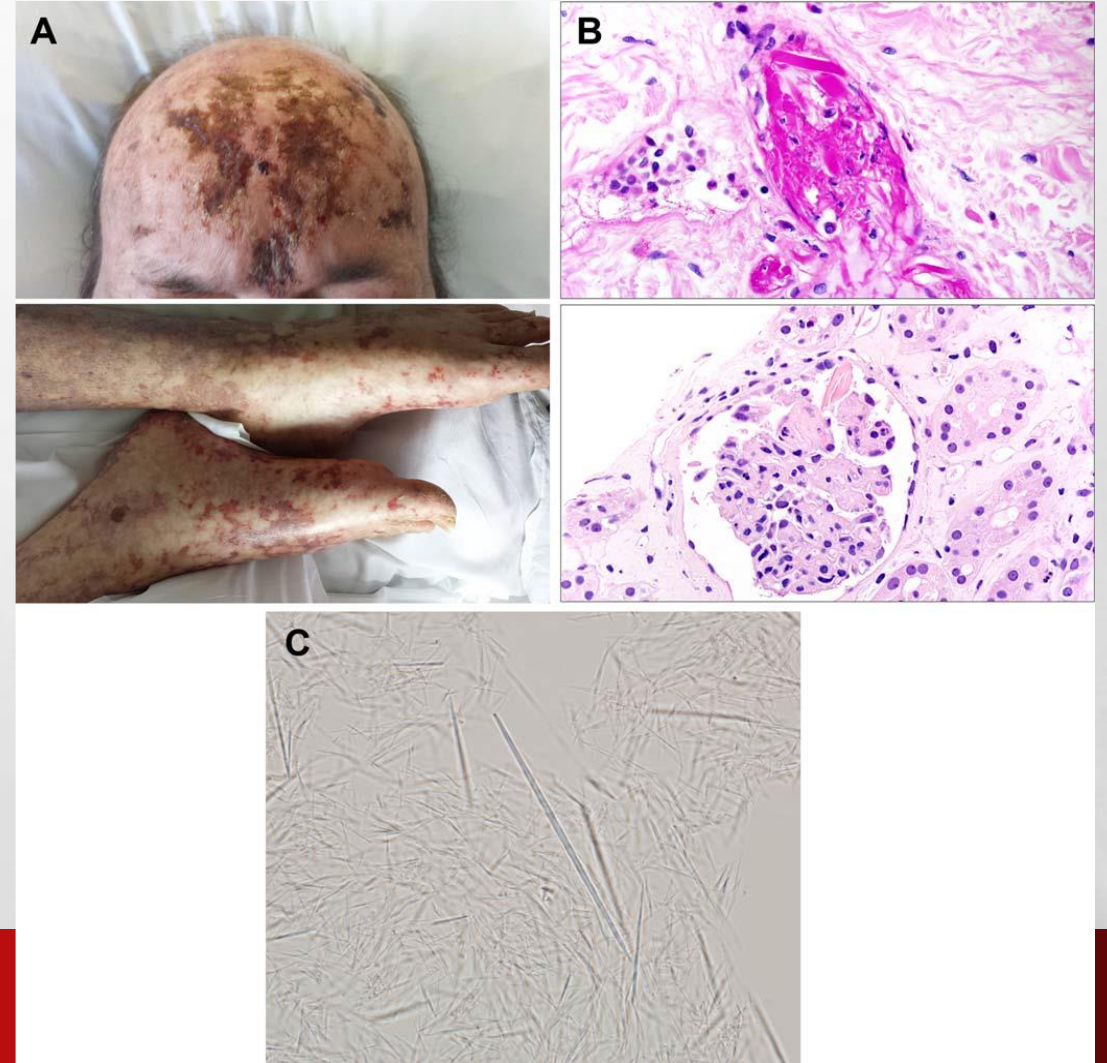
INTRACELLULAR ASSASSIN

- ***A. PHAGOCYTOPHILUM***
 - CAVEOLAE-MEDIATED ENDOCYTOSIS DIRECTS BACTERIA TO AN INTRACELLULAR COMPARTMENT, OR INCLUSION
 - INCLUSION DOES NOT ACQUIRE COMPONENTS OF NADPH OXIDASE NOR SHOW SIMILARITY TO LATE ENDOSOMES OR LYSOSOMES
 - *A. PHAGOCYTOPHILUM* INCLUSIONS ACQUIRE EARLY-AUTOPHAGOSOME CHARACTERISTICS



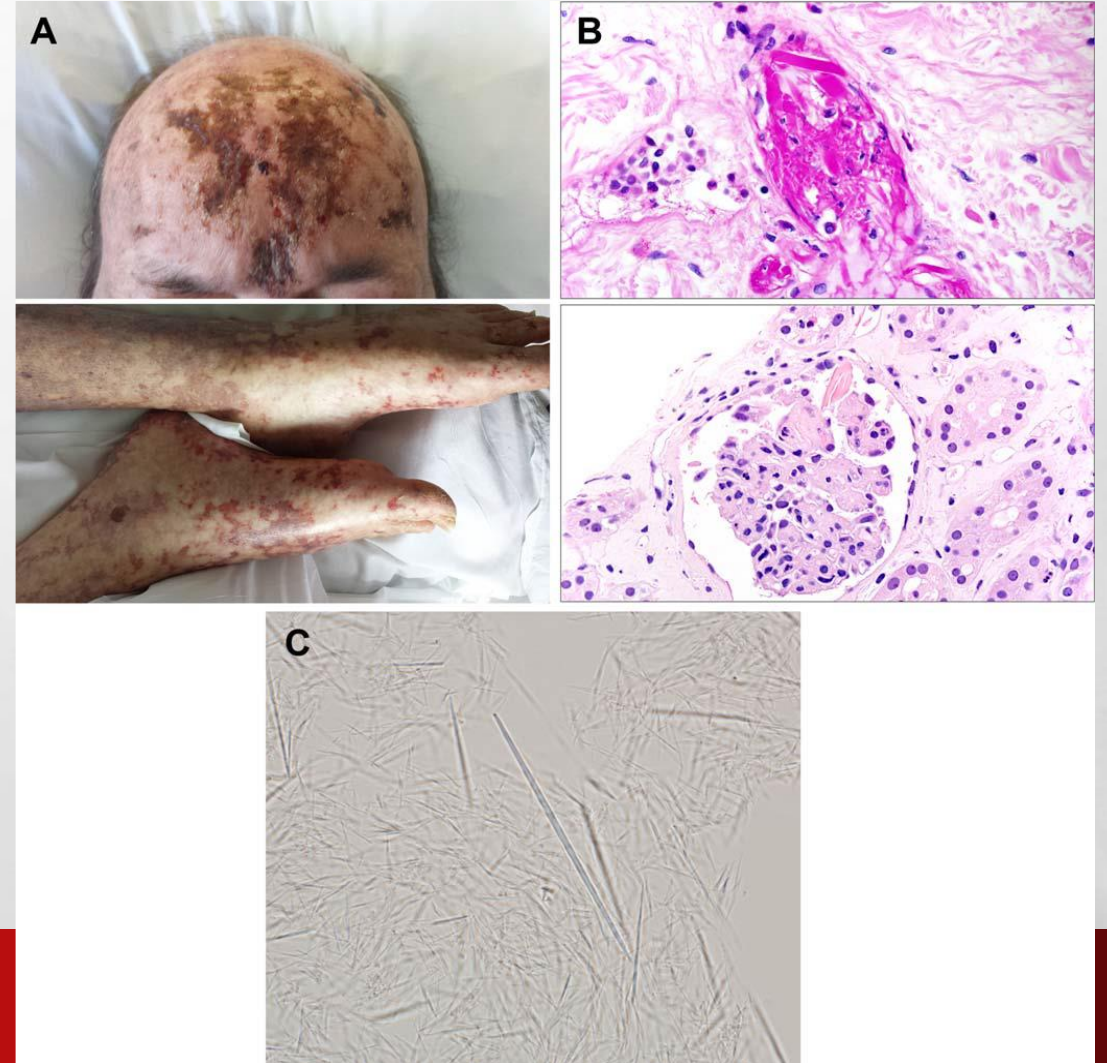
CRYSTAL DEATH

- **AVEDSCHMIDT, S. FARKASH, E. & YAMADA, C. (2016) A CASE OF CRYOCRYSTALGLOBULINEMIA. *TRANSFUSION*, 56, 1678-1679.**



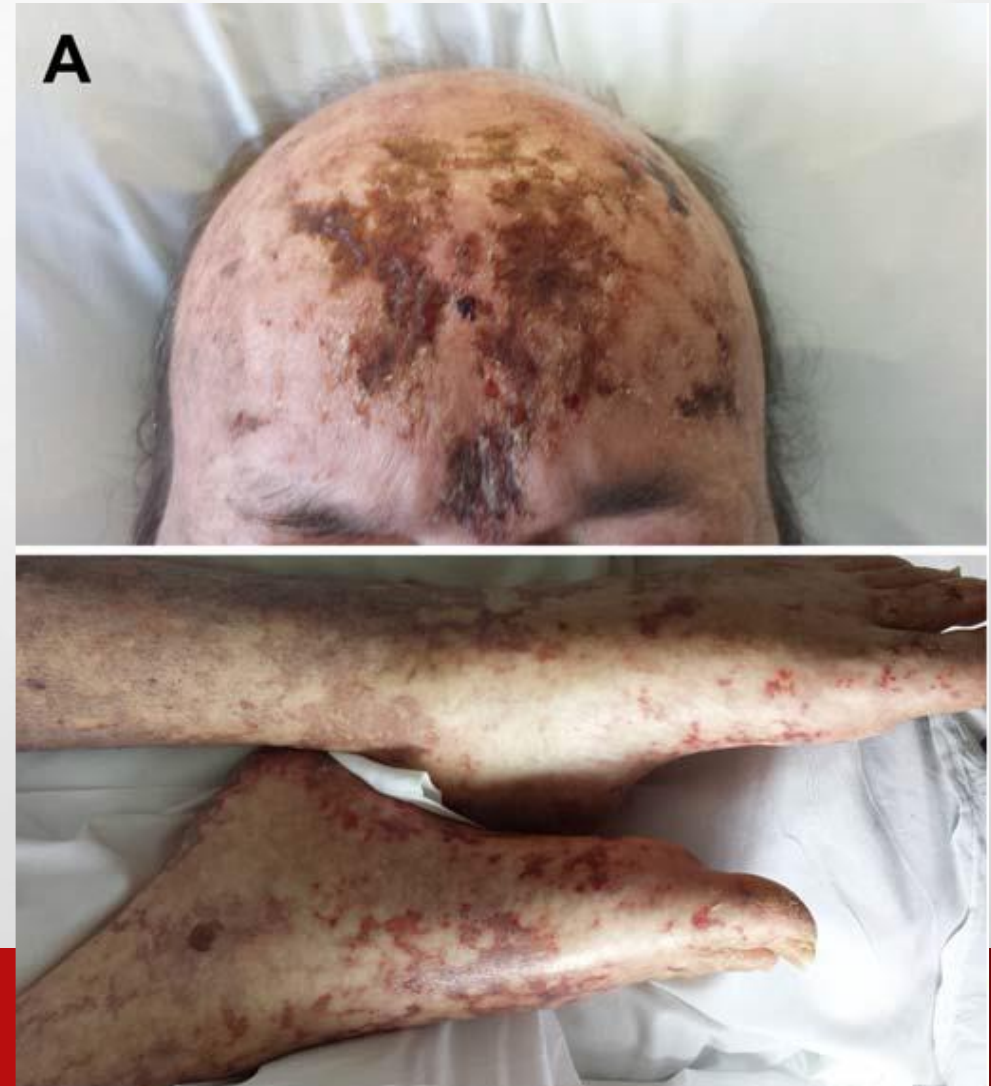
CRYSTAL DEATH

- **52-YEAR OLD MALE WITH MULTIPLE HOSPITALIZATIONS DUE TO:**
 - **COPD**
 - **RESPIRATORY/RENAL FAILURE**
 - **DEEP VEIN THROMBOSIS**
 - **BASAL GANGLIA HEMORRHAGE**
 - **SEPSIS**
- **IGGK MONOCLONAL GAMMOPATHY**
 - **SERUM PROTEIN ELECTROPHORESIS – 0.1G/DL M-PROTEIN**



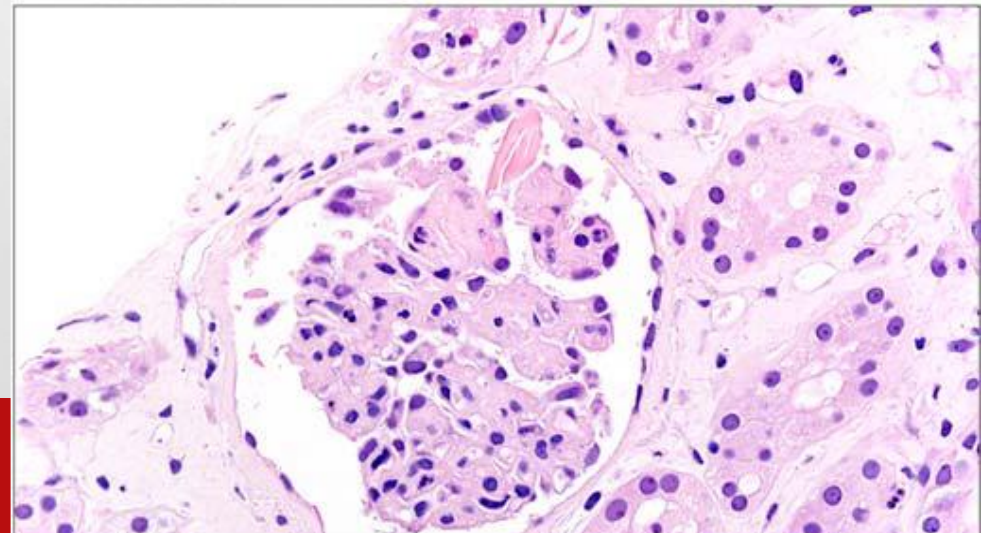
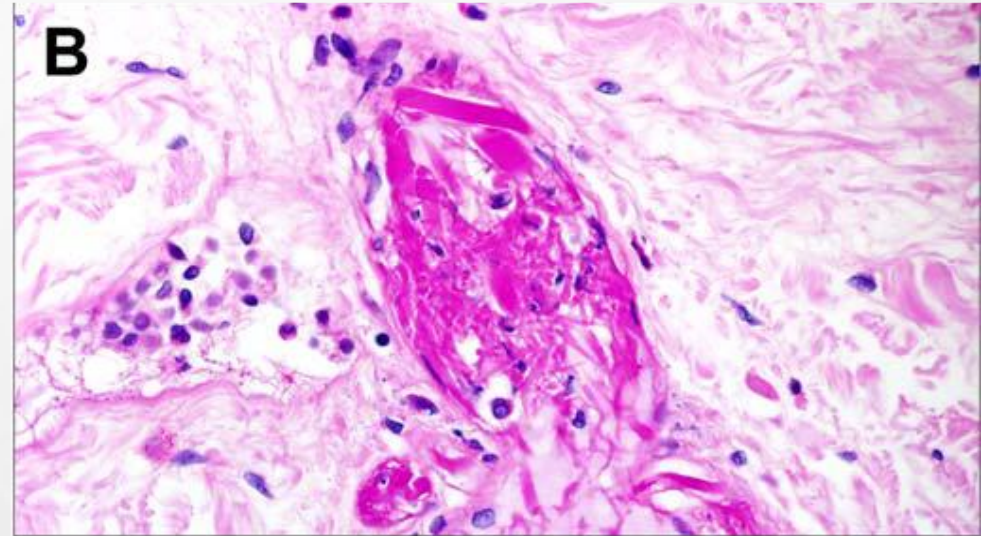
CRYSTAL DEATH

- **7 MONTHS LATER PRESENTED WITH DIFFUSE PURPURIC RASH, RENAL FAILURE, MILD THROMBOCYTOPENIA**
 - **IAT AND DAT WERE NEGATIVE**
 - **HEMOLYSIS LABS WERE NEGATIVE**
 - **BM BIOPSY WAS UNREMARKABLE**
 - **ADAMTS13 WAS 53% (NORMAL IS >68%)**



