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of the procedure. The equivalence of technical performance with hemoglobin S (HbS) parameters after treatment, fraction of remaining cells (FCR), procedure duration, processed volumes of blood and anticoagulant and consumption of RBC units were confirmed.23-26 The RBCD/RBCX protocol in SPO and the hemodilution/RBCX isovolemic protocol in Cobe are comparable.23,24 No significant differences were found in HbS, hematocrit, FCR, and number of platelets. Interestingly, rbcdd made with SPO requires a significantly lower saline replacement volume and rinse volumes (P<0.001). However, a longer process time should be noted in the OPS.24 An effective reduction in HbS was confirmed by Daniel et al. in a simple exchange of erythrocytes, initial HbS levels were reduced from 73%-85% to 22%-29%.25 The main objective in RBCX SPO was to confirm that the patient's expected RCF at the end of the procedure reflects the actual FCR, measured in HbS%. The average expected RATIO of FCR/FCRp was 0.90 (95% CI, 0.86 to 0.94) within the pre-defined acceptable range of 0.75 to 1.25. The safety profile of RBCD/RBCX in 60 patients examined by Quirolo et al. remained effective without severe AE and unexpected adverse side effects.26 However, hypotension in RBCD/RBCX procedures with SPO occurred more often in 7% of cases compared to 1.8% in Cobe. Erythrocyte depletion is mentioned as a method of removing iron in patients with hereditary hemochromatosis. To this end, a small study was published where an average blood volume of 857.3 ± 22.3 mL with a short average treatment period of 12.0 ± 0.4 min was treated, and the average hematocrit per session was lowered by 6%. Iron of 405.2 ± 23.2 mg per procedure was removed.27 Collection of mononuclear cells (MNC) For collection MNC, 2 different options are available. The MNC procedure performs successively several phases of accumulation and collection. Here, a high centrifugal force is exerted to achieve an optimal packaging factor (default is 20) for anticoagulated anticoagulated to enter the separation channel. The Automatic Interface Management System (AIM) regulates the flow of the plasma pump to control the concentration of cells circulating through the collection port. The collection pump transfers the MNC and platelets from the regulatory chamber to the second chamber where the platelets are selectively removed by elutriation and returned to the patient. The optimal positioning of the interface should be defined via the collection preference based on the desired target performance of the product cells and the content of the erythrocytes.30,34 The second option is the continuous MNC procedure (CMNC) in which the patient's blood is pumped into the tube set and the centrifuge rotates to reach the default packing factor of 4.5 on a slightly larger volume separation channel with a less interface layer. Minor. The AIM system regulates the flow of the plasma pump to control the concentration of cells passing through the collection port, depending on the collection preference. If the cells are detected by the AIM system, the valve of the collection pipe moves to the collection position, and the collection pump pushes the MNC into the collection bag. The platelets are suspended in the plasma layer and are returned directly to the patient without the need for a secondary separation chamber.34 The indications for MNC and cmnc procedures are identical in principle. Examples of clinical application are the offline method of extracorporeal photopheresis (ECP), collection of autologous peripheral blood stem cells, and other new immunotherapies such as immature dendritic cell treatment or chimeric antigen receptor-modified cell therapy. Two of the most common indications are apheresis of allogeneic peripheral blood stem cells and donor lymphocytes, which will not be discussed further here. All of these approaches require reliable MNC collection with a defined cell population composition. In the database analysis with PubMed and Medline, MNCs were identified for apheresis with OPS in 28 studies between June 2011 and December 2017. ECP extracorporeal photopheresis is an established cell therapy for the treatment of T lymphoma skin lymphoma, graft against host disease, and organ rejection after organ transplantation.29 Several studies have been performed on MNCs apheresis comparing the efficacy of OPS with its precursor cobe and other ahesia features (Table 3). These showed the collection of high quality MNC with low platelet and erythrocyte contamination.29-31 In non-simulated collection procedures for secondary treatment of patient safety, including a short collection time and a small volume of product, its patients received a series of procedures. With regard to secondary treatment, the product hematocrit was more critical because the MNC cell count.30 Unfortunately, the prediction