

Patient Blood Management: Challenges and Approaches to Blood Health

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Summary

Anaemia is among the most common global health problems. When taking into account related blood health disorders such as blood loss and coagulopathies, the picture becomes even broader: According to recent estimates, more than 3 billion people worldwide are affected. Particularly in perioperative settings, patients with anaemia face a greater need for blood transfusions and increased risk of postoperative complications. The concept of Patient Blood Management (PBM) addresses these issues by providing therapeutic measures aimed at preserving valuable resources and enhancing patient safety. PBM is a multidisciplinary, patient-centred approach to optimizing blood health and rests upon three pillars: (1) (preoperative) anaemia management, (2) minimization of blood loss, and (3) rational use of blood products. Since anaemia has multifactorial causes, it is essential to consider expanding the existing treatment paradigm, which predominantly focus on iron deficiency therapy alone. An intersectoral and interdisciplinary approach – particularly involving primary care providers – is crucial to sustainably enhancing patient safety, quality of care, and resource efficiency.

Challenges to blood health

The high prevalence of anaemia, insufficient implementation of perioperative anaemia management, and liberal han-

dling of blood transfusions pose significant challenges to health care systems worldwide. Global epidemiological data show that roughly 25% of the world's population is affected by anaemia. Approximately 1.9 billion people suffered from this disease in 2021 [1]. If other related blood health conditions like blood loss and coagulation disorders are factored in, the picture will become even broader: current estimates show that there are more than 3 billion people affected worldwide whose blood health is chronically or acutely impaired. This widespread blood health crisis has far-reaching effects on the global disease burden, the cost of health care, and the quality of life of those affected [2]. Anaemia is an especially important risk factor in the surgical context that can significantly affect the prognosis of patients undergoing surgery. Studies document that around 10–48 % of patients show signs of anaemia prior to surgery [3,4], which will be associated with significant risks if left untreated. A current study by Warner and Colleagues shows that anaemic patients (n = 1234) who undergo elective cardiac surgery have higher rates of postoperative acute renal failure (8 % vs. 18 %) and a prolonged stay in the hospital (5.8 days vs. 6.7 days) compared to non-anaemic patients [5].

There is currently a controversy as to whether the gender-specific haemoglobin cutoffs defined by the World Health Organisation (WHO) (men <13 g/dl and women <12 g/dl) are appropriate for defining anaemia. Earlier studies [6,7] as

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Keywords

Anaemia – Patient Safety – Perioperative Management

well as a more recent investigation by Netz et al. [8] suggest that adjusting the haemoglobin cutoff to <13 g/dl for women who are undergoing major surgical interventions is associated with a reduction in the need for transfusions and a lower rate of postoperative complications. Despite the known risks of allogeneic blood transfusions [9], anaemia not associated with a massive blood loss or life-threatening conditions is still frequently treated with red blood cell transfusion.

Early diagnostic evaluation, for instance by primary care physicians prior to the scheduled surgical intervention, combined with targeted causal therapy, can effectively reduce perioperative complications and significantly minimize the requirement for allogeneic blood transfusions. But this would urgently require a cross-sector treatment concept with clearly defined responsibilities and suitable reimbursement. The various aspects can be addressed and improved simultaneously through the targeted optimisation of blood health. Patient Blood Management (PBM) is a promising medical approach that focus on improving blood health in order to improve outcomes during the perioperative period. In spite of the WHO's call in recent years [2,10–12] for the comprehensive implementation of PBM, it has yet to be adequately put into practice [13–15].

What is Patient Blood Management?

PBM is a patient-oriented, evidence-based and multidisciplinary concept for optimising blood health [2] and increasing patient safety [16].

PBM measures are based on three pillars (Fig. 1) [17]:

1st Pillar: (Preoperative) anaemia management

2nd Pillar: Minimisation of blood loss

3rd Pillar: Rational use of blood products.