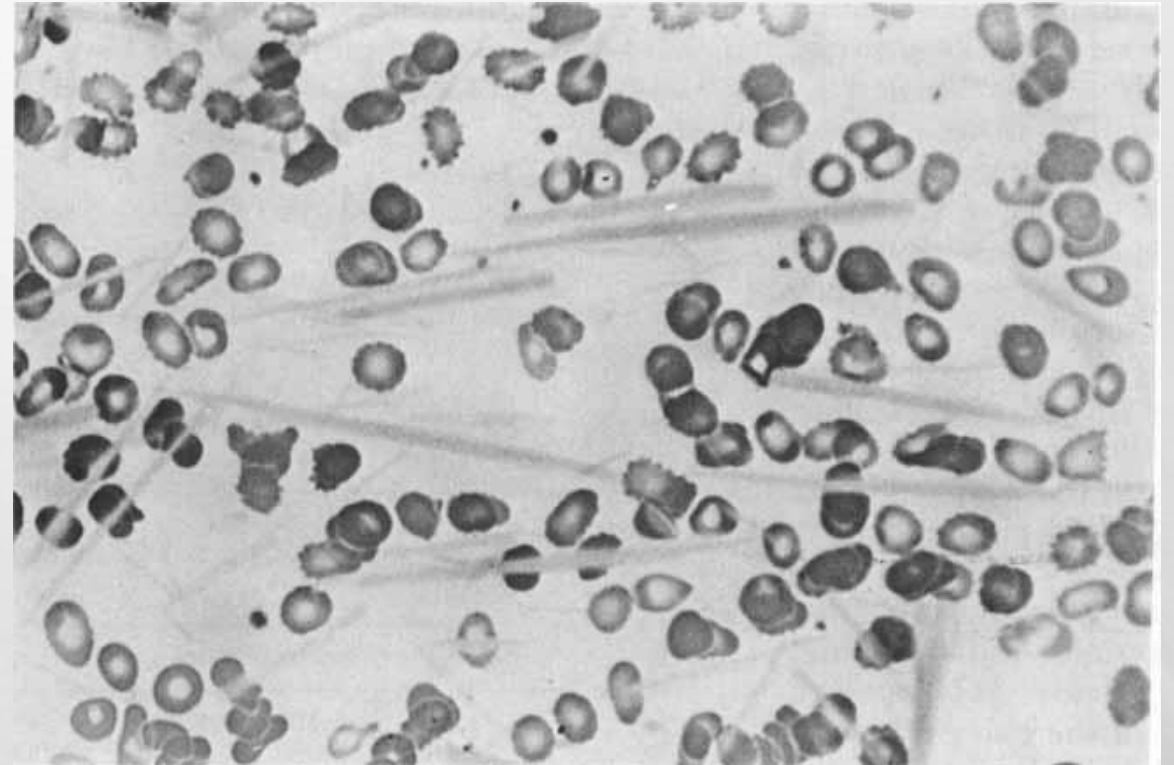
CRYSTAL DEATH

- **SKIN BIOPSY SHOWED NEUTOPHILIC THROMBOTIC VASCULOPATHY WITH CRYSTALS (TOP)**
- **KIDNEY BIOPSY SHOWED CORTICAL NECROSIS, FIBROCELLULAR CRESCENTS, ARTERIAL THROMBI, EXTENSIVE TUBULAR INJURY WITH OCCASIONAL TUBULAR/GLOMERULAR CRYSTALS (BOTTOM)**
- **CRYOCRIT WAS 12% (NORMAL <1%)**
 - **DIAGNOSED WITH TYPE I CRYOCRYSTALGLOBULINEMIA**



CRYSTAL DEATH

- **CRYOCRYSTALGLOBULINEMIA**
 - **RARE TYPE OF CRYOGLOBULINEMIA**
 - **RESULTS IN MICROVASCULAR LESIONS**
 - **CRYSTALLIZATION OF MONOCLONAL IMMUNOGLOBULINS CAUSES MORE SEVERE MANIFESTATIONS**
- **TREATMENT BY THERAPEUTIC PLASMA EXCHANGE (TPE) – CATEGORY I APPLICATION FROM ASA**



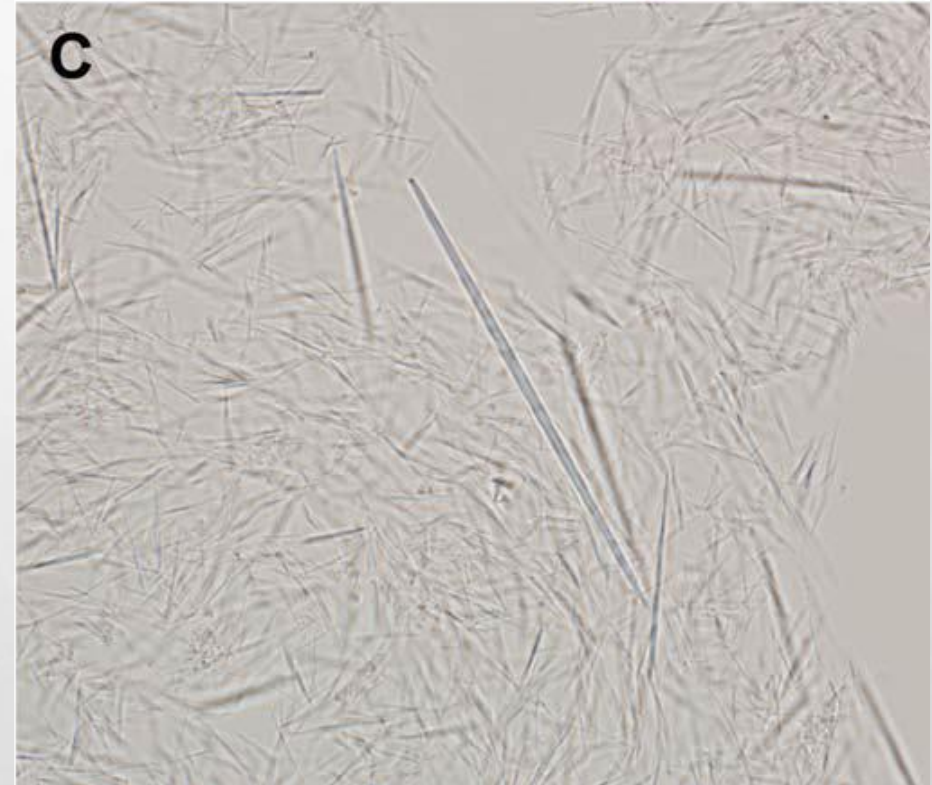
CRYSTAL DEATH

- **PATIENT UNDERWENT 8 TPE PROCEDURES WITH STEROID, CYCLOPHOSPHAMIDE AND BORTEZOMIB THERAPY FOR 1 MONTH**
 - **CRYOGLOBULIN LEVEL UNDETECTABLE**
 - **RENAL FUNCTION IMPROVED**
- **UNFORTUNATELY DEVELOPED GI BLEEDING FROM CRYOGLOBULINEMIC VASCULOPATHY**
 - **RELAPSED WITH CRYOCRIT OF 42%, RENAL/RESPIRATORY DYSFUNCTION, MENTAL STATUS CHANGE**



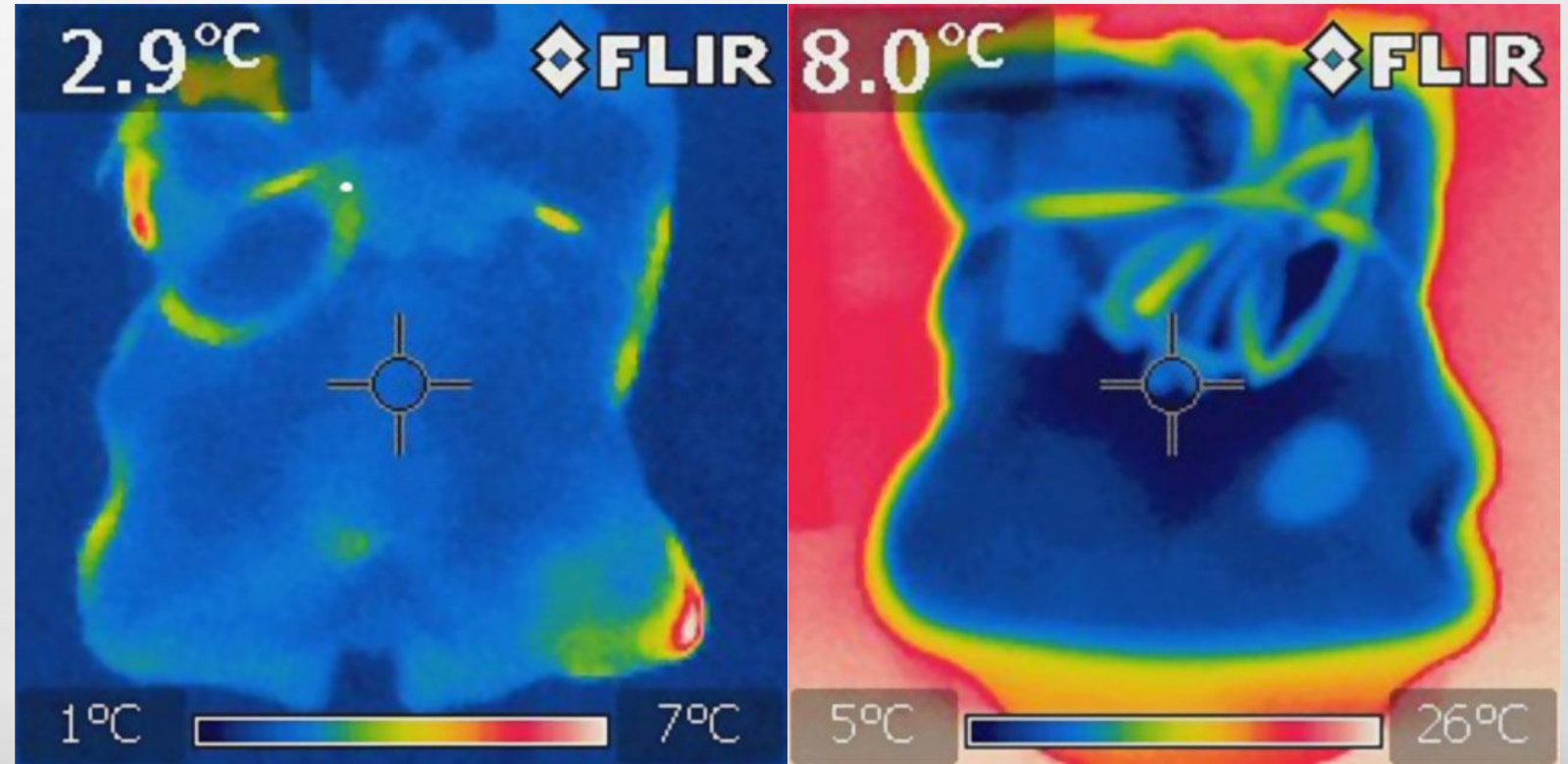
CRYSTAL DEATH

- **CRYOPRECIPITATE FROM CRYOCRIT SHOWED SPINDLED CRYSTALS**
- **PATIENT UNDERWENT 11 MORE TPE PROCEDURES WITH MEDICATION THERAPY AND DIALYSIS OVER 1 MONTH**
 - **CRYOCRIT DECREASED TO 5%**
 - **MENTAL STATUS IMPROVED**
 - **ORGAN FUNCTION DID NOT IMPROVE**
- **PATIENT WITHDREW FROM FURTHER TREATMENTS AND EXPIRED 12 MONTHS AFTER INITIAL DIAGNOSIS**



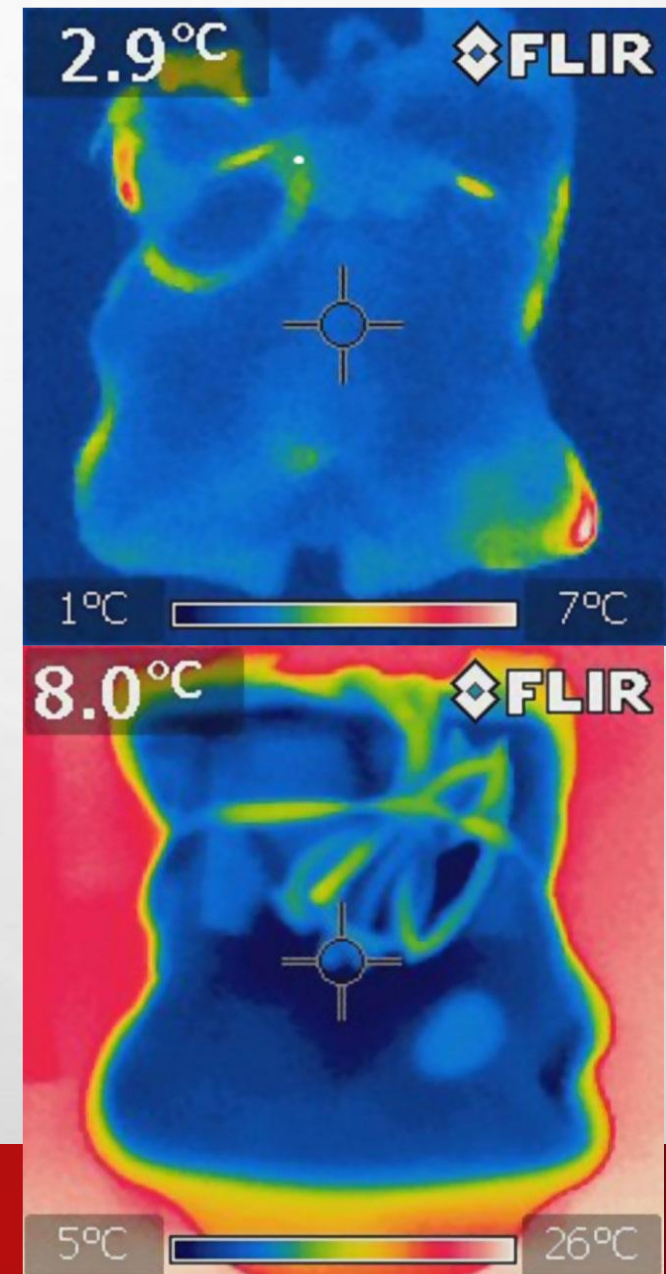
THE 30-MINUTE RULE

- **BOOTH, G. SPAIN, T, & GEHRIE, E. (2014) THERMAL IMAGING OF A RED BLOOD CELL UNIT DURING STORAGE AND AT TIME OF ISSUE. *TRANSFUSION*, 54, 2803.**



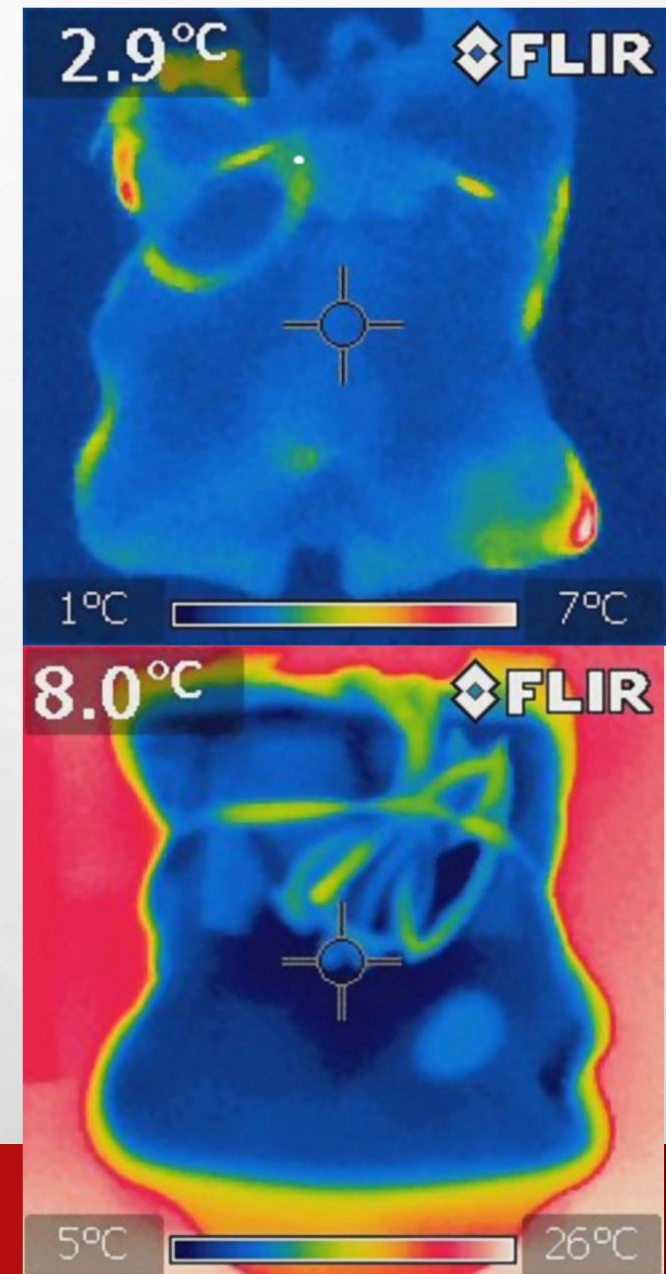
THE 30-MINUTE RULE

- **THERMAL IMAGES OF RED BLOOD CELL PRODUCTS**
 - RBC UNIT TEMPERATURE DURING STORAGE IN TEMPERATURE-CONTROLLED WALK-IN REFRIGERATOR (TOP)
 - RBC UNIT TEMPERATURE AT TIME OF ISSUE BEFORE RELEASE AND AFTER ELECTRONIC CROSSMATCH (BOTTOM)
- **ELECTRONIC CROSSMATCH ONLY REQUIRED 10 MINUTES**
 - EFFECT OF EXPOSURE TO ROOM TEMPERATURE ON RBC UNIT
 - DEMONSTRATES LOGISTICAL CHALLENGE ADHERING TO FDA/AABB GUIDELINES



