Transurethral Resection of the Prostate and Pre-Operative Anemia, in Men Older Than 60 Years

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Abstract:

Background/Aim: In their everyday clinical practice, anesthesiologists often deal with patients suffering from preoperative anaemia. The objective of the study was to determine the incidence and influence of preoperative anemia in the perioperative course in patients undergoing transurethral resection of the prostate.

Methods: The clinical study included 206 patients, who were divided into two groups. In the first group there were the patients with preoperative anemia (hemoglobin level less than 13g/dL), and the control group with the patients without preoperative anemia (hemoglobin level $\geq 13g/dL$). The study was conducted at the Clinical Center of Vojvodina from June, 2011 to July, 2012. We analyzed the following: demographic factors, the ones related to the surgical procedure and the factors associated with anemia. Within the perioperative course we analyzed: the perioperative complications and the length of hospitalization.

Results: From the total of 206 patients, 54 (26.21%) were the patients from the group with preoperative anemia (hemoglobin level <13g/dL), and the control group of 152 (73.79%) patients without anemia (hemoglobin level $\geq 13g/dL$). The number of patients with perioperative complications was 15 (27.78%) from the group with anemia and 22 (14.47%) from the group without anemia. The complications were significantly more common in patients from the group with the preoperative anemia in comparison with the ones without anemia (x2=4.786, p<0.05). In the patients from the group with preoperative anemia, the average length of hospitalization was 6.44±3.489 (5-29) days whereas in the patients from the control group it was 5.61±1.51 (3-15) days. The patients from the group with anemia had significantly longer hospital stay, in comparison with the ones from the group without anemia (t=2.386, p<0.05).

Conclusion: The incidence of preoperative anemia in our clinical study was 26.21%. The examined patients with preoperative anemia had perioperative complications far more frequently and substantially longer hospital stay.

Keywords: prostate, surgery, anemia, perioperative course.

1. INTRODUCTION

In their everyday clinical practice, anaesthesiologists often deal with patients with pre-operative anemia that is manifested with lower hemoglobin levels. Its significance and influence on perioperative morbidity and mortality rates are the subject of numerous clinical studies (1-3). The anemia evaluation is especially important in elderly patients (over 65 years of age), in whom preoperative anemia is associated with increased morbidity and mortality, for this group of patients often suffers from cardiovascular, respiratory, cerebrovascular and other co-existing diseases (4-7). Several factors link the age of the patient and anemia as the risk factors for perioperative morbidity and mortality. The cardiac reserve in elderly patients is reduced (8), which can decrease the regular compensatory mechanisms due to lower hemoglobin levels (9,10). Comorbidities, especially the coronary artery disease, in elderly patients reduce their tolerance to anemia (11). In addition, the preoperative anemia is quite significant for being a strong predictor variable for the transfusion of blood and its components (12-14). Transurethral resection of the prostate (TURP) is the second most common surgical procedure in men older than 65 years. The indications for surgical treatment are as follows: benign prostatic hyperplasia (BPH) and prostate cancer, at an advanced stage, with urine retention, where palliative TURP, as a method of treatment, is applied. From the anesthesiologist's point of view, the specificity of the patients undergoing the transurethral resection of the prostate is that they are the elderly patients, most commonly the ones with multiple comorbidities,

cardiovascular, respiratory and cerebrovascular ones, as well as the fact that the above mentioned interventions are most commonly associated with intraoperative and postoperative bleeding (15,16).

We have decided to undertake the following clinical study, after taking into consideration the above mentioned specificities of the population undergoing the TURP treatment method and, due to the fact that there have been no published studies dealing with preoperative anemia in this particular group of patients recently.

The aim of this study was to determine the incidence and influence of preoperative anemia on perioperative course (perioperative complications and length of hospitalization) in patients undergoing transurethral resection of the prostate.