Pillar 1: (Preoperative) anaemia management

The first pillar of the PBM approach promotes patients' own haematopoiesis

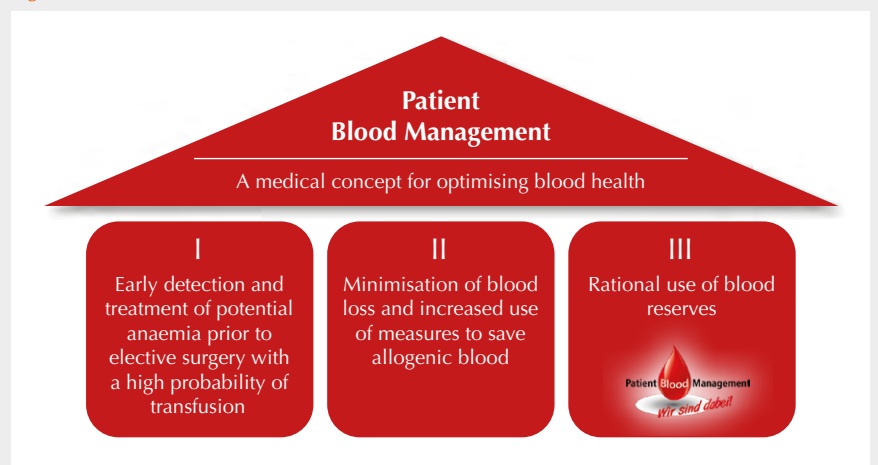
with the aim of raising the haemoglobin level in advance of planned surgical interventions. The focus is on the early diagnosis and treatment of preoperative anaemia. One of the main causes of anaemia is iron deficiency [18], which is generally easy to treat. Neef and colleagues showed that intravenous iron supplementation is most effective if it is administered at least ten days before surgery – ideally, treatment should be applied three to four weeks before surgery. This period of time enables an optimal rise in haemoglobin and allows for an additional iron supplementation if necessary [28]. In particular, various studies indicate that the correction of preoperative anaemia is associated with lower complication rates, a shorter stay in the hospital and a lower transfusion rate [4,19,20].

However, the latest studies show that a holistic concept for treating anaemia is necessary, and correcting iron deficiency only is not always sufficient. The causes of anaemia are multifactorial and, aside from iron deficiency, also include vitamin B12 and folic acid deficiency, chronic kidney disease, persistent inflammations, infections and, in rare cases, even genetic hemoglobinopathies (e.g. thalassemia, sickle cell anaemia) [21]. Furthermore, primary diseases such as primary osteomyelofibrosis, secondary bone marrow depression (e.g. after

radiation) and toxic influences (e.g. poisoning) can trigger anaemia [22–24]. Spahn et al. developed a multimodal therapeutic approach in which cardiac surgery patients with (iron deficiency) anaemia received a combination therapy consisting of intravenous iron, erythropoietin alpha (EPO), vitamin B12 and folic acid before surgery. Even though the combination therapy is theoretically accompanied by risks such as thromboembolic events, infections and other complications, these did not occur more frequently during the course of the study. This multidimensional treatment concept led to a reduction in the need for transfusions [25].

Early diagnosis and therapy of anaemia in case of elective surgery should ideally be commenced in the primary care setting. In Germany, there is a structural gap in primary care when it comes to preoperative anaemia: Standardised diagnostics do not usually take place, which is attributable to a lack of legal requirements, unclear responsibilities and insufficient reimbursement. General practitioners, who are often the first point of contact before planned surgery, rarely perform relevant blood tests to determine iron deficiency parameters, such as ferritin or transferrin saturation. Furthermore, communication structures across sectors are lacking, and PBM has yet to be integrated at the national

Figure 1



Concept of Patient Blood Management (source modified: <https://www.patientbloodmanagement.de>).

level. Additionally, knowledge of the clinical significance and implementation of PBM has not yet been sufficiently embedded in training, continuing education, or everyday practice.

The general practitioner will play a central role in the future, as they usually care for the patients on a long-term basis, knows their pre-existing conditions and often is the first to notice symptoms of anaemia such as fatigue, decline in performance, heart palpitations or dyspnoea under exertion. The physician will usually have the most lab values on hand in chronological order, which can be used to detect changes in haematologic parameters at an early stage. In addition to early diagnosis, measures that can be taken by the general practitioner also include clarification of the causes and initiation of targeted therapy. Depending on severity, therapy can take place directly in the GP's practice, e.g. oral or intravenous supplementations of iron, vitamin B12 or folic acid pre-

parations. This way, it can be ensured that the patients are optimally prepared for elective surgery. Close cooperation between the general practitioner and the treating hospital is also useful and desirable. Further clarification and treatment can be continued in a hospital equipped with interdisciplinary specialist departments, especially in more complex cases. Different forms of anaemia require different therapeutic approaches, ranging from dietary supplementation and erythropoiesis-stimulating agents to transfusion medicine and specialised treatment measures (Fig. 2).