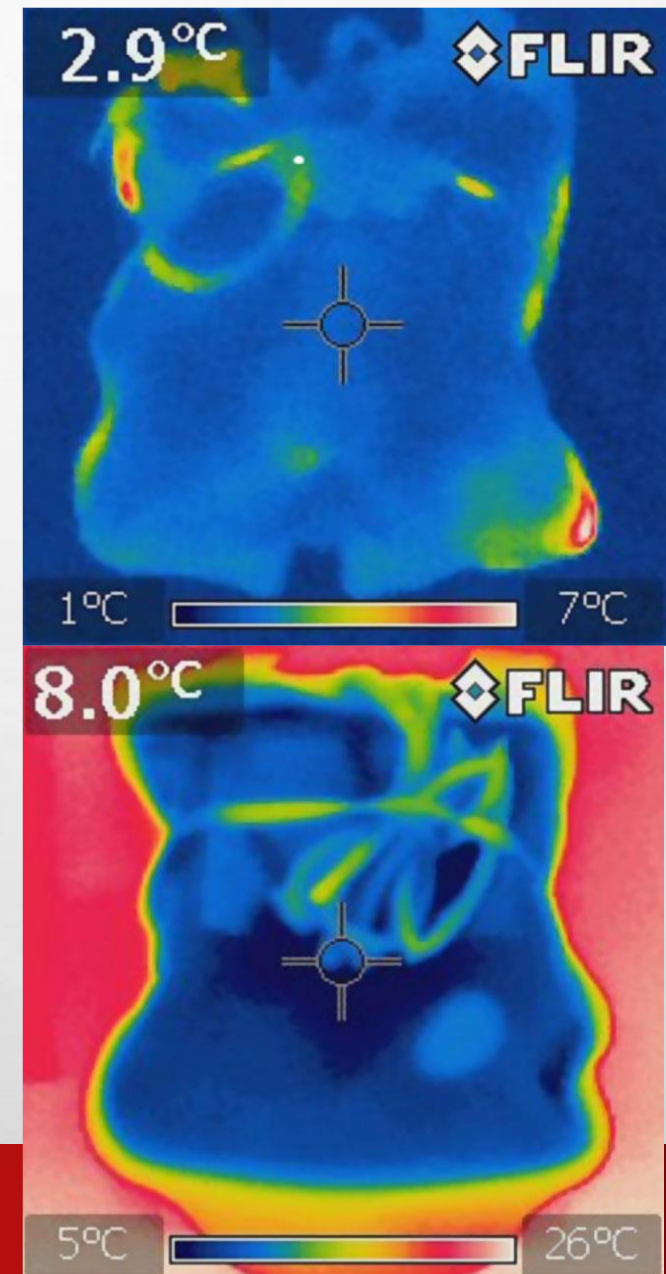
THE 30-MINUTE RULE

- **“30-MINUTE RULE”**
 - **PERHAPS DERIVED FROM ARMY IN EARLY 20TH CENTURY**
 - **OLD WIVES TALE**
- **MUST VALIDATE PROCESS FOR TRANSPORT OF PRODUCTS AND ACCEPTABILITY BACK INTO INVENTORY**
 - **STORAGE VS. TRANSPORT**



THE 30-MINUTE RULE

- **FDA TITLE 21, SECTION 640**
 - **LAW OF THE LAND**
- **AABB 32ND EDITION OF BB/TS STANDARDS**
 - **5.1.8A – TABLE LISTING ACCEPTABLE STORAGE AND TRANSPORT TEMPERATURES FOR BLOOD AND BLOOD COMPONENTS**
 - **5.6.5 – TRANSPORT FROM COLLECTION SITE TO PROCESSING SITE**
 - **5.26 – REISSUE OF BLOOD AND BLOOD COMPONENTS**



THROMBUS AMONG US

- **JACOB, E. BURGSTALER, E. WINTERS, J. STUBBS, J. & GANDHI, M. (2009) THROMBUS FORMATION DURING PLATELET DONATION. *TRANSFUSION*, 49, 2021.**



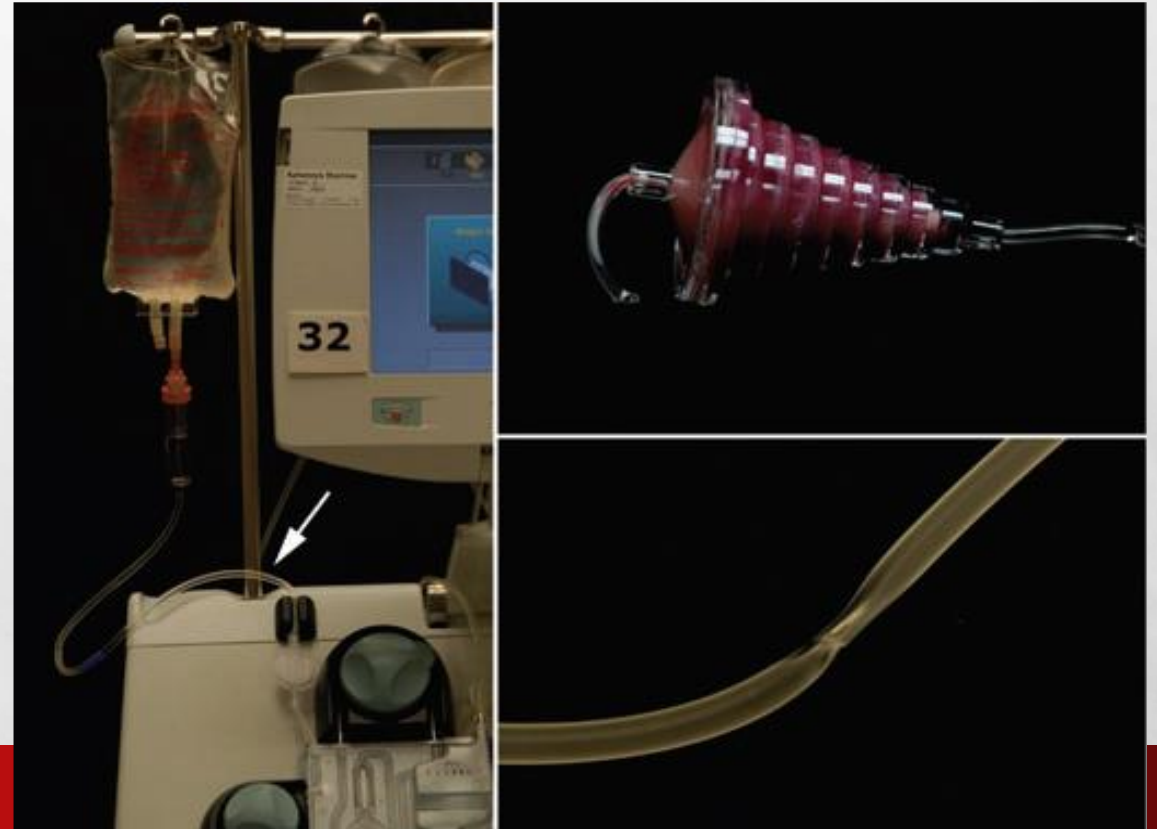
THROMBUS AMONG US

- **AUTOMATED PLATELET DONATION USING COBE SPECTRA**
- **ALARM SOUNDED INDICATING LOW PLATELET COUNT IN COLLECTION LINE**
- **DONATION STOPPED AND INVESTIGATION INITIATED**



THROMBUS AMONG US

- **COMPRESSION NOTED IN ANTICOAGULANT TUBING LINE (BOTTOM RIGHT)**
 - **LOCATION INDICATED IN LEFT PICTURE**
- **COMPRESSION RESULTED IN INADEQUATE ANTICOAGULATION**
 - **THROMBI FORMED IN LEUKOREDUCTION CHAMBER (TOP RIGHT)**



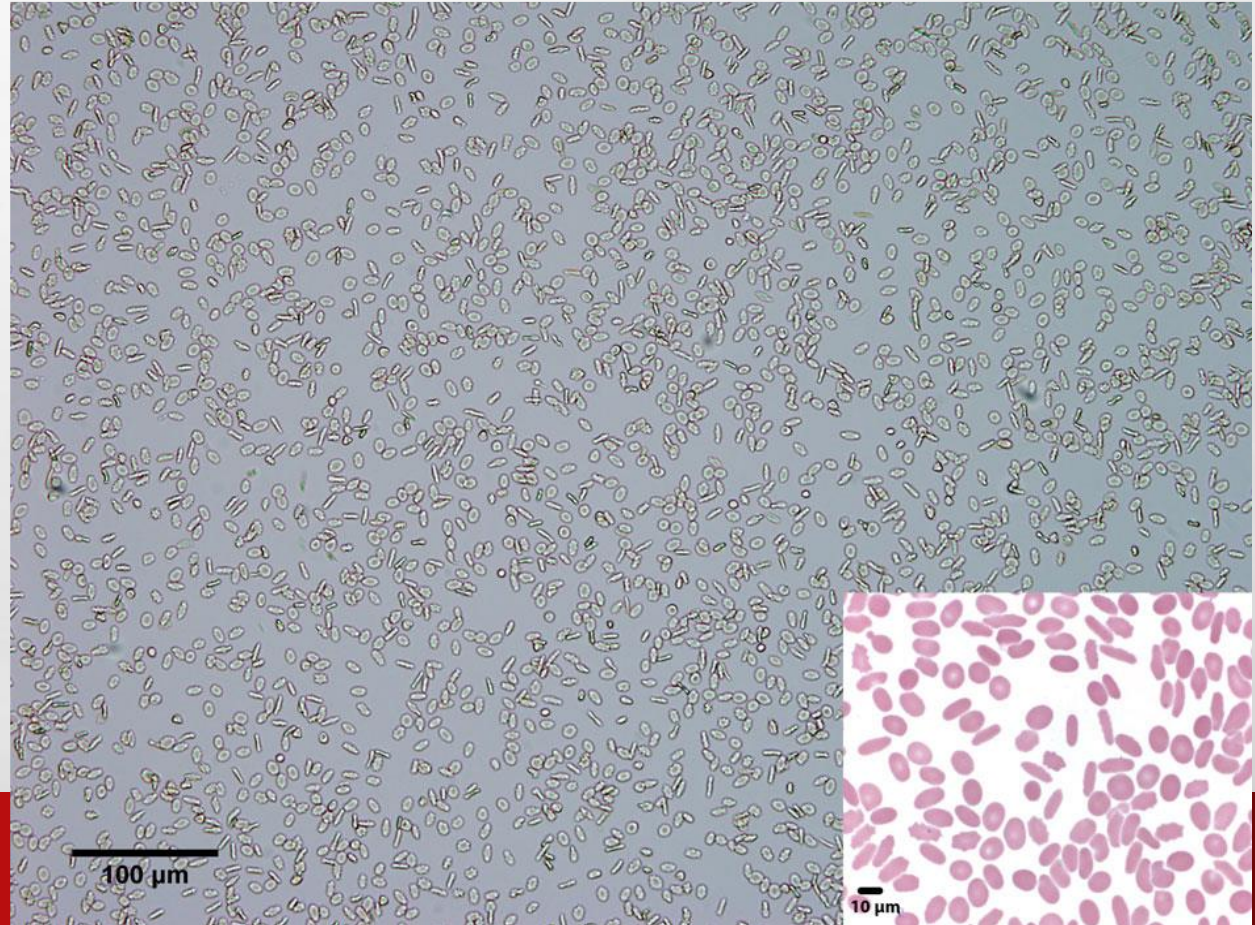
THROMBUS AMONG US

- **SENSOR FROM LEUKOREDUCTION CHAMBER TRIGGERED**
- **THROMBI FORMED IN COLLECTION BAG**
- **NO THROMBI NOTED AT RETURN SITE**
 - **DONOR SUFFERED NO COMPLICATIONS**
- **COMPRESSIONS SHOULD BE NOTED AND CORRECTED USING DISPOSABLES DURING BLOOD COLLECTION**



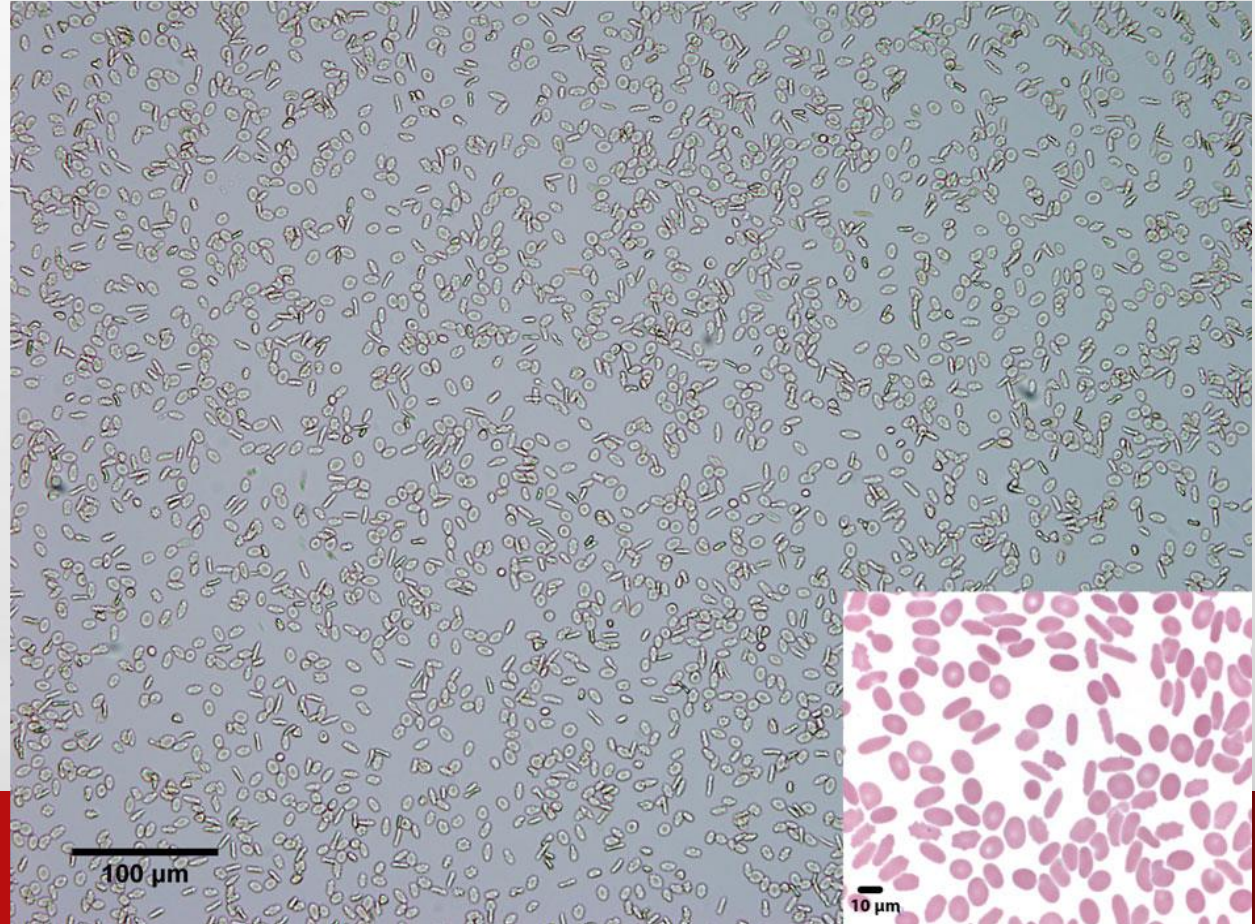
ELLIPTICAL TRAINING

- **DESIMONE, R. ONG, S. CROWLEY, K. VASOVIC, L. SENALDI, E. KESSLER, D. GOEL, R. & HSU, Y. (2018) HEREDITARY ELLIPTOCYTOSIS OF DONOR RED BLOOD CELL UNIT DETECTED DURING COOMBS CROSSMATCH. *TRANSFUSION*, 59, 446-447.**