2. METHODS

A retrospective clinical study was conducted at the Clinic of Anesthesia and Intensive therapy and the Clinic of Urology of the Clinical Centre of Vojvodina, Novi Sad from 1st June, 2011 to 30th June, 2012. During the above mentioned period the total of 229 patients underwent a transurethral resection of the prostate. Due to the lack of the complete medical documentation, 23 (10.04%) patients were excluded from the study. Therefore, 206 (89.96%) patients with the complete medical documentation entered the study. The data were collected from the existing medical documentation (case history, anaesthesiological data, therapy lists, decursus, releasing forms). This study was approved by the Ethics Comittee of the Clinical Centre of Vojvodina. In the course of the clinical study, the following parameters were analyzed: 1) demographic factors, 2) surgical procedure related factors 3) anemail related factors and 4) within the perioperative course we analyzed the perioperativne complications and the length of hospitalization. As for the demographic factors, the following ones were analyzed: the age of the patients, general health assessment according to the ASA classification (American Society of Anaesthesiologists, physical status class) (17), and comorbidities. As for the surgical procedure related factors, the following was analyzed: the main disorder as an indication for the TURP, the type of anaesthesia and surgical operation time. Further factors in relation to anaemail were analyzed: preoperative hemoglobin (Hb) levels in blood, percentage decrease in the Hb level on postoperative day 1 (POD1-Hb change) which indicates the blood loss during the surgical procedure, the mean value of postoperative Hb until the postoperative day 3 and the blood transfusion administration quantity during the hospitalisation (18). The criteria for blood transfusion used at our institution was the transfusion trigger, $Hb \le 10 \text{ g/dL}$ (liberal transfusion strategy) (19). Within the perioperative course, the following was analyzed: perioperative complications (intraoperative and postoperative until the hospital release, new occurrence of perioperative outcomes), noted in the medical documentation and the length of hospitalization. The perioperative complications that were being examined were the following, systemic ones: cardiac, including ischemic cardiac complications (new occurrence of chest pain, electrocardiogram changes, or cardiac enzyme changes), myocardial infarction, dysrhythmias, or heart failures. Respiratory complications included pulmonary edema, tracheal re-intubation, pulmonary consolidation on the chest X-ray, or pleural effusion. Renal complications were defined as the serum creatinine elevation. Neurological complications were defined as the new occurrence of transient ischemic attack or stroke, delirium, or confusion. Infection required a documentation of the positive culture (20). Intrahospital mortality rate was analyzed, as well.

We defined preoperative anemia as a Hb level lower than 13 g/dL in men according to the World Health Organization (21). Hb levels were measured within 30 days before the planned prostate surgical procedure. According to the preoperative Hb levels, the 206 examined patients were divided into two groups. 54 patients with preoperative anemia were in the first group (Hb level < 13 g/dL), whereas the control group consisted of 152 patients with no preoperative anemia (Hb level \geq 13 g/dL).

The examined patients were given general or regional anaesthesia depending on the general health of the patient. During statistical data processing, standard statistical research methods were used (descriptive statistics and frequency distribution). Numerical data were presented by medium arithmetic values and standard deviation, a comparison was done by the t- test. For the frequency of the observed parameters difference (distribution) hypotheses testing, x^2 test was used. Univariate and multivariate logistic regression analyses were performed to identify the risk factors influencing the perioperative course. A *p* value of <0.05 was considered statistically significant.

3. RESULTS

From the total of 206 patients included in the clinical study, 54 (26.21%) ones had preoperative anemia (a group of patients with Hb < 13g/dL) and 152 (73.79%) patients did not have preoperative anemia (a group of patients with Hb \geq 13g/dL). The mean value of preoperative Hb in patients with anemia was 11.72 ± 1.03 g /dL and in the control group it amounted to 14.49 ± 0.94 g /dL. Table 1 compares the demographic variables and the ones associated with surgical procedure among the patients with anemia and the ones without anemia. The average age of the patients was 71.12 ± 8.35 years. In comparison with the patients without anemia, the ones with anemia were significantly older (t= 3.626, p <0.01). As for the ASA status, there is a significantly larger number of ASA class 3 patients, in the group of patients with anemia, in comparison with the patients from the group without anemia ($x^2=10.690$, p<0.05). The incidence of comorbidities in the examined patients was 81.1% and it was similar in both groups of patients (x^2 =1,699, p>0.05). As for the underlying disease, for the indication of the transure thral resection of the prostate, the patients from the group with anaemia had significantly more malignant prostate disease incidence than the ones from the group without anemia $(x^2=26,694, p<0.01)$. As for the type of anaesthesia and surgical operation time there was no significant difference between the groups (Table 1). Table 2 shows anemia associated factors. Transfusion was received by significantly more patients from the group with anemia in comparison with the ones without anemia ($x^2=31.323$, p<0.01). There was no statistically significant difference in postoperative decrease in Hb values between the groups. As far as the mean value of postoperative Hb is concerned, significantly larger number of patients, from the group with anemia, had a postoperative Hb <13g/dl (postoperative anemia) in comparison with the ones without anemia (96.3% vs 41.4%). The number of patients who had perioperative complications was 15 (27.78%) from the group with anemia (Hb <13g/dL) and 22 (14.47%) from the group without anemia (Hb >13g/dL). The complications were significantly more common in patients from the group with preoperative anemia in comparison with the ones from the group without anemia ($x^2 = 4.786$, p<0.05).