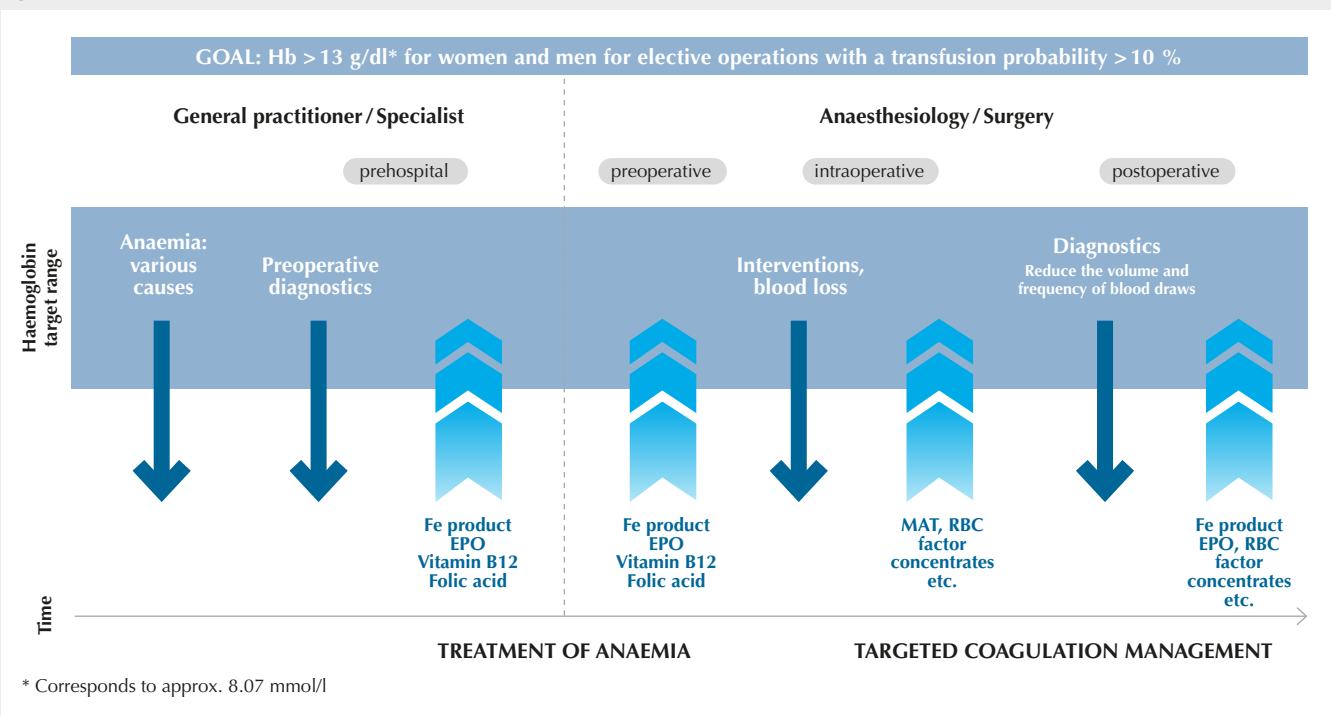
Pillar 2: Minimisation of blood loss

The second pillar is composed of measures to reduce avoidable blood loss with the aim of maintaining the body's own blood reserves. For example, diagnostic-related blood loss can be associated with an unfavorable clinical course [26,27]. Frequent blood draws

lead to a significant blood loss, which can reach up to 1,428 ml if the patient spends more than eleven days in the intensive care ward. Low-volume blood collection tubes and closed blood collection systems can significantly reduce the amount of blood loss needed for diagnostic purposes [28–30].

Another major aspect is a comprehensive coagulation management, which is supported by the careful correction of physiological parameters (body temperature, ionised calcium, pH value). The use of point-of-care diagnostics (POCD) is recommended for targeted therapy with coagulation factor concentrates, since it enables both a rapid and a bedside analysis [31]. Nonetheless, it must be kept in mind that POCD systems can have limitations despite their practical advantages. Their use is particularly valuable in situations in which rapid diagnostics are required – for example, when a central laboratory is not immediately available. The use of cell salvage

Figure 2



The ups and downs of haemoglobin. Influencing factors and potential patient blood management (PBM) measures in the perioperative context (source: own illustration).