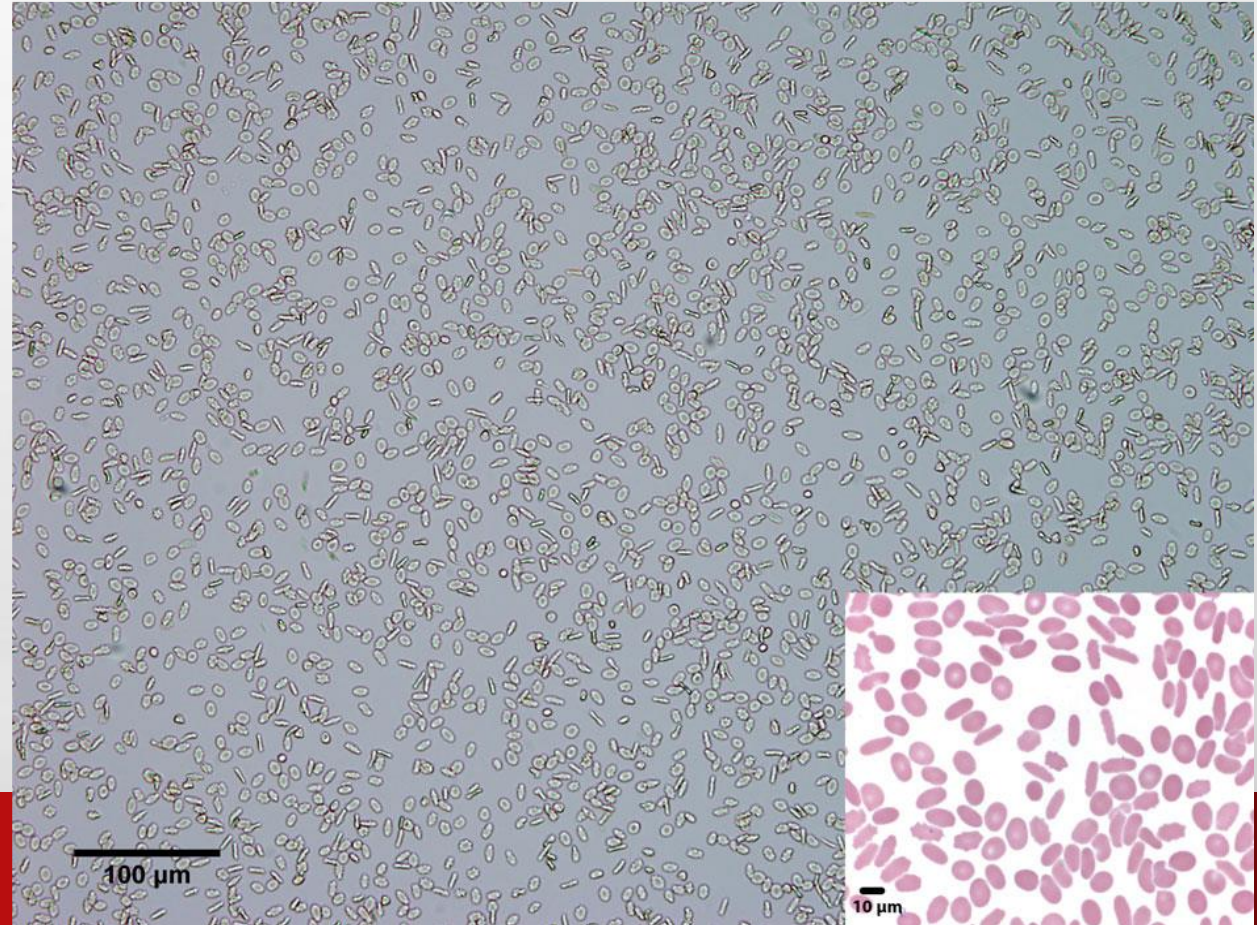
ELLIPTICAL TRAINING

- **RBC TRANSFUSION ORDERED FOR PATIENT**
 - **NEGATIVE IAT**
 - **HISTORY OF RBC ALLOANTIBODIES**
- **FULL CROSSMATCH WAS NEGATIVE**
 - **MICROSCOPIC EXAM SHOWED NUMEROUS PENCIL-SHAPED RBCS**
- **PERIPHERAL SMEAR FROM SEGMENT REVEALED 50% ELLIPTOCYTES**



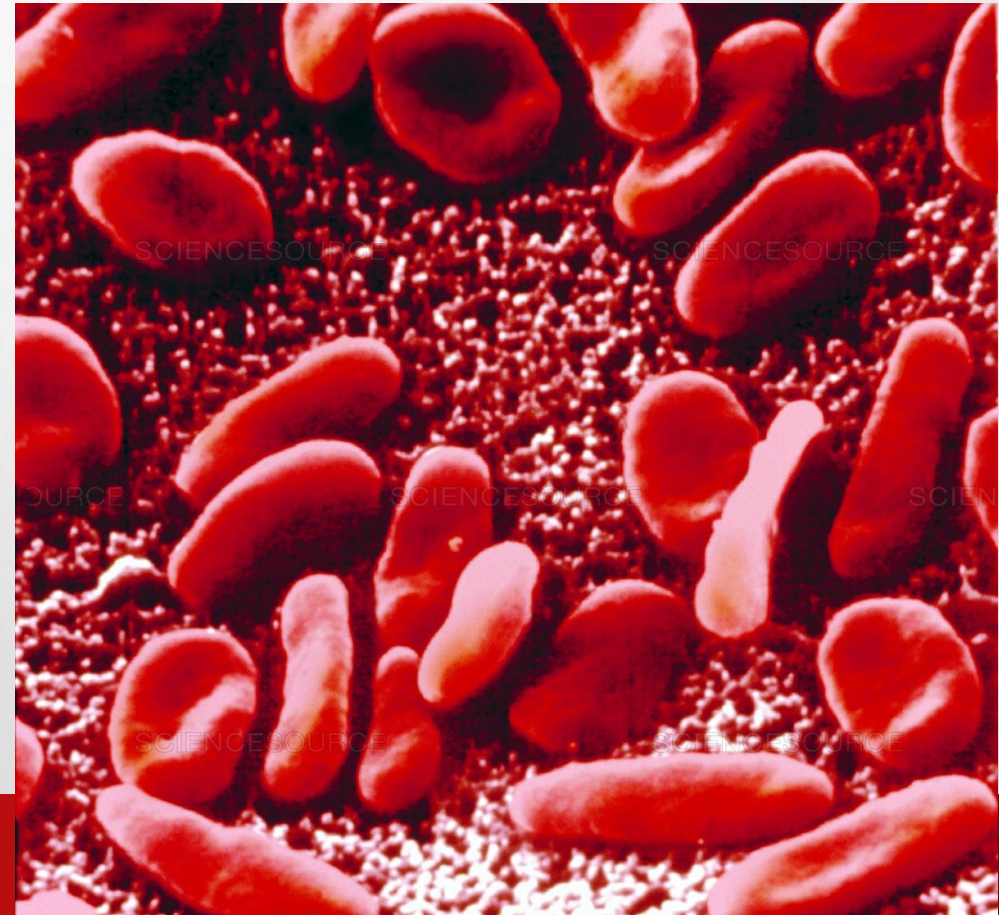
ELLIPTICAL TRAINING

- **DONOR WAS HEALTHY 22-YEAR OLD ASIAN FEMALE**
 - **2X WB DONOR**
 - **PRE-DONATION HCT = 39%**
 - **DEFERRED FROM FUTURE DONATIONS**
- **LATER LEARNED FATHER AND SISTER HAVE HEREDITARY ELLIPTOCYTOSIS (HE)**



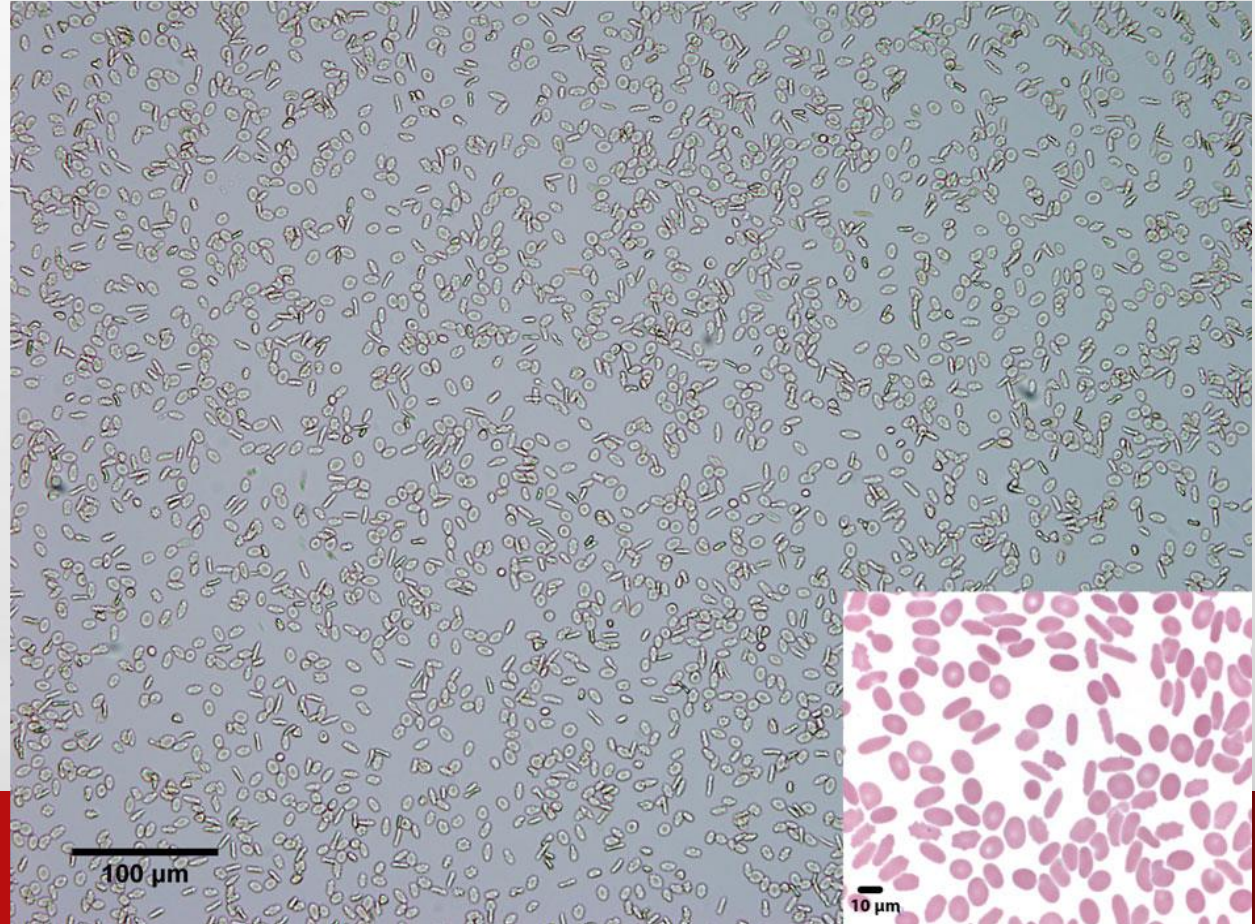
ELLIPTICAL TRAINING

- **ELLIPTOCYTOSIS CAN BE:**
 - **ACQUIRED (IRON, VITAMIN B12, OR FOLATE DEFICIENCY)**
 - **MALIGNANT (MDS OR MYELOPHTHISIC ANEMIA)**
 - **HEREDITARY (SPECTRIN/PROTEIN MUTATIONS)**
- **PEOPLE WITH HE ARE USUALLY ASYMPTOMATIC AND NOT ANEMIC**
- **RBCS FROM HE DONORS HAVE BEEN SHOWN TO HAVE REDUCED SURVIVAL (19-21 DAY HALF-LIFE *IN VIVO*)**



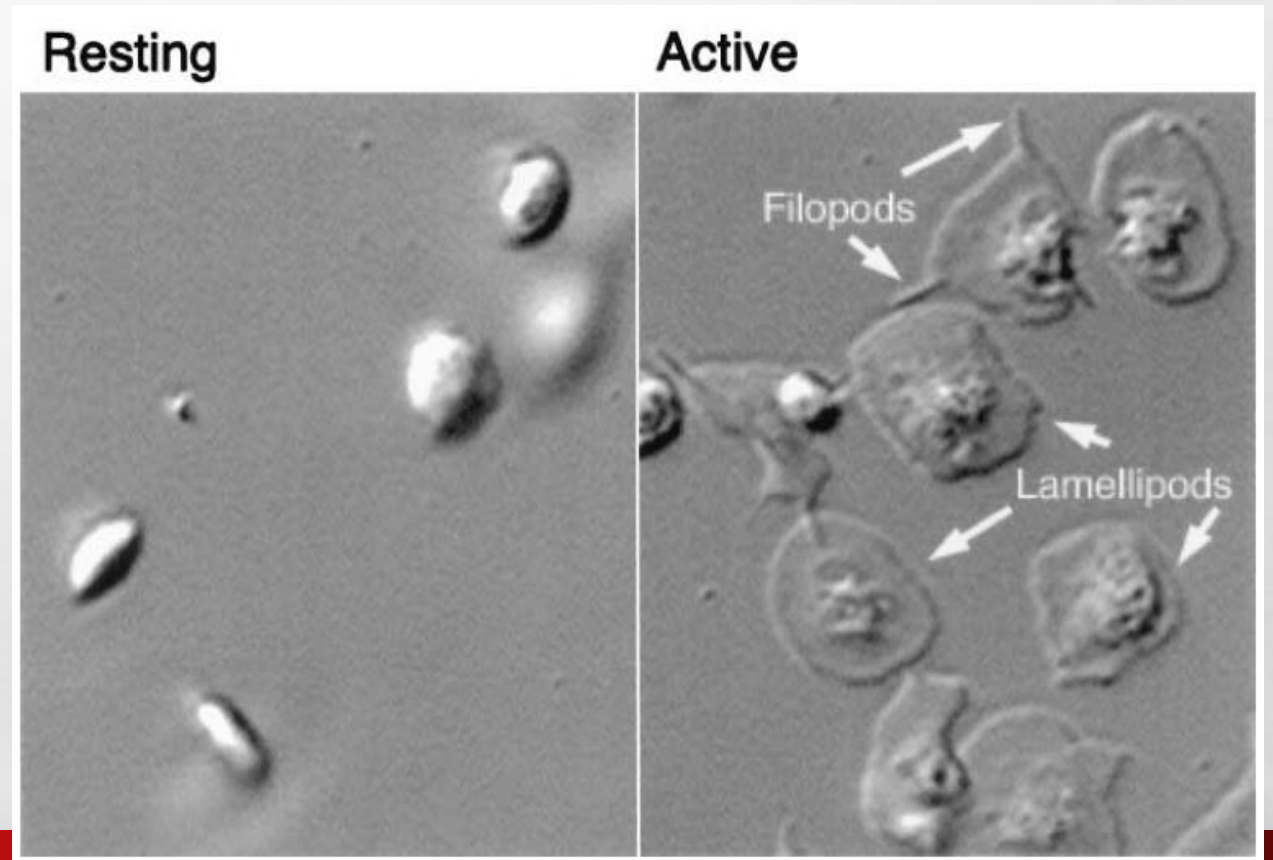
ELLIPTICAL TRAINING

- **DONOR UNIT WAS QUARANTINED AND NOT ISSUED**
- **UNKNOWN IF PATIENTS WITH INHERITED RBC DEFECTS OR HEMOGLOBINOPATHIES CAN TOLERATE RBC UNITS WITH ELLIPTOCYTES**
- **RBC MORPHOLOGY NOT ACTIVELY SCREENED IN DONORS**