General variables	No anemia (n=152)	Anemia (n=54)
Age (years), $\overline{X} \pm SD$	69,89±8,57	74,56±6,65
ASA class [*] , n (%)		
1-2	65 (42,8)	10 (18,5)
3	85 (55,9)	42 (77,8)
4	2 (1,3)	2 (3,7)
Comorbidity, n (%)		
Yes	120 (78,9)	47 (87,0)
No	32 (21,1)	7 (13,0)
Main disorder, n(%)		
BPH	136 (89,5)	31 (57,4)
Malignant tumor	16 (10,5)	23 (42,6)
Type of anaesthesia, n(%)		
General	43 (28,3)	21 (38,9)
Regional	109 (71,7)	33 (61,1)
Surgical operation time (min), $\overline{X} \pm SD$	53,99±31,39	60,28±21,33

Table1.	Demografic	factors and	d surgical	procedure	related fact	ors
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n-number of patients; SD- standard deviation; ASA- American Society of Anaesthesiologists; *ASA scores: 1 = healthy patient, 2 = mild systemic disease but no functional limitations, 3 = severe systemic disease with definite functional limitations, 4 = severe systemic disease that is a constant threat to life, 5 = moribund patient unlikely to survive 24h with or without an operation; BPH- Benign Prostatic Hyperplasia;

Variable	No anemia (n=152) n (%)	Anemia (n=54) n (%)	p value
Perioperative transfusion, yes	8 (5,3)	19 (35,2)	0,000
POD1- Hb change [*]			
Change ≤20%	119 (78,3)	27 (50)	0,332
Change >20%	12 (7,9)	5 (9,3)	
Postoperative Hb≥13g/dL	89 (58,9)	2 (3,7)	0,000
Postoperative Hb<13g/dL	63 (41,4)	52 (96,3)	

POD1-Hb change - percentage drop in the Hb level on postoperative day 1; Hb- hemoglobin

Table 3 shows the incidence of individual complications in relation to the total number of patients in both groups. The table shows that cardiac complications were the most common ones in both groups of patients. The patients from the group with anemia had significantly more cardiac complications in comparison with the ones from the group without anemia ($x^2=7.594$, p <0.05).

Complications	No anaemia (n=152) n (%)	Anemia (n=54) n (%)
Cardiac	16 (10,53)	14 (25,93)
Respiratory	2 (1,32)	1 (1,85)
Neurological	2 (1,32)	1 (1,85)
Renal	1 (0,66)	1 (1,85)
Infection	4 (2,63)	0 (0)
Death	1 (0,66)	0 (0)
Total number of patients with		
complications	22 (14,47)	15 (27,78)
0 Complication	130 (85,53)	39 (72,22)
1 Complication	18 (11,84)	13 (24,08)
≥ Complications	4 (2,63)	2 (3,70)

Table3. The frequency of complications related to the total number of patients in both groups

In the patients from the group with preoperative anemia, the average length of hospitalization was 6.44 ± 3.489 (5-29) days whereas in the patients from the control group it was 5.61 ± 1.51 (3-15) days. The patients from the group with anemia had significantly longer hospital stay, in comparison with the ones from the group without anemia (t = 2.386, p <0.05).

In order to compare the predictor parameters, the examined variables were tested by the logistic regression analysis where the dependent variable were the perioperative complications as well as the length of hospitalization (hospitalization longer than five days). The univariate logistic regression analysis included the following independent variables: the age of the patients, comorbidities, ASA status, preoperative Hb, transfusion, POD1-Hb changes and postoperative Hb. In addition, the multivariate logistic regression analysis included the variables associated with the perioperative complications as well as the length of hospitalization, according to the univariate analysis.

In Table 4 it is shown that there is a statistically significant univariate connection between the preoperative Hb<13g/dL and postoperative Hb<13g/dL with perioperative complications. According to the results of the multivariate regression analysis the postoperative Hb<13g/dL is an independent predictor of the perioperative complications incidence (Table 5). The patients who had the postoperative anemia had 2.5 times greater chance of perioperative complications incidence.