of the number of MNC cells is difficult due to each patient's own peculiarity. Del Fante et al.'s study showed that the use of the MNC SPO procedure may have an average product volume 10% lower with possible benefit for subsequent irradiation and lower liquid load with low body weight.29 Table 3 MNC collection in non-simulated patients for secondary treatment to perform offline ECP or for donor lymphocyte infusion Notes: amixed anticoagulation with 5 Heparin IU/mL ACD-A; bCE for lymphocytes (CE- is based on the average number of target cells before and afteraphed); cCE for MNC. Abbreviations: ACD-A, anticoagulant solution citrate dextrose A, CE, Collection Efficiency; CMNC, a continuous collection of mononuclear cells; DDI, infusion of donor lymphocytes; ECP, extracorporeal photopheresis; MNC, mononuclear cells; Untested; OPS, Spectra Optia®; TPE, therapeutic plasma exchange. An interesting strategy to reduce ACD consumption is the initial 1:12 to 1:20 increase in MNC procedures described by Del Fante et al The operator has modulated the ACD-A ratio based on the number of platelets, Clotting status, and comorbidities.29 An anticoagulation proportion sufficient to maintain adhesion platelet function is essential if differentiation of dendritic cells in ECP occurs by transient engagement of monocytes with device-joining activated platelets and their ligands.32 To what extent the anticoagulation regimen in ECP may influence the clinical outcome is unknown. For ecp, no WBC threshold is set. Nevertheless, up to 1.0x1010 total lymphocytes with high purity seem possible to be collected within a procedure.29,30,33,34 The importance of product purity is described only for patients with an indication for ECP due to lung involvement after lung involvement after transplantation.29 Punzel et al. compared the MNC procedure with the CMNC procedure in the context of collecting lymphocytes for donor lymphocyte infusion.34 It has been observed that the CMNC procedure takes a shorter treatment time and a lower product volume but a higher number of hematocrit and platelet in the product. With lower centrifugal forces using a packing factor of 4.0 instead of 4.5 in cmnc, it is possible to reduce platelet contamination in the cell product while keeping all target cells at the same yield.34 Nevertheless, we do not prefer the use of the CMNC procedure in the offline ECP setting because a significantly higher hematocrit in the product may hinder subsequent treatment with UVA illumination. CMNC with small product volume, shorter procedure time, and thrombocyte savings may be the preferred setting for the collection of peripheral blood stem cells. Brosig et al. have made an important contribution to their analysis of ECP methods in offline33. Side effects were not observed in the ECP methods tested. For offline methods, calcium was substituted as needed by anticoagulant with ACD-A, effectively preventing the most common side effect.33 The report on the preparation of mini buffy buffy for adult patients who can't receive an online ECP or an offline ECP seems interesting. Whole blood separation with bone marrow treatment program of the OPS ahesia system in 1 group achieved a higher lymphocyte yield than the standard mini buffy coat preparation method with the fully automated Compomat G4 separator device. This technique, which has been examined in healthy whole blood donors, should be further studied in a clinical setting.27 With regard to AEs, only mild hypocalcemia in the MNC collection for ECP has been reported. The occurrence of mild hypocalcemia has been reported by study groups of Del Fante et al. and Schulz and others.29,30 white blood cell depletion (WBCCD) and platelet depletion (PLTD) For WBCCD depletion, different procedures performing with SPO are available. It is possible to use the granulocyte collection procedure (PMN) as well as the WBCCD procedure. There are no substantial differences between the two methods; only the default settings are different. The set to be used is the SPO (IDL-Set) intermediate density layer. This set is particularly developed to collect a large amount of CWB or platelets in a short period of time. Clinical manifestation of leukostasis due to hyperleukocytosis in acute lymphoblastic (ALL) or acute myeloid (LAM) procedures could motivate physicians to perform leukocytapheresis procedures. The depletion of the WBC in the symptomatic ALE with wbc counts >100x109/L or ALL with wbc count >400x109/L is recommended with respect to ASFA guidelines as category II grade 1b.5 In a PubMed research, we found only 3 reports related to WBCCD or PLTD with SPO.30,35,36 WBCCD performance with SPO was first described by Schulz and others.30 They treated 5 patients with leukostase due to AML and performed a total of 8 PMN treatments. Double the total volume of blood, a product equivalent to one-fifth of the total blood volume or no more