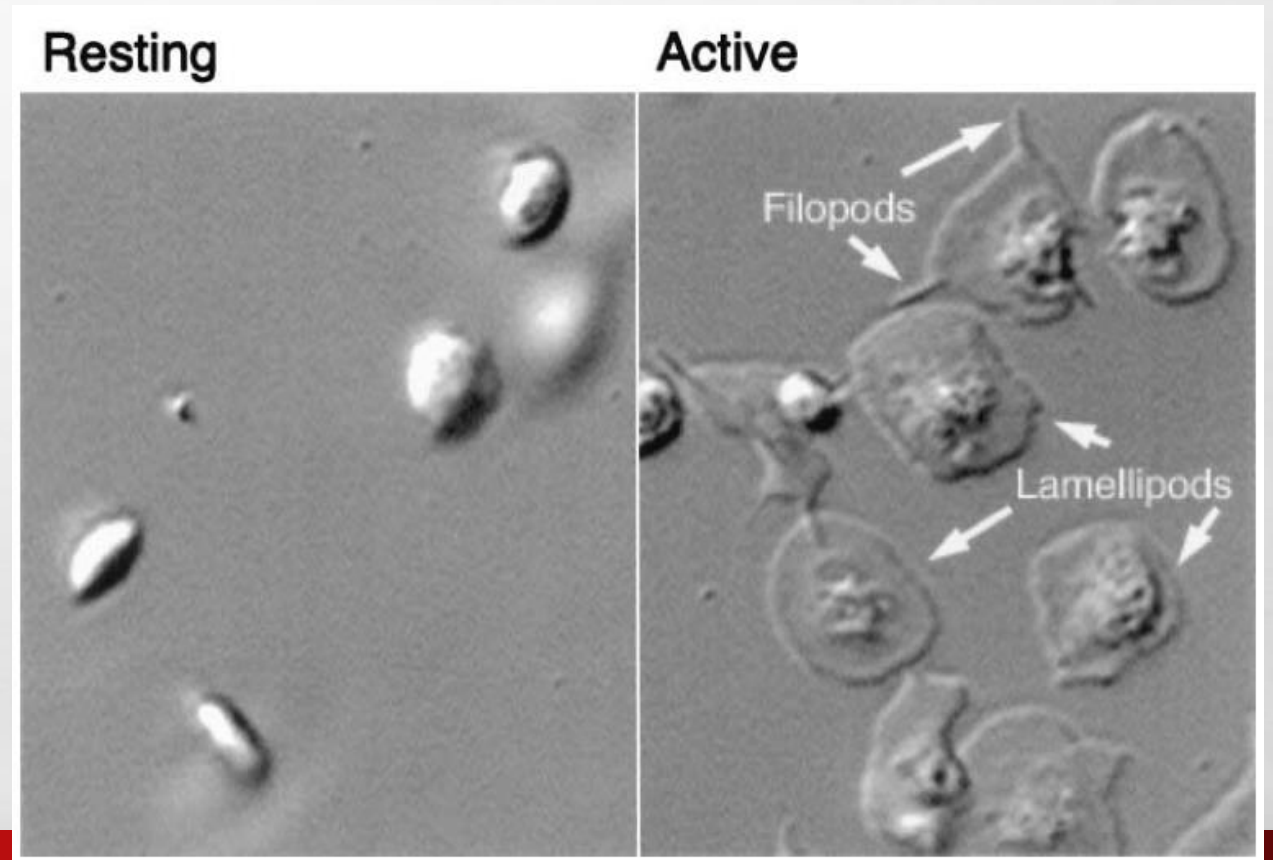
REACHING OUT

- **HARTWIG, J. (2001) THE HUMAN BLOOD PLATELET, RESTING AND ACTIVE. *TRANSFUSION*, 41,723.**



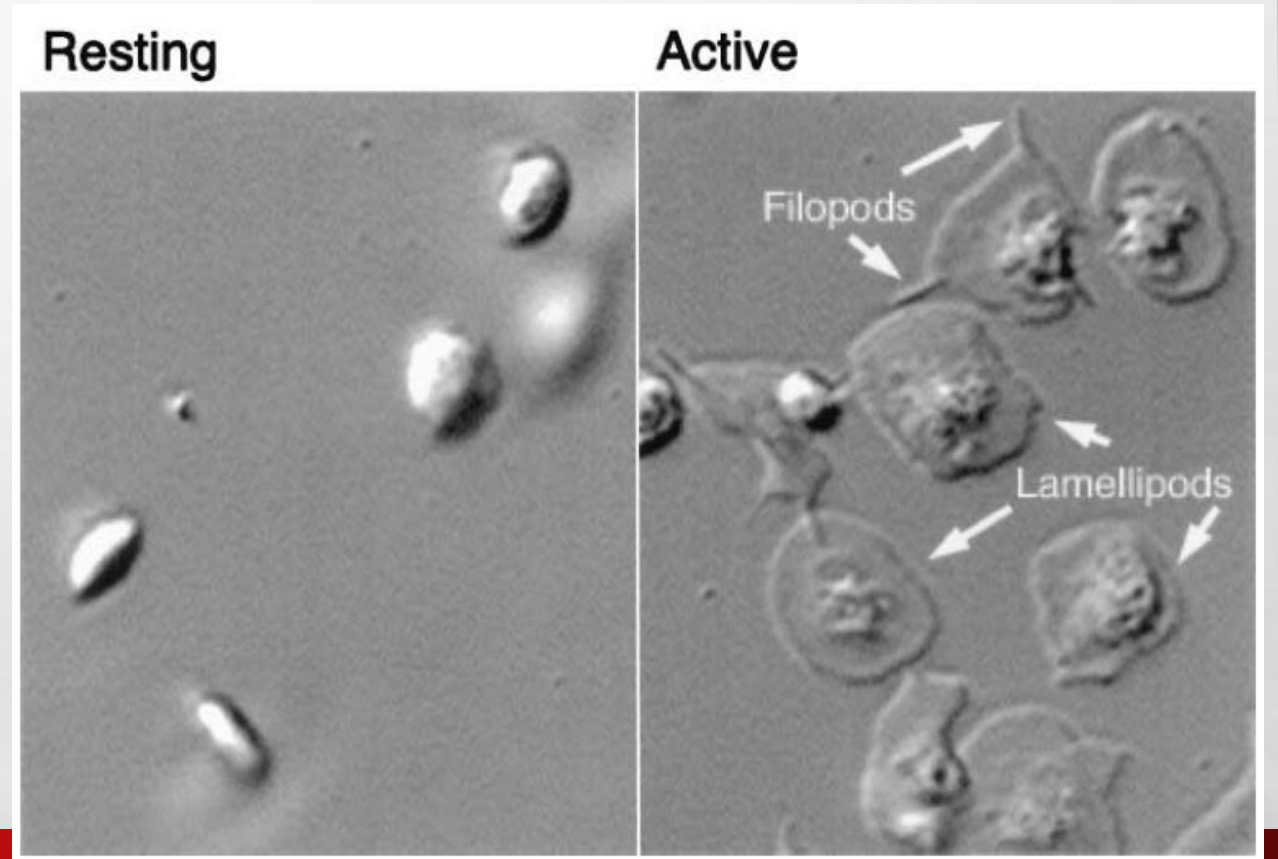
REACHING OUT

- **RESTING PLATELETS CIRCULATE AS SMALL NUCLEATED DISCS**
 - **PHOTOGRAPHED IN SUSPENSION USING DIFFERENTIAL INTERFERENCE CONTRAST OPTICS**



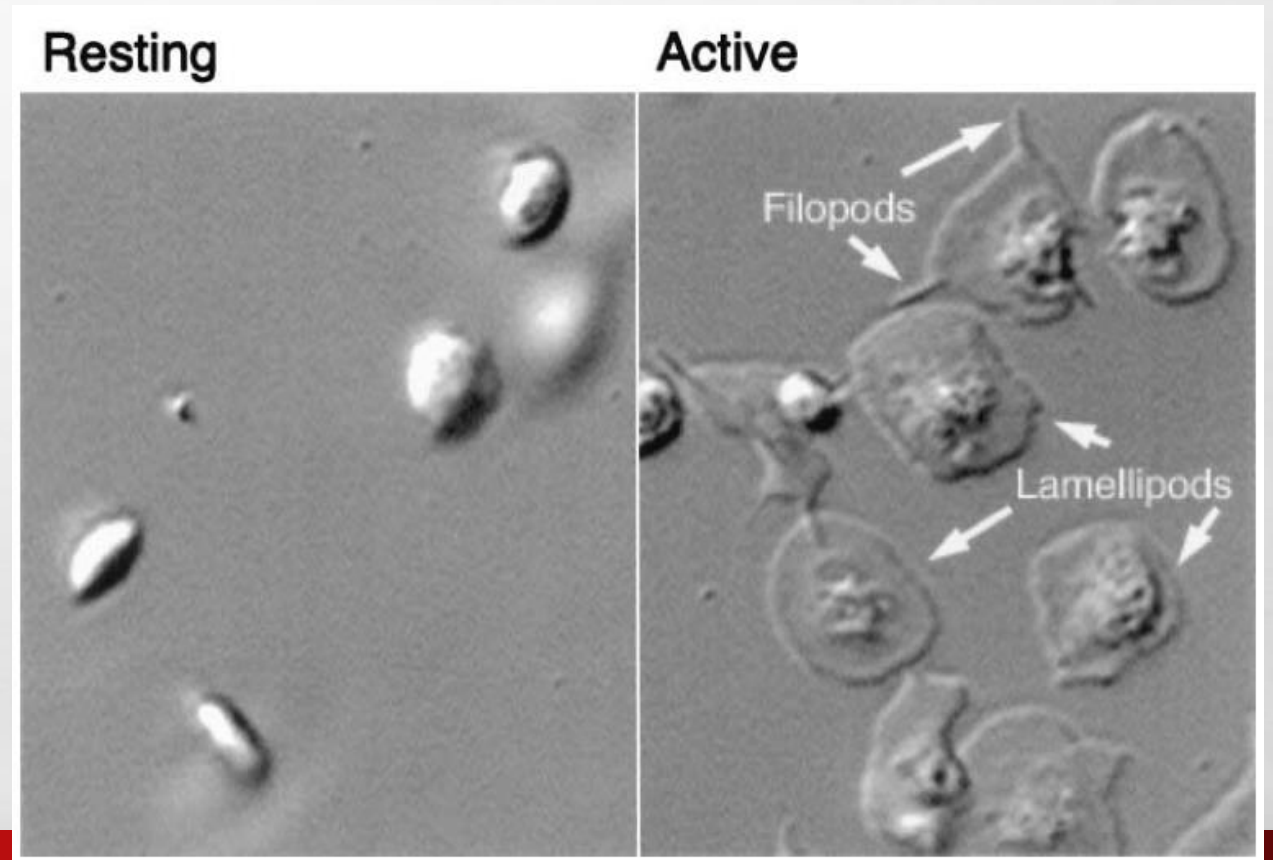
REACHING OUT

- **PLATELETS ACTIVATED BY ATTACHMENT TO SURFACE COATED WITH CRP-LIGAND FOR COLLAGEN RECEPTOR (GPVI)**
- **ACTIVATED PLATELETS ELABORATE FILOPODIA**
 - **TRAVEL AROUND CELL PERIPHERY THEN SPREAD USING LARGE FLAT LAMELLAE**
 - **ASSEMBLY OF CORTICAL ACTIN FILAMENTS DRIVES BOTH GROWTH OF FILOPODIA AND SPREADING OF CELLS**



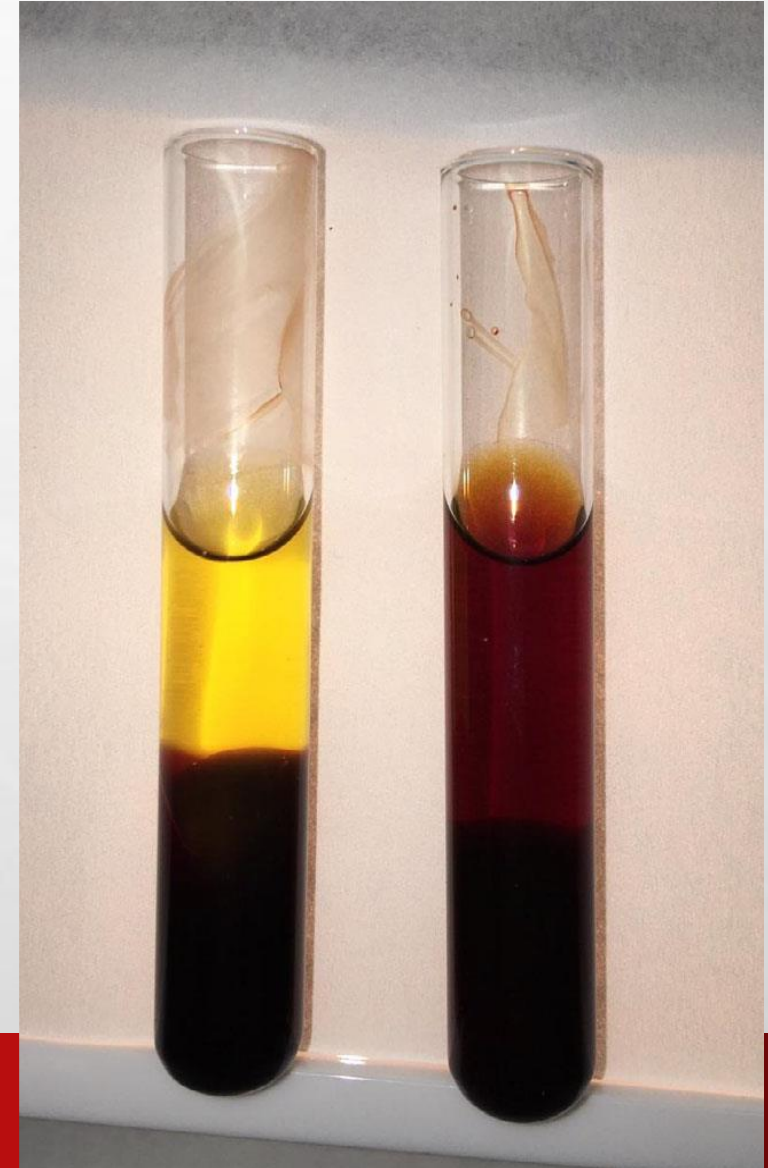
REACHING OUT

- **FILOPODIA ARE USED BY PLATELETS TO FIND AND BIND FIBERS OF FIBRIN AND RECRUIT PLATELETS TO AREA OF DAMAGE**
- **SPREADING OF PLATELETS MAXIMIZES AMOUNT OF DAMAGED SURFACE COVERED BY PLATELETS**



ADD A LITTLE DROP

- **PINEYROA, J. GOMEZ-HERNANDO, M. RODRIGUEZ-ALIBERAS, M. CID, J. MOLINA, A. & LOZANO, M. (2020) HEMOLYSIS IN THE PLASMA OF A SPECIMEN? ADD A DROP TO CLARIFY IT. *TRANSFUSION*, 60, 2783-2784.**



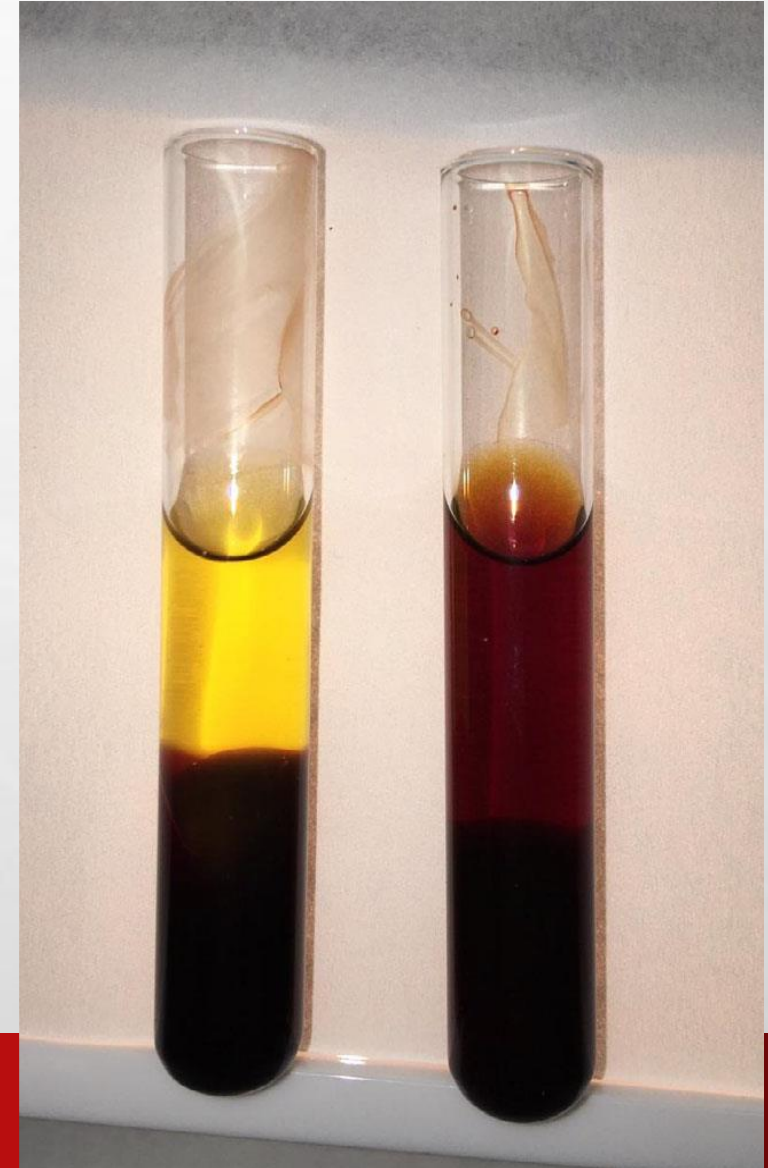
ADD A LITTLE DROP

- **38-YEAR OLD FEMALE WITH APLASTIC ANEMIA**
 - **ADMITTED FOR 3-WEEK HISTORY OF EPIGASTRIC PAIN AND WEIGHT LOSS**
- **TRANSFUSION REQUEST AND SPECIMEN SUBMITTED TO BLOOD BANK**
 - **SPECIMEN SHOWED HEMOLYSIS AFTER CENTRIFUGATION**
 - **RECOLLECT SAMPLE SHOWED SAME HEMOLYSIS**
 - **NO SIGNIFICANT INTERFERENCE IN TESTING**