Variable	Number	Complication n (%)	p value
Total number of patients	206	37 (17,96)	
Age (years)			
<65	41	5 (12,1)	0,366
≥65	165	32 (19,4)	
Comorbidity			
Yes	167	33 (19,8)	0,245
No	39	4 (10,3)	
ASA class			
<3	75	11 (14,7)	0,451
≥3	131	26 (19,8)	
Preoperative Hb			
<13g/dL	54	15 (27,8)	0,038
≥13g/dL	152	22 (14,5)	
Perioperative transfusion			
Yes	27	8 (29,6)	0,107
No	179	29 (16,2)	
POD1-Hb change*			
Change ≤20%	146	23 (15,8)	0,448
Change >20%	17	4 (23,5)	
Postoperative Hb			
<13g/dL	111	28 (24,6)	0,010
≥13g/dL	95	9 (9,9)	بل ب

Table4. Univariate analysis of risk factors for perioperative complications

n- number of patients; ASA- American Society of Anaesthesiologists; Hb- hemoglobin; *POD1-Hb* change – percentage drop in the Hb level on postoperative day 1

Transurethral Resection of the Prostate and Pre-Operative Anemia, in Men Older Than 60 Years

Table5. Multivariate analysis of risk factors for perioperative complication	Table5. Mt	ultivariate analv	sis of risk	factors for	perioperative	complications
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Variable	p value	OR	95% CI
Preoperative Hb < 13g/dL	0,384	1,451	0,628-3,352
Postoperative Hb < 13g/dL	0,046	2,505	1,016-6,177

OR - Odds Ratio; CI - Confidence Interval; Hb- hemoglobin

In Table 6 it is shown that there is a statistically significant univariate connection between the preoperative Hb<13g/dL, transfusion and postoperative Hb<13g/dL with the length of hospitalization. According to the results of the multivariate analysis, in predicting the length of hospitalization as the dependent variable, there is not any predictor that provides a unique, statistically significant, contribution to the model (p>0.05) (Table 7).

Variable	Number	Length of stay >5 days n (%)	p value
Total number of patients	206	75 (36,4)	
Age (years)			
<65	41	10 (24,4)	0,102
≥65	165	65 (39,4)	
Comorbidity			
Yes	167	61 (36,5)	1,000
No	39	14 (35,9)	
ASA class			
<3	75	25 (33,3)	0,548
≥3	131	50 (38,2)	
Preoperative Hb			
<13g/dL	54	27 (50,0)	0,021
$\geq 13g/dL$	152	48 (31,6)	
Perioperative transfusion			
Yes	27	16 (59,3)	0,010
No	179	59 (33,0)	
POD1-Hb change [*]			
Change ≤20%	146	47 (32,2)	0,279
Change >20%	17	8 (47,1)	
Postoperative Hb			
<13g/dL	111	49 (43,0)	0,041
$\geq 13 \text{g/dL}$	95	26 (28,6)	

Table6. Univariate analysis of risk factors for the length of hospitalization

n- number of patients; ASA- American Society of Anaesthesiologists; Hb- hemoglobin; POD1-Hb change – percentage drop in the Hb level on postoperative day 1

 Table7. Multivariate analysis of risk factors for the length of hospitalization

Variable	p value	OR	95% CI
Preoperative Hb < 13g/dL	0,302	1,489	0,699-3,174
Perioperative transfusion	0,109	2,106	0,846-2,44
Postoperative Hb < 13g/dL	0,430	1,320	0,622-2,634

OR - Odds Ratio; CI - Confidence Interval; Hb- hemoglobin

4. DISCUSSION

Our clinical study, that included the patients who underwent the transurethral resection of the prostate between June 2011 and July 2012 showed that the preoperative anemia was a relatively common state which is significantly in connection with the perioperative complications and the length of hospitalization. The prognostic value of preoperative anaemail in surgical procedures has been examined in many populations of the patients in both cardiac and non-cardiac surgery. In addition, numerous clinical studies, which included a relatively small number of patients who had undergone a particular type of surgical procedure have been conducted (18, 22). These examinations have shown that anemia is an independent risk factor for morbidity and mortality in surgical patients (3, 4, 23), which is the reason why it should be considered a serious medical condition that can be treated (3).