than 300 minutes, were treated. ACD-A was used for anticoagulation in a 12:1 report (whole blood: ACD-A). Depending on the clinical situation, the report has been adapted. It is important to note that, under normal operating conditions, the extracorporeal volume of the IDL-Set does not exceed 253 mL. Under alarm conditions, extracorporeal volume can increase to 297 mL. Therefore, it is simple to overcome the permitted extracorporeal volume limit. However, in addition to mild hypocalcemia, no AE, including hypotonia, was observed.30,36 A collection efficiency of 47.3% ± 7.4% and platelet attrition of 32.8% ± 2.8% was achieved, but the authors admitted that a very variable number (up to 50% of cells was mobilized into the bloodstream during leukocytapheresis. Despite a low decrease in WBC count (22%), Cline et al. described a high volume of collected cells, 3.51x1010 WBC per volume of treated blood using the PMN procedure.36 In the transfusion medicine apheresis unit at Jena University Hospital, we treated 7 patients with hyperleukocytosis; AML and 1 case was with ALL. We used the WBCCD procedure exclusively. The number of WBCCs was reduced by an average of 27% (5% to 55%) relative to the baseline. All wbcd procedures were completed as planned. Despite a high extracorporeal volume at the end of the WBCCD, there was no EA. Only 1 case of PLTD in a pregnant woman at 35 weeks gestation was reported.35 thrombocytapheresis procedures were performed within 2 weeks. An average of 1.3 ± 0.3 total blood volume was treated, and collection efficiency was 50.6% ± 2.6% while preserving the level of leukocytes. No AE and no other pregnancy complications have been documented. Conclusion There is little data available on the clinical risks and benefits of OPH ahesia treatments. All published data showed clinical benefits, which were more pronounced than clinical risks. The OPS works efficiently and achieves high plasma extraction rates, even with a low input rate. TPE procedures are safe and well tolerated, with AEs occurring in 5%-7% of cases.14,15 AEs generally include reactions to replacement fluid, citrate, and hypotension. Platelet loss using moderate rates of input flow is no longer a clinical risk, even in patients with TTP. Evaluation of published clinical results results were adduced in effective removal of pathological substances like IgG-antibodies, and triglycerides. SPO SPD-LA treatments achieved an effective removal of lipoproteins such as LDL cholesterol and lipoprotein up to 59% and 64%, respectively, compared to baseline.21,22 In particular, no cardiac events were observed. In SPO SPD-IA, there is almost no published clinical data on the results. However, the described reduction in IgG appears satisfactory. To date, in plasma-based procedures, the optimal use of citrate has not been evaluated. The use of ACD-A in TPE differs between apheresis centres from 3.11 mL/min to 6.61 mL/min. Nevertheless, bleeding or clotting has not been observed.8-14,17 PLTD, PMN, and WBCCD procedures are suitable for rapid blood cell depletion. Assessing the effectiveness of collection is difficult, due to the mobilization of CWB in the bloodstream in WBCCD or PMN procedures. In addition, these procedures often exceed the permitted extracorporeal volume limit. Nevertheless, no severe AE was observed.30,36 Described PLTD procedures could reduce platelet in a pregnant woman with thrombocythemia without AEs.35 RBCD/RBCX protocols in SPO are comparable to the RBCX hemodilution isovolemic protocol in Cobe Spectra.23 No significant difference was found. SPO-MNC procedures unsimulated patients showed high quality MNC collection with low platelet and erythrocyte contamination, and 10% lower product volume with possible benefit for subsequent illumination in ECP. Unfortunately, the efficiency values of the collection are not comparable. Apheresis centres used different calculation formulas.29,30,33,34 In cell-based OPS procedures, cell-based anticoagulation differs slightly between apheresis centres. Sweet AEs are reported. In general, for further evaluation of the OSS, more data on clinical outcomes is needed. Acknowledgements The authors thank Tom Westhauser for the excellent support in the arrangement of this review. Disclosure Silke Rummier works as a country reviewer for Terumo BCT, Inc. The authors do not report any other conflicts of interest in this work. References 1.Bramlage CP, Schroeder K, Bramlage P, et al. Predictors of complications in therapeutic plasma exchange. J Clin Apher. 2009;24(6):225-231. 2.Sanchez M, Guadilla J, Fiz N, Andia I. Plasma injections rich in ultrasound-guided platelets for the treatment of osteoarthritis of the hip. Rheumatology. 2012;51(1):144-150. 3.Pusey CD, Levy JB. Plasmapheresis in immunological kidney disease. Blood purif. 2012;33(1-3):190-198. 4.Paton E, Baldwin IC. 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