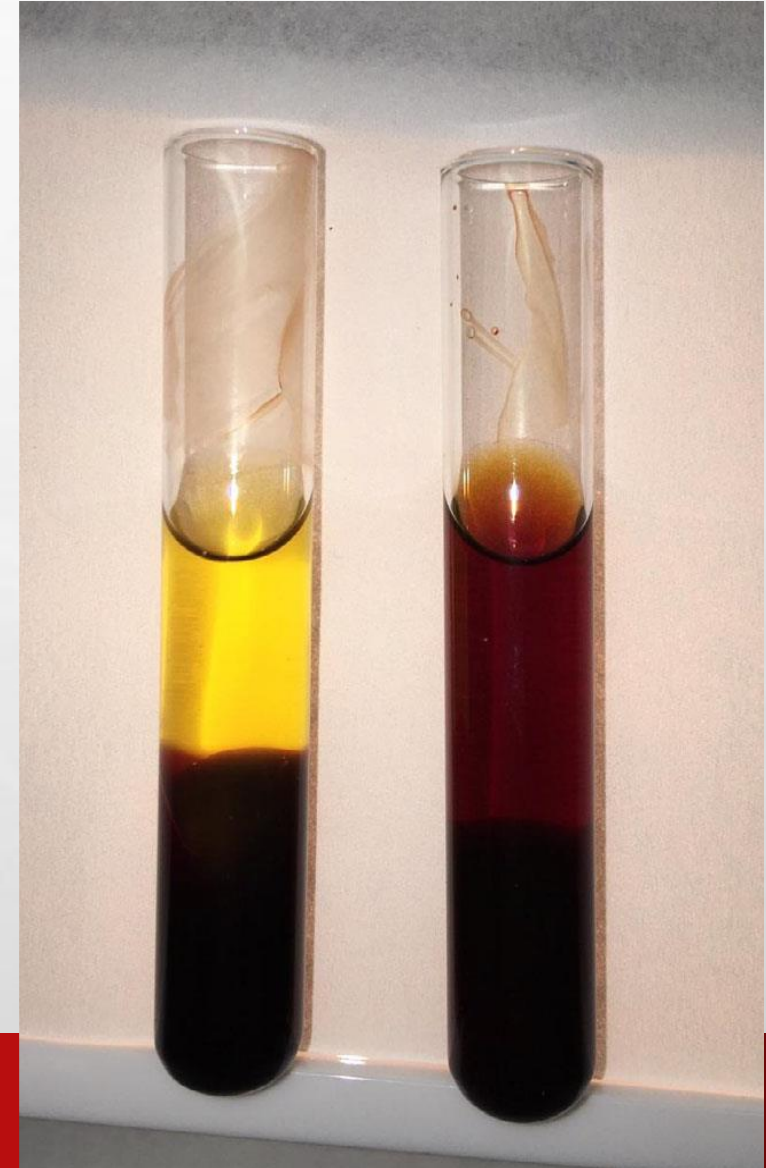
ADD A LITTLE DROP

- **MD WAS CONTACTED ABOUT CONCERN FOR HEMOLYSIS**
 - **PATIENT WAS ASYMPTOMATIC AND BEING DISCHARGED**
- **OTHER LAB VALUES NORMAL:**
 - **LDH, BILIRUBIN, HAPTOGLOBIN**
 - **NO HEMOGLOBINURIA DETECTED**



ADD A LITTLE DROP

- **PATIENT'S MEDICATIONS:**
 - **CYCLOSPORINE, CLONAZEPAM, PANTOPRAZOL, ELTROMBOPAG**
- **ELTROMBOPAG HAS BEEN REPORTED TO CAUSE DARK BROWN PLASMA AT HIGH DOSES**
 - **ORAL THROMBOPOIETIN RECEPTOR AGONIST**
 - **DIACID WITH CARBOXYL & PHENOLIC FUNCTIONAL GROUPS**
 - **COLOR IN SOLUTION VARIES BY PH**



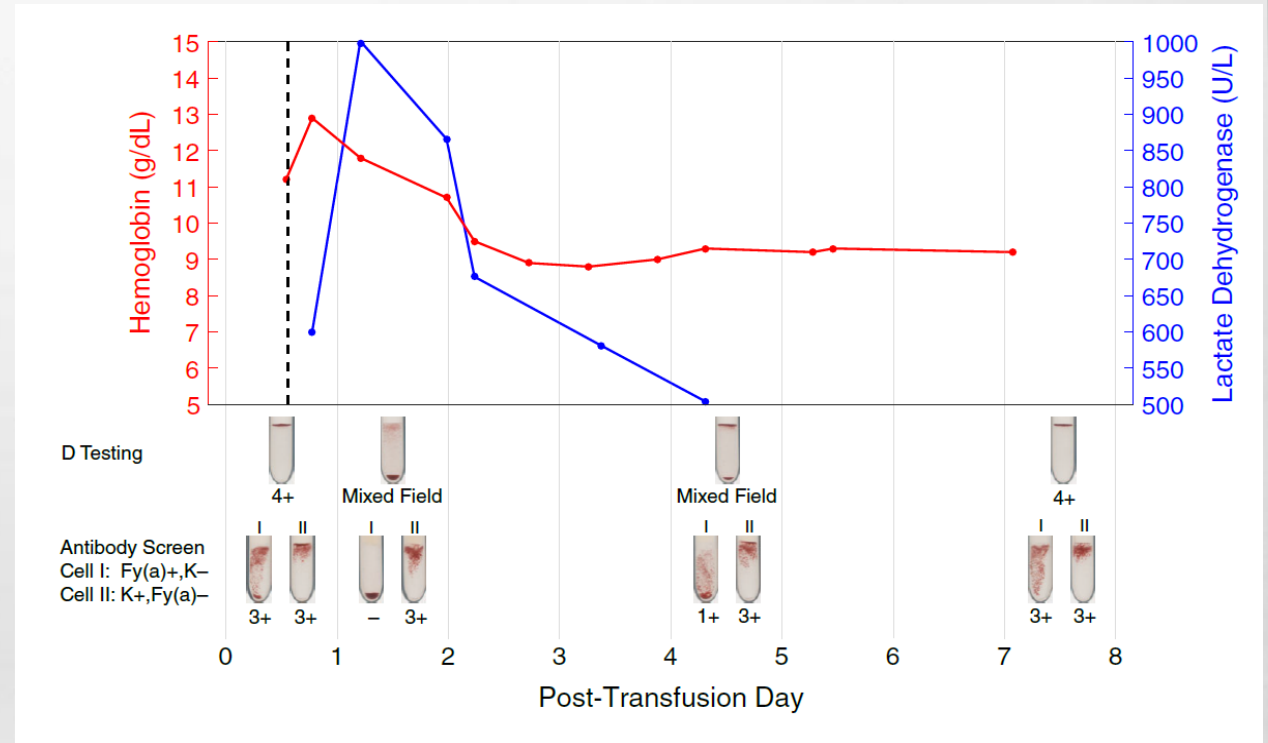
ADD A LITTLE DROP

- **LAB CONDUCTED EXPERIMENT:**
 - **RIGHT TUBE – PATIENT'S PLASMA**
 - **MIDDLE TUBE – PT'S PLASMA + 6M HCL**
 - **LEFT TUBE – PT'S PLASMA + 6M NaOH**
- **COLOR CHANGE DUE TO ACIDIFICATION/ALKALINATION OF PLASMA CONFIRMED ELTROMBOPAG WAS CAUSE OF ABNORMAL PLASMA COLOR**
- **ADDING ACIDIC SOLUTION TO HEMOLYZED SAMPLE CAUSES WHITE PRECIPITATE TO APPEAR**



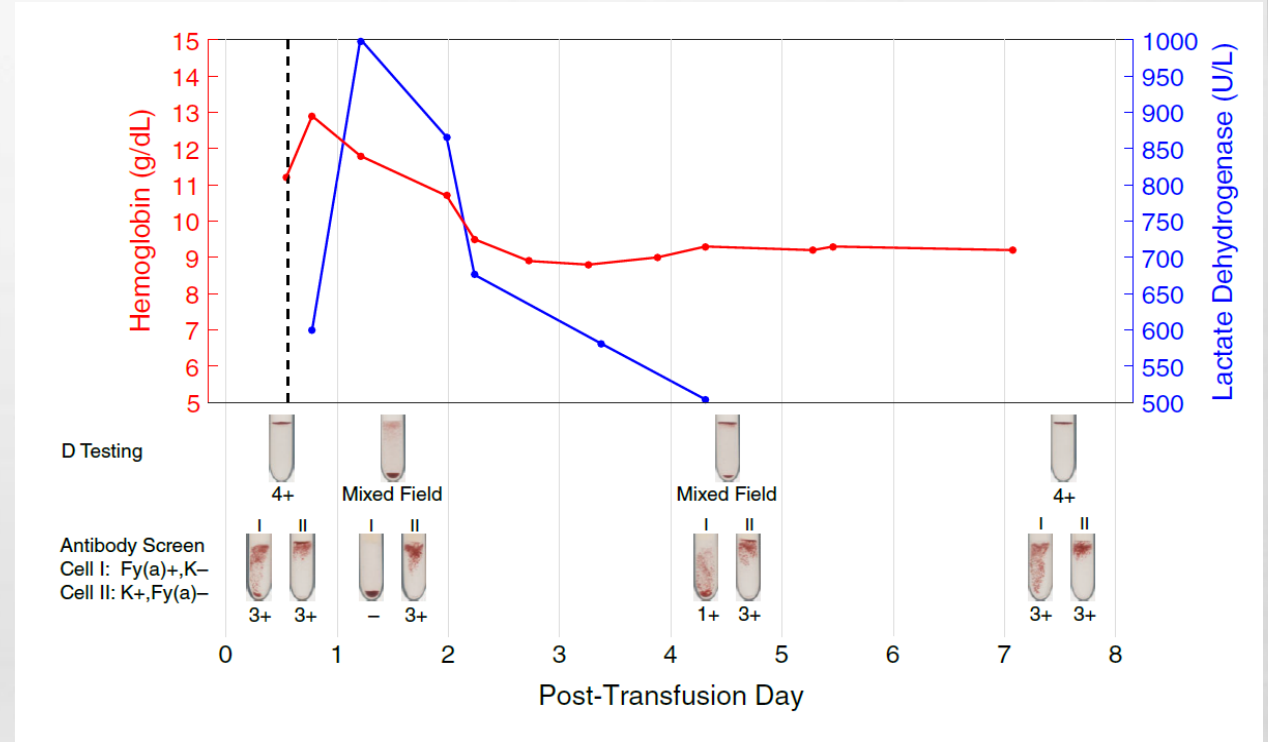
DONOR CLEARANCE

- **CANVER, M. CUSHING, M. VAGELATOS, G. & VASOVIC, L. (2019) VISUAL EVIDENCE OF A HEMOLYTIC TRANSFUSION REACTION IDENTIFIED BY BLOOD BANK TESTING AFTER EMERGENCY BLOOD TRANSFUSION.**
TRANSFUSION, 59, 3301-3302.



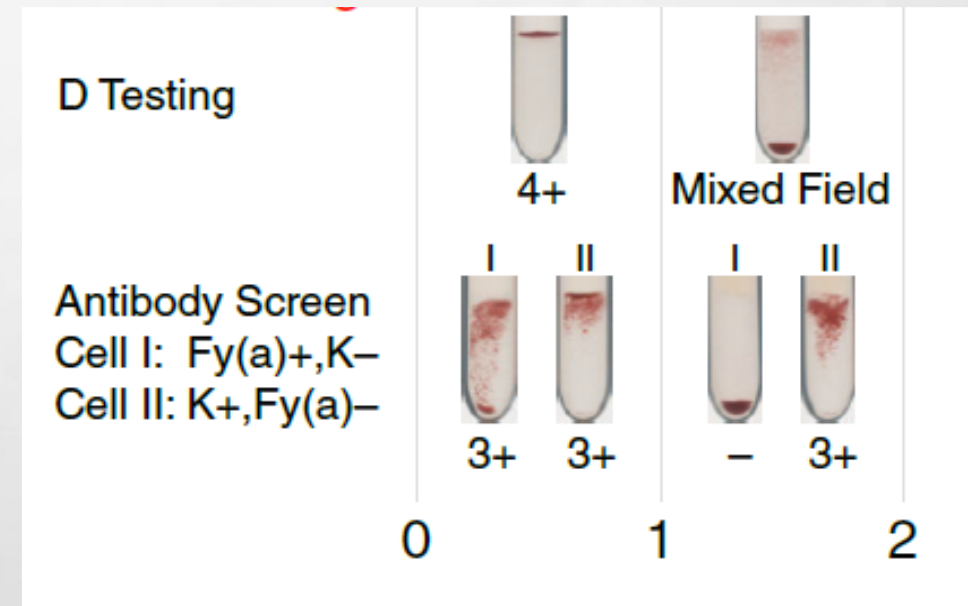
DONOR CLEARANCE

- **76-YEAR OLD O+ FEMALE WITH HEMATEMESIS, HYPOTENSION, AND TACHYCARDIA**
 - **HISTORY OF GASTRITIS**
 - **BROUGHT TO EMERGENCY DEPARTMENT**
- **BP IMPROVED WITH NORMAL SALINE INFUSION**
- **2 UNITS OF O- RBCS TRANSFUSED EMERGENTLY**
 - **BP WAS 101/70 AT TIME OF INFUSION**



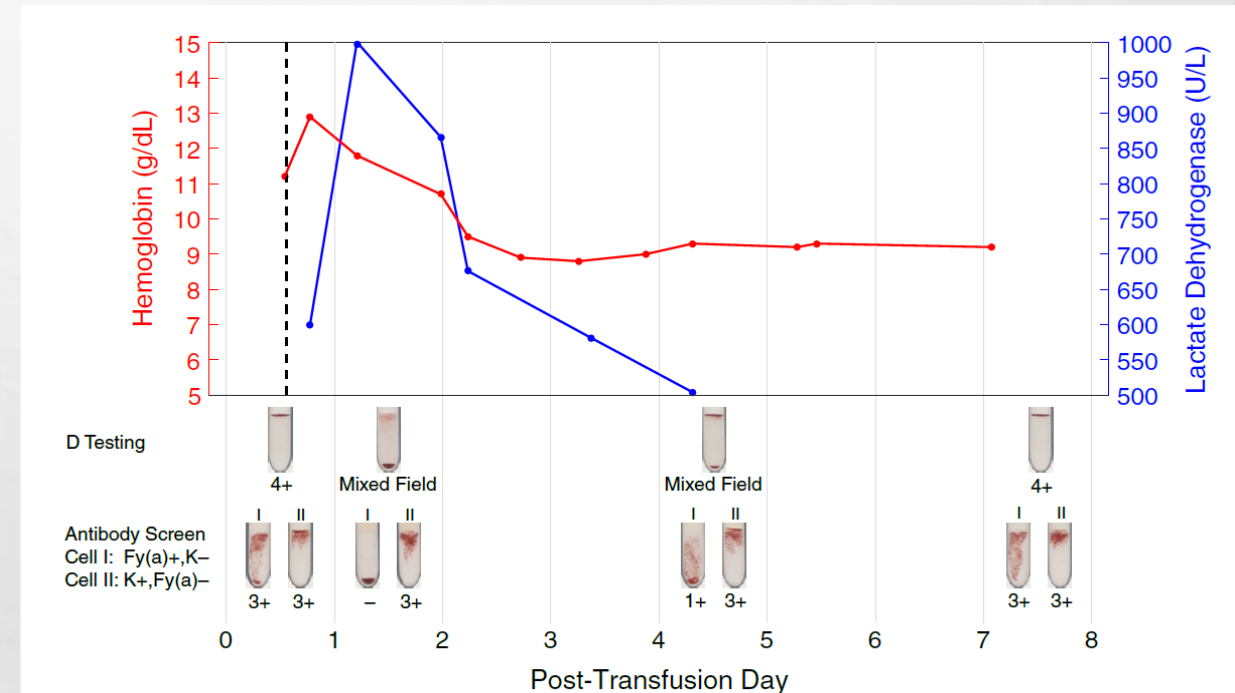
DONOR CLEARANCE

- **PATIENT HAD HISTORY OF ANTI-K AND ANTI-FYA**
- **ABSC AND ABID WERE POSITIVE FOR ANTI-K AND ANTI-FYA**
- **POST-TX SAMPLE HAD POSITIVE DAT**
 - **ANTI-K AND ANTI-FYA IDENTIFIED IN ELUATE**
- **RETROSPECTIVE XM SHOWED INCOMPATIBILITY WITH TRANSFUSED UNITS**
 - **ONE TYPED K+, FY(A-); OTHER K-, FY(A+)**



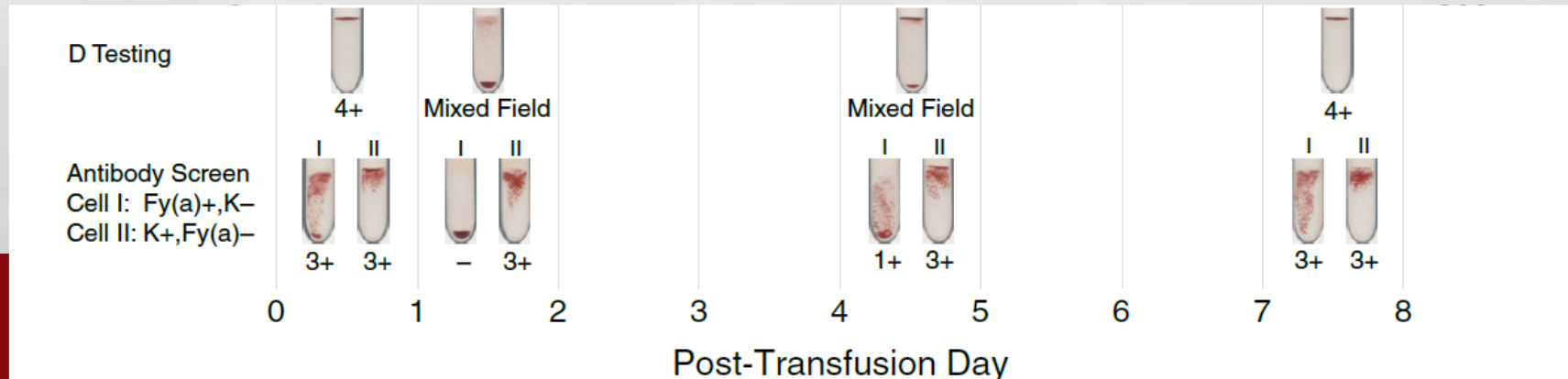
DONOR CLEARANCE

- **PATIENT DEVELOPED HEMOLYTIC TRANSFUSION REACTION FOLLOWING TRANSFUSION**
 - **PRE-HGB WAS 11.2 G/DL; POST-HGB WAS 12.9 G/DL**
 - **HGB DECREASED DESPITE NO ADDITIONAL BLEEDING**
 - **LACTATE DEHYDROGENASE INCREASED**
 - **TOTAL BILIRUBIN INCREASED 0.3 TO 1.3 G/DL**
 - **CREATININE INCREASED 0.8 TO 1.14 MG/DL**
 - **HAPTOGLOBIN WAS 86 MG/DL 5 HOURS POST-TX AND BECAME UNDETECTABLE WITHIN 48 HOURS**



DONOR CLEARANCE

- PATIENT WAS AFEBRILE THROUGHOUT HOSPITALIZATION WITH NO SIGNS/SYMPTOMS OF HEMOLYSIS
- D-TYPING FROM DAYS 1-4 SHOWED MIXED FIELD AGGLUTINATION
- DAY 7 SHOWED NO MIXED FIELD AGGLUTINATION DUE TO HEMOLYSIS OF ALL DONOR RBCS
- SCI WAS NON-REACTIVE ON DAY 1 DUE TO *IN VIVO* ADSORPTION OF ANTI-FYA ONTO DONOR CELLS
 - SCI BECAME POSITIVE AGAIN ON DAY 4



THAT'S A LOT OF BLOOD!