EPO: Erythropoietin; RBCC: RBC concentrate; Fe product: Iron product; MAT: Machine autotransfusion.

also makes a major contribution towards reducing the need for allogenic blood transfusions. A recent meta-analysis [32] showed that the use of cell salvage in surgery patients not only significantly reduces the need for allogenic blood transfusions by 39 %, but is also associated with a 28 % reduction in the risk of postoperative infection and a reduction of hospitalisation time by 2.31 days.

The implementation of various blood-saving methods, such as the use of low-volume blood collection tubes, closed blood collection systems, cell salvage, POCD as well as comprehensive coagulation management significantly improve haemoglobin levels and secure valuable blood reserves for emergencies (Fig. 2).

Pillar 3: Rational use of blood products.

The third pillar of the PBM concept consists of the rational use of blood products in compliance with the guidelines. The primary goal of a transfusion of allogenic blood products is to ensure an adequate oxygen supply to the tissue in order to avoid complications resulting from acute severe anaemia. However, it must be carefully considered whether the potential benefit of the transfusion outweighs the risks it entails. This decision should always be made considering individual patient factors. With patients in stable haemodynamic condition, even low haemoglobin values under 8 g/dl can often be tolerated without clinically relevant symptoms. Despite extensive studies, a significant benefit of a liberal transfusion practice (haemoglobin target values between 9 and 11 g/dL) compared to a rational strategy (haemoglobin target values between 7 and 9 g/dL) has not yet been seen in most clinical situations [33–35]. Therefore, a preventive approach is preferable [36].

Conclusion

The early diagnosis and treatment of preoperative anaemia by general practitioners is crucial, ideally in cooperation with specialists or specialised anaemia outpatient clinics. Comprehensive anaemia

management which pays particular attention to iron, vitamin B12 and folic acid deficiencies makes a major contribution towards increasing patient safety, preventing perioperative complications and ensuring effective and sustainable patient care. An interdisciplinary approach is of central importance here. In individual cases with a short time frame, it should be considered whether to postpone elective surgery in order to enable adequate preoperative anaemia management. Just as important as interdisciplinary cooperation is the implementation of practicable measures in everyday primary care practice. These include structured recording of elective surgery, targeted requesting of relevant laboratory parameters (such as haemoglobin, reticulocyte-haemoglobin, ferritin and transferrin saturation) as well as cause-oriented therapy.

The integration of standardised processes – such as through short Standard Operating Procedures (SOPs), patient questionnaires or digital interfaces – also enables an efficient and permanent integration of preoperative anaemia management into the existing primary care processes.

Conflict of interest

KZ received fees for taking part in advisory board meetings from Haemonetics and CSL Vifor and received presentation fees from CSL Behring, Masimo, Pharmacosmos, Boston Scientific, Salus, iSEP, Edwards, HemoSonics and GE Healthcare. He is principal investigator in the EU-Horizon 2020 project ENVISION (intelligent digital plug and play tool for real-time monitoring of COVID19 patients and intelligent decision-making in intensive care units) and the Horizon Europe 2021 project COVend (biomarkers and AI-assisted FX06 therapy to prevent mild and moderately severe stages of COVID19 from progressing to severe stages) and partner for the EU Horizon 2023 project EDiHTA. KZ is CEO of the Christoph Lohfert Foundation and also heads the Stiftung für Gesundheit, Patientensicherheit und Patient Blood Management (PBM Foundation).

SC received fees for writing articles from the publishing house Thieme Verlag.

PM and his clinic received funding for research projects and clinical trials from the German Federal Ministry of Education and Research (BMBF, 01KG1815), the German Federal Ministry of Health (BMG, ZMVI1 2520DAT10E, ZMII2-2523FEP50A), CSL Behring and the German Research Foundation (DFG, ME 3559/1-1, ME 3559/3-1, ME 6094/3-2).

PM received presentation fees and reimbursement of travel expenses from CSL Behring, CSL Vifor, Pharmacosmos and Werfen. PM is a board member of the board of directors of the Stiftung für Gesundheit, Patientensicherheit und Patient Blood Management (PBM Foundation), the Network for the Advancement of Patient Blood Management, Haemostasis and Thrombosis (NATA) and a member of the standing work group of the scientific advisory board "Cross-sectional Guidelines for Therapy with Blood Components and Plasma Derivatives".

DM does not have any conflict of interest.

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