In one of the biggest observational studies (Wu et al.), which involved 310311 patients over 65 years who had undergone non-cardiac surgical procedure, it was concluded that 42.8% of them had preoperative anemia (4). According to the *Beatlie* et al. study, which involved 7679 patients over 18 years, who had also undergone non-cardiac surgical procedure, 39.8% of the men had preoperative

anemia that was defined as Hb<13g/dL in men (1). The results of our study involving men undergone transurethral resection of the prostate, have shown that the frequency of preoperative anemia is a little lower in relation to other studies and it is 26.21%, and is defined as Hb<13g/dL value.

Wu et al. demonstrate that preoperative anemia, even when discrepancies from physiological values of hematocrits are minimal, is linked to the increase in risk of postoperative mortalities and cardiac complications. The results of this study on a large sample of patients, mostly men older than 65 years, show that the increased cardiac complications and mortality risk is in proportion to the degree of anemia (4). Our study which included patients of similar demographic features also showed that patients with anemia had statistically significantly more cardiac complications in comparison with the group without anemia. As for the mortality rate in our study, there was one case of death in a patient who did not have anemia. The reason why our result is different in relation to other studies on mortality is likely due to the fact that we examined intra-hospital mortality rate only, and it is a small sample.

The study by *Dunne* et al., which involved patients of similar demographic features as ours, has examined the influence of preoperative anemia on the postoperative outcome in non-cardiac surgeries. It involved about 6301 patients, 95% of whom were men of the average age 61 ± 13 and the average ASA status of (2.6 ± 0.7) . 33.9% of them had preoperative anemia, anemic patients were administered five times as much blood in comparison to non-anemic ones (24). In our study preoperatively anemic patients received significantly more blood than the non anemic ones (x^2 = 31.323, p <0.01). In the study by *Dunne* et al. it has been shown that the low preoperative hematocrit level and blood transfusion administration are significant independent predictors of the length of hospitalization (24). Our clinical study found that there is a statistically significant difference in the length of hospitalization between the preoperative anemic and non-anemic patients undergoing transurethral resection of the prostate. It is shown that there is a statistically significant univariate association between the preoperative Hb<13g/dL, transfusion and the postoperative Hb<13g/dL in relation to the length of hospitalization.

The results of the *Beattie* et al. study, have shown that the proportion of blood administration is three times greater (30.4% vs. 10.6%) in anemic patients than in the ones who had no preoperative anemia (1). In addition, the results of our examination tests are in accordance with the above mentioned ones. We showed that the blood transfusion was applied in 35.2% patients with preoperative anemia, in relation to the 5.3% of the ones who were not anemic prior to their surgical procedure.

However, there are several limitations of our clinical study that should be taken into consideration when interpreting the one. First, this is a retrospective clinical study. Furthermore, the fact that only intra-hospital morbidity and mortality rates were examined is also considered a disadvantage. Then, the fact that no intraoperative blood loss was recorded is also considered to be the shortage of our study. However, in the TURP, it is difficult to assess intraoperative blood loss due to the application of large amounts of irrigation fluid during the surgical procedure. For these reasons, in practice, it is found that frequent post-operative measurement of Hb would be a useful indicator of blood loss, which was used in our study, as well. Despite the above mentioned limitations the results of our study correspond to the results of the previous studies.

Considering the incidence of preoperative anemia and its influence on the perioperative course, which was established in the group of the examined patients, we believe that it is necessary to determine the values of hemoglobin at least 28 days before the scheduled surgery of the prostate, which would enable a timely anemia correction because the hemoglobin level determination is a rather simple and generally quite available test analysis, and anemail treatment is not expensive in many situations. Within a preoperative preparation, anemail etiology should be determined and its treatment should start before the scheduled surgical procedure (3). There are diverse therapeutic options depending on the etiology, such as iron therapy, erythropoietin therapy or autologous transfusion administration such is preoperative autologous blood donation (3,25,26). This would significantly reduce perioperative complications and the length of hospitalization. So, perioperative alogenous blood transfusion administration that is by itself linked to numerous complications would be thus reduced (26, 27). Due to the limitations of our study, further prospective study in larger sample of patients is necessary.

5. CONCLUSION

In our clinical study, the incidence of preoperative anemia was 26.21%. The results of our study showed that the examined patients who had preoperative anemia had significantly more perioperative complications and increased length of hospitalization. The cardiovascular complications were the most common ones in both groups of patients.

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TRANSPARENCY DECLARATIONS

Competing interests: none to declare

Transurethral resection of the prostate = TURP

Benign prostatic hyperplasia = BPH

American Society of Anaesthesiologists = ASA classification

Hemoglobin = Hb

Hb level on postoperative day 1= POD1-Hb change

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