- **HULSE, M. FADER, A. SHAN, H. BENNEH, M. & TOBIAN, A. (2016) MASSIVE TRANSFUSION IN AN OBSTETRIC EMERGENCY. *TRANSFUSION*, 56, 23.**



THAT'S A LOT OF BLOOD!

- **HEALTHY 37-YEAR OLD FEMALE**
 - **GRAVIDA 3, PARA 2 (2 PRIOR CESAREAN SECTIONS)**
 - **BLOOD GROUP A+ WITH NO ALLOANTIBODIES**
 - **DIAGNOSED WITH PLACENTA ACCRETA AND HIGH SUSPICION OF PLACENTA PERCRETA**



THAT'S A LOT OF BLOOD!

- **PLACENTA ACCRETA OCCURS WHEN THE PLACENTA GROWS TOO DEEPLY INTO THE UTERINE WALL**
 - **PART OR ALL OF PLACENTA REMAINS ATTACHED TO UTERINE WALL AFTER BIRTH**
 - **CAN CAUSE SEVERE BLOOD LOSS AFTER DELIVERY**
- **PLACENTA PERCRETA INVOLVES INVASION OF CHORIONIC VILLI THROUGH THE MYOMETRIUM**
 - **OCCASIONALLY INTO OTHER ORGANS**
 - **CAUSES SIGNIFICANT MATERNAL MORBIDITY AND MORTALITY**



THAT'S A LOT OF BLOOD!

- **PATIENT SCHEDULED FOR PLANNED C-SECTION AND HYSTERECTOMY**
- **WENT INTO SPONTANEOUS LABOR AT 34 WEEKS**
 - **PRESENTED WITH PROFUSE VAGINAL BLEEDING**
- **EMERGENT C-SECTION DELIVERY AND EXTENSIVE ABDOMINAL OPERATION WAS PERFORMED**
 - **SAFE, HEALTHY INFANT DELIVERED SUCCESSFULLY**



THAT'S A LOT OF BLOOD!

- **SURGICAL TEAM FOUND EXTENSIVE PLACENTAL INVASION THOUGH:**
 - **BLADDER**
 - **VAGINA**
 - **BILATERAL PARAMETRIA**
 - **UTERUS**
 - **PELVIC SIDEWALL INCLUDING MAJOR VASCULATURE STRUCTURES**



THAT'S A LOT OF BLOOD!

- **SURGICAL TEAM COLLABORATED WITH OBSTETRICS, GYNECOLOGIC ONCOLOGY, TRAUMA SURGERY, UROLOGY, INTERVENTIONAL RADIOLOGY, AND VASCULAR SURGERY TEAMS**
- **PATIENT HAD TOTAL ABDOMINAL HYSTERECTOMY, UPPER VAGINECTOMY, BILATERAL ILIAC ARTERY EMBOLIZATION, RIGHT URETERAL STENT, LEFT URETERAL LIGATION, TRIGONAL BLADDER RESECTION, AND INFRARENAL AORTA REPAIR**



THAT'S A LOT OF BLOOD!

- **MASSIVE TRANSFUSION PROTOCOL WAS INITIATED AT SURGERY BEGINNING AND REMAINED ACTIVE FOR 15 HOURS**
- **PATIENT TRANSFUSED WITH:**
 - **195 RBCS**
 - **155 PLASMAS**
 - **26 APHERESIS PLATELETS**
 - **8 CRYOPRECIPITATE POOLS**
 - **3 DOSES OF FVIIA**
 - **7L OF CELL SAVER**



THAT'S A LOT OF BLOOD!

- **ESTIMATED BLOOD LOSS WAS 180L**
- **INTRAOPERATIVE HGB VALUES RANGED BETWEEN 8.1 – 13.9 G/DL**
 - **NEAR NORMAL COAGULATION PROFILE OBSERVED**
- **HEMOSTASIS ACHIEVED AFTER 14 HOUR SURGERY**
 - **PATIENT TRANSFERRED TO SURGICAL ICU WITHOUT VASOPRESSOR SUPPORT**
 - **PATIENT EXTUBATED POST-OP DAY 3 AND MET HEALTHY NEWBORN**



THAT'S A LOT OF BLOOD!

- **PATIENT UNDERWENT ADDITIONAL BLADDER RECONSTRUCTION SURGERY**
- **PATIENT DID WELL AND HAD UNEVENTFUL HOSPITAL COURSE**
- **MASSIVE TRANSFUSION PROTOCOLS HELP BLOOD BANKS ISSUE CORRECT QUANTITY AND PROPORTION OF BLOOD PRODUCTS IN STRESSFUL SITUATIONS**



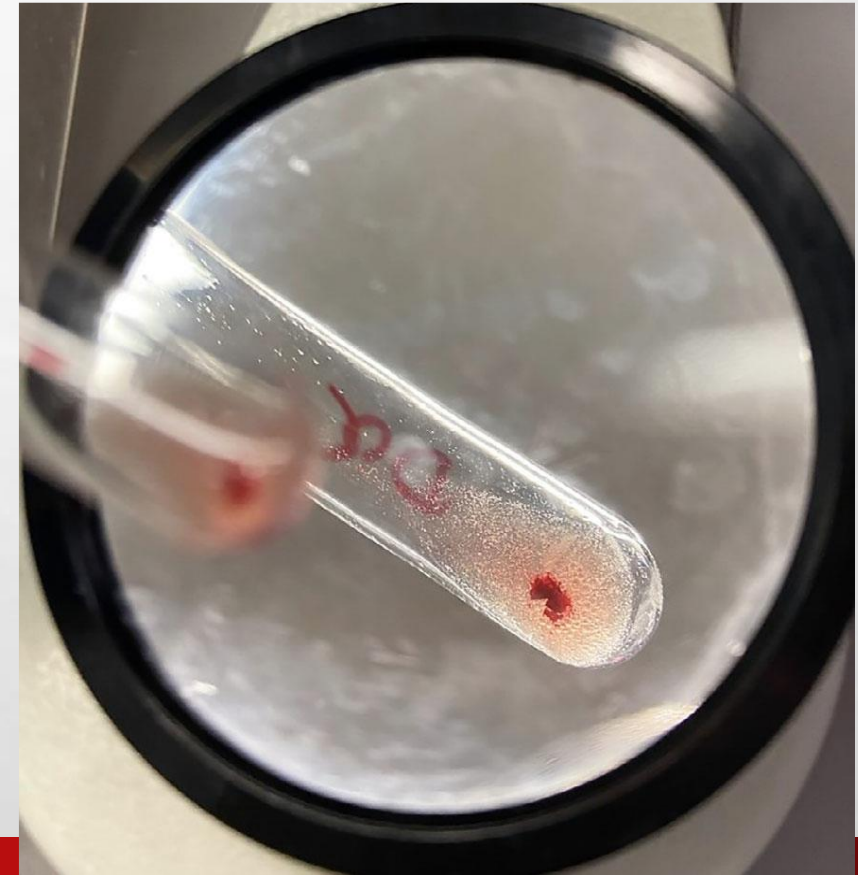
BLOODY EMULSION

- **MAYER, R. WARD, M. MERRILL, V. PEISACH, N. SHIMER, L. & MURPHY, C. (2020) AN INTRAOSSEOUS SAMPLE INITIALLY RESEMBLING PERIPHERAL BLOOD. *TRANSFUSION*, 60, 1346-1347.**



BLOODY EMULSION

- **BLOOD BANK RECEIVED SAMPLE LABELED “PERIPHERAL BLOOD” FROM TRAUMA PATIENT**
 - **PATIENT HAD OPEN BOOK PELVIC RING DISRUPTING FRACTURE**
- **SAMPLE APPEARED NORMAL AFTER CENTRIFUGATION**
 - **MANUAL TUBE TESTING WAS PERFORMED**



BLOODY EMULSION

- **PATIENT'S BACKTYPE SHOWED “SALAD DRESSING”-LIKE APPEARANCE OF FAT DROPLETS**
- **BLOOD BANK MD SUSPECTED SAMPLE WAS INTRAOSSEOUS (IO) SAMPLE**
 - **CONFIRMED BY TRAUMA UNIT**
 - **DRAWN FROM LEFT TIBIAL PLATEAU IO DEVICE**



BLOODY EMULSION

- **PATIENT TYPED A+ WITH MIXED-FIELD IN THE FORWARD ANTI-A REACTION**
 - **DUE TO 6 O+ RBCS EMERGENCY RELEASED PRIOR TO TESTING**
- **NEGATIVE ANTIBODY SCREEN**
- **FAT DROPLETS ONLY APPARENT IN TUBE TESTING**
- **IO SAMPLE CONSIDERED INVALID**
 - **O+ RBCS / A OR AB PLASMA ISSUED DURING INITIAL RESUSCITATION**



BLOODY EMULSION

- **PATIENT TYPED A+ WITH NEGATIVE ANTIBODY SCREEN ON A PERIPHERAL BLOOD SAMPLE ON DAY 4**
- **BONE MARROW ASPIRATES ALSO HAVE BEEN PREVIOUSLY SHOWN TO BE CONCORDANT WITH PERIPHERAL BLOOD**



PETECHIAE-AY-AY!

- **PRITCHARD, A. & LOCKHART, E. (2017) RUMPEL-LEEDE PHENOMENON IN A PATIENT BEING TREATED WITH PRASUGREL. *TRANSFUSION*, 57,1642.**



PETECHIAE-AY-AY!

- **73-YEAR OLD MALE WITH WALDENSTROM MACROGLOBULINEMIA**
- **UNDERGOING THERAPEUTIC PLASMA EXCHANGE FOR HYPERVISCOSITY**
- **DEVELOPED RUMPEL-LEEDE SIGN AFTER PLACEMENT OF BLOOD PRESSURE CUFF**



PETECHIAE-AY-AY!

- **PATIENT'S HISTORY:**
 - **MYOCARDIAL INFARCTION**
 - **TAKING ASPIRIN AND PRASUGREL**
 - **NORMAL PLT COUNT AND NORMOTENSIVE**
 - **NO HISTORY OF DIABETES, CONNECTIVE TISSUE DISORDERS, OR BRUISING/BLEEDING ISSUES**
- **PATIENT DEVELOPED DIFFUSE PETECHIAE AND SCATTERED PURPURA DISTAL TO BP CUFF INFLATION**
 - **RESOLVED WITHIN 1 WEEK**



PETECHIAE-AY-AY!

- **RUMPEL-LEEDE SIGN IS ACUTE DERMIS CAPILLARY RUPTURE**
 - **DEVELOPMENT OF PETECHIAE AFTER SUDDEN INCREASES IN VENOUS PRESSURE (SUCH AS TOURNIQUET OR BP CUFF)**
 - **ASSOCIATED WITH THROMBOCYTOPENIA, PLT DYSFUNCTION, AND CAPILLARY FRAGILITY**
 - **PREVIOUSLY DESCRIBED IN PATIENTS WITH DIABETES, EHLERS-DANLOS SYNDROME, AND HYPERTENSION**



PETECHIAE-AY-AY!

- **IN THIS PATIENT, IT WAS SUSPECTED THE ASPIRIN AND PRASUGREL'S ANTI-PLT ACTIVITY MAY HAVE CONTRIBUTED TO DEVELOPMENT OF HIS RUMPEL-LEEDE SIGN**
 - **WALDENSTROM MACROGLOBULINEMIA-ASSOCIATED HYPERVISCOSITY COULD NOT BE RULED OUT AS A FACTOR**



NO, THAT IS INCORRECT

- **MOIZ, B. ADIL, S. & SULEMAN, M. B SUBGROUP MISTYPED AS O. (2007) *TRANSFUSION*, 47, 753**



NO, THAT IS INCORRECT

- **44-YEAR OLD MALE PRESENTED TO EXECUTIVE CLINIC WITH CHEST PAIN**
- **MD REQUESTED ABORH ALONG WITH OTHER VARIOUS ROUTINE LAB TESTING**
- **PATIENT'S PREVIOUS ABORH TYPINGS BY VARIOUS LABS ALL REPORTED AS O POSITIVE**
- **GEL TYPING SHOWED FRONT TYPE AS O POS, BUT BACK TYPE AS B**



NO, THAT IS INCORRECT

- **AUTOCONTROL WAS NEGATIVE AND PROVISIONAL INTERPRETATION WAS B SUBGROUP OR O WITH LOW-LEVEL ANTI-B**
- **RECOLLECT REQUESTED AND TESTED USING MANUAL TUBE AND GEL METHODS WITH INCLUSION OF ANTI-A,B**
 - **RECOLLECT RESULTED IN AGREEMENT WITH PREVIOUS RESULT**



NO, THAT IS INCORRECT

- **WEAK AGGLUTINATION OBSERVED IN ANTI-B AND ANTI-A,B TUBES AFTER 4-6 HOUR INCUBATION AT ROOM TEMPERATURE**
- **PROLONGED INCUBATION OF REVERSE GROUPING AND AUTOCONTROL AT RT AND 4°C DID NOT ALTER RESULTS**
 - **BOTH RESULTS SUGGEST PATIENT HAS WEAK B SUBGROUP**



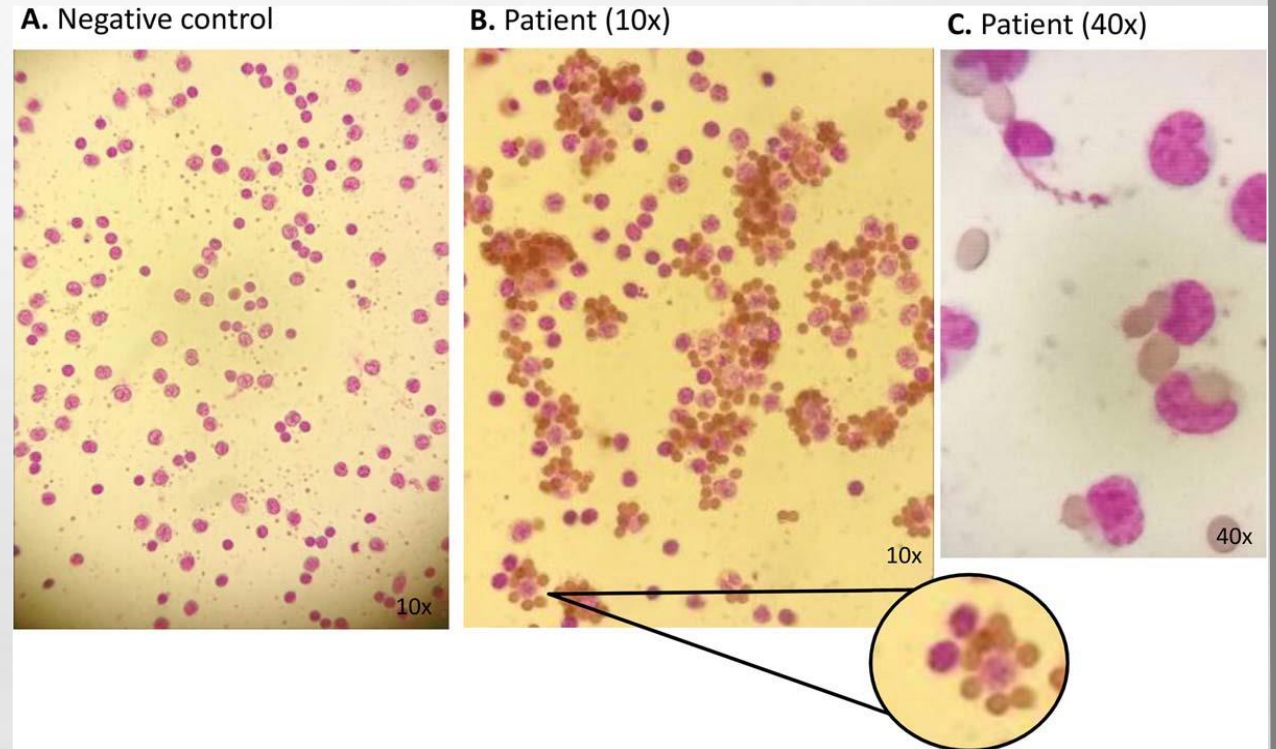
NO, THAT IS INCORRECT

- **ADSORPTION AND ELUTION STUDIES WITH ANTI-B ANTISERA SHOWED PRESENCE OF B ANTIGEN ON PATIENT'S RBCS**
 - **ELUATES REACTED WITH 3 B TEST CELLS BUT NON-REACTIVE WITH 3 GROUP O CELLS**
- **B SUBGROUPS ARE INFREQUENT BUT RECOGNITION OF THEM IS IMPORTANT**
 - **FAMILY STUDIES AND/OR MOLECULAR GENETICS CAN ALSO BE USEFUL TOOLS FOR ABORH TYPING RESOLUTIONS**



MMA IN THE BLOOD BANK

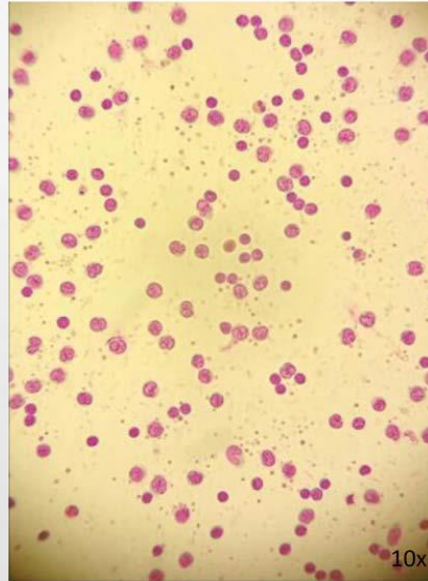
- **CONRADO, M. BONIFACIO, S. NOGUEIRA, F. SARAIVA-FILHO, J. DEZAN, M. CHINOCA, K. GOMES, F. FONSECA, G. GUALANDRO, S. ROCHA, V. MENDRONE-JUNIOR, A. & DINARDO, C. (2018) MASSIVE AUTOIMMUNE HEMOLYSIS DOCUMENTED BY MONOCYTE MONOLAYER ASSAY IN A MULTIPLY TRANSFUSED PATIENT USING RETICULOCYTES ISOLATED BY SIMPLE CENTRIFUGATION IN MICROHEMATOCRIT TUBES. *TRANSFUSION*, 58, 1578-1579.**



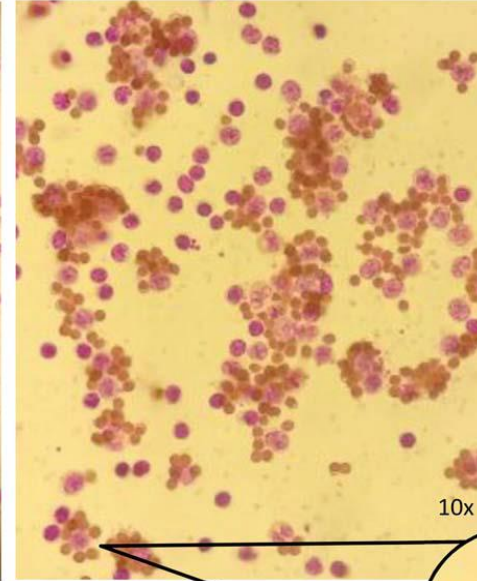
MMA IN THE BLOOD BANK

- **34-YEAR OLD MALE POSITIVE FOR HIV ADMITTED FOR INTENSE ANEMIA AND HEMODYNAMIC INSTABILITY**
 - **HGB WAS 1 G/DL, MCV WAS 110 FL/CELL, AND RETICULOCYTE COUNT WAS 12K /MM³**
- **TRANSFUSED WITH 5 GENOTYPE-COMPATIBLE RBCS**
 - **RORO (RHD*01N.01), FY(A-), JK(B-), S-**

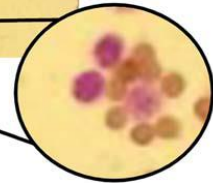
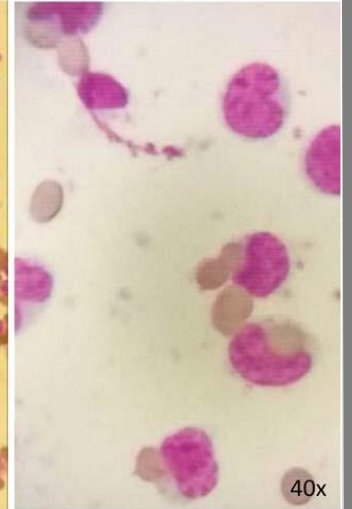
A. Negative control



B. Patient (10x)



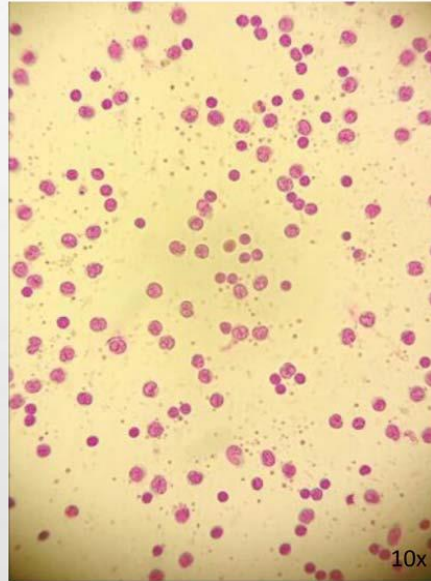
C. Patient (40x)



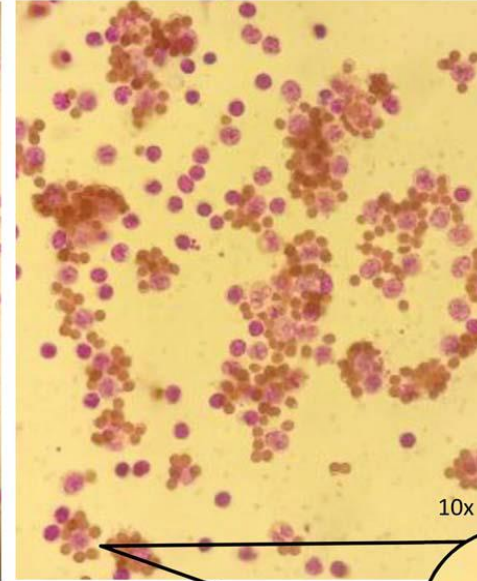
MMA IN THE BLOOD BANK

- **PRE-TRANSFUSION TESTING:**
 - **4+ DAT (IGG, IGM, AND C3)**
 - **4+ AUTOCONTROL**
 - **SERUM PANAGGLUTINATION**
- **PERIPHERAL BLOOD SMEAR REVEALED INTENSE ANISOCYTOSIS BUT NO ERYTHROPHAGOCYTOSIS**
- **MONOCYTE MONOLAYER ASSAY (MMA) TESTING INDICATED FOR AUTOIMMUNE RBC LYSIS**

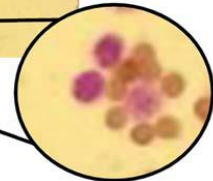
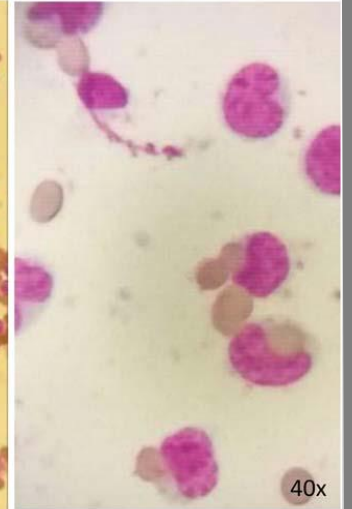
A. Negative control



B. Patient (10x)



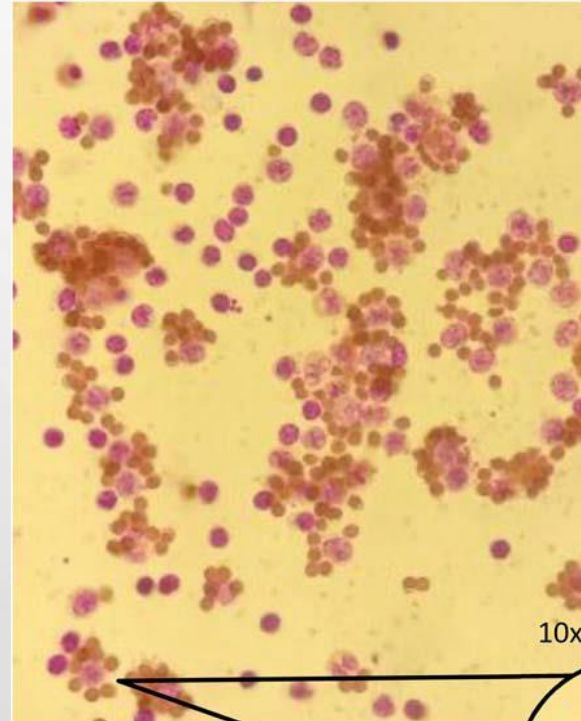
C. Patient (40x)



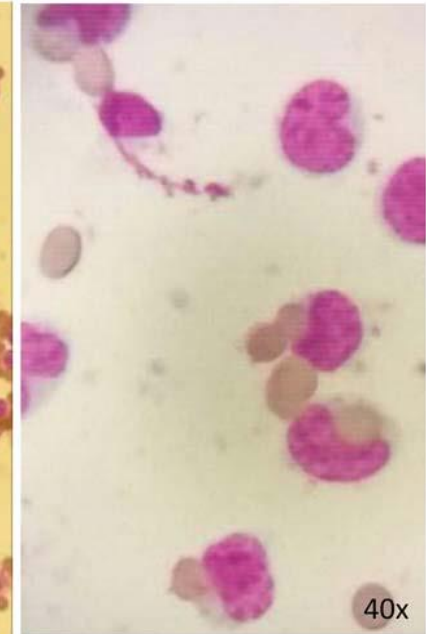
MMA IN THE BLOOD BANK

- **MMA TEST TYPICALLY USED TO GAUGE *IN VIVO* REACTIVITY OF DONOR RBCS WITH PATIENT'S SERUM**
 - **HIGH-PREVALENCE ANTIBODY**
- **PATIENT SERUM MIXED WITH DONOR RBCS AND DONOR MACROPHAGES**
 - **PT AB ATTACHED TO DONOR RBCS**
 - **MACROPHAGES ATTACH (ROSETTES) AND PHAGOCYTIZE DONOR RBCS**

B. Patient (10x)

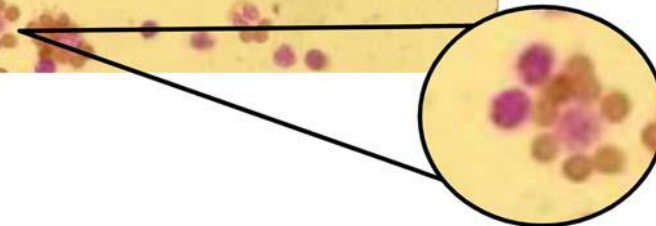


C. Patient (40x)



10x

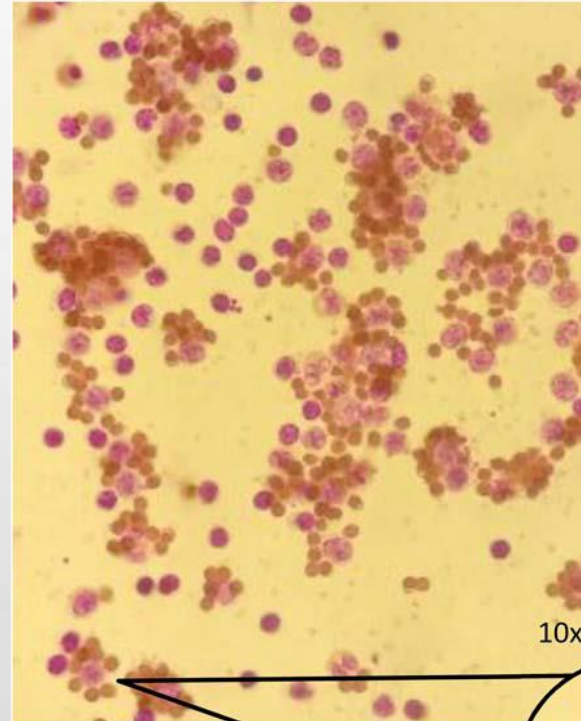
40x



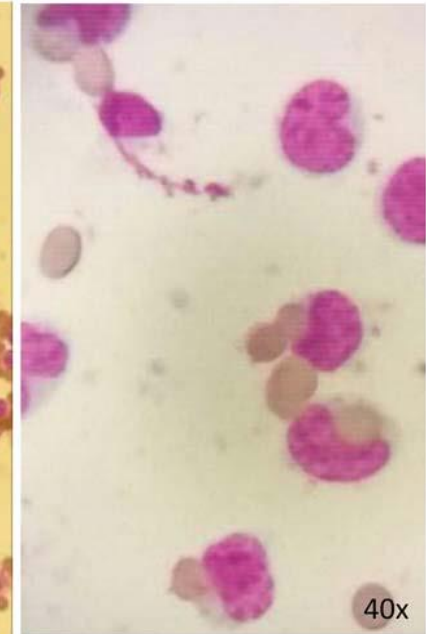
MMA IN THE BLOOD BANK

- **IN THIS CASE, PATIENT RBCS USED INSTEAD OF DONOR RBCS**
 - **RECENTLY TRANSFUSED**
 - **RETICULOCYTES ISOLATED FROM PATIENT SAMPLE USING CENTRIFUGED MICROHEMATOCRIT TUBES**
- **MMA OBTAINED IMPRESSIVE RESULTS**
 - **ALMOST ALL RBCS WERE PHAGOCYTIZED OR FORMED ROSETTES**

B. Patient (10x)

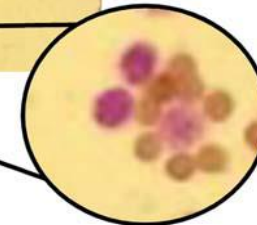


C. Patient (40x)



10x

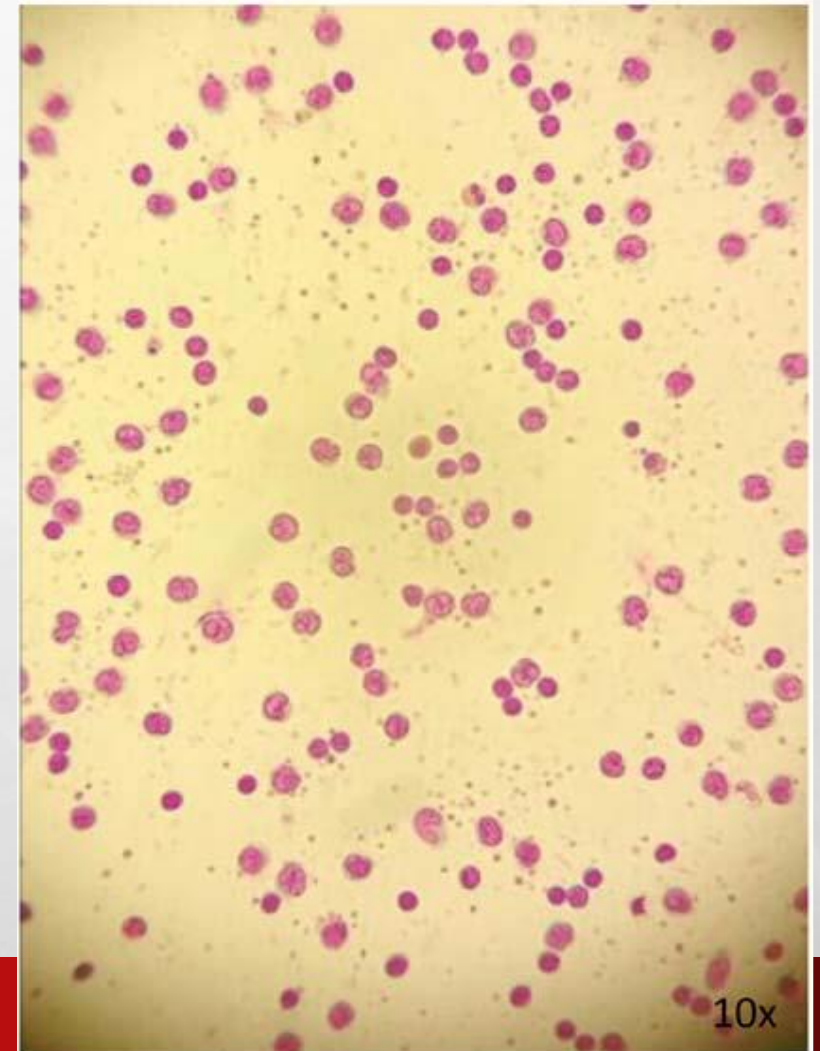
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MMA IN THE BLOOD BANK

- **MMA TEST USING NORMAL RETICULOCYTES WAS NEGATIVE**
- **DIAGNOSIS OF AGGRESSIVE AUTOIMMUNE HEMOLYSIS WAS CONFIRMED**

A. Negative control



QUESTIONS?



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TRANSFUSION

