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## A transfusion-independent method for treating ICU-associated anemia

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## Keywords

Anemia, blood substitutes, blood transfusion, erythropoietin, intensive care unit, transfusion medicine

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## Comments

This study is admirable for its design and questioning of a common clinical scenario. It suggests a practical, clinically available alternative to blood transfusion in this 'at risk' population which would avoid the hazards of blood transfusion. The authors, using the results from the study, were able to suggest reasons for the anemia seen in ICU patients which need further investigation to fully elucidate the pathophysiology involved. Note however, that in the method, the authors did vary from routine clinical practice within the UK, by giving all enrolled patients oral iron from the start of the study, or as soon as bowel sounds were present.

## Introduction

Much recent interest has been focused on the transfusion requirements of intensive care unit (ICU) patients and how best to reduce patient exposure to unnecessary transfusions (and so avoid the associated risks). Previous work has identified the ICU population as having a transfusion requirement of 2-4 units/week. ICU patients, particularly those with sepsis, appear unable to mount an erythropoietic response to compensate for their anemia. It is uncertain whether this is as a result of inappropriately low erythropoietin production for the hemoglobin level or a failure to respond to endogenous erythropoietin.

## Aims

This study aimed to determine whether the administration of recombinant human erythropoietin (rHuEPO) to critically ill patients would reduce the requirements for red blood cell (RBC) transfusions.

## Methods

The study was a prospective, randomized, double-blind, placebo-controlled trial carried out in three tertiary care ICUs and studied a total of 160 patients (80 in each of the placebo and rHuEPO groups). Patients who fulfilled the study criteria were randomized and entered the trial on day 3 of their ICU admission. rHuEPO was administered subcutaneously at a dose of 300 units/kg for a total of 5 days. Subsequently, the study drug was administered on alternate days for a minimum of 2 weeks, or until ICU discharge. The rHuEPO was temporarily withheld when the hematocrit (Hct) reached  $> 38\%$  and restarted when the Hct fell below  $38\%$ . Outcomes examined were cumulative blood transfusion requirements during the study period and transfusion independence between study days 8 and 42.

## Results

Of 1778 patients admitted to the three ICUs over the study period, 329 were eligible but only 160 were enrolled (most commonly due to refusal to consent). The study populations were well matched and did not differ from the patients who did not consent to participate in the study. Cumulative transfusion requirements for the rHuEPO group were significantly less than the placebo group (166 total units in the rHuEPO group versus 305 total units for the placebo group). Each patient received a mean of  $8.3 \pm 4.5$  doses of rHuEPO. When transfusion independence was examined, 45% of patients in the rHuEPO group either received a blood transfusion between day 8 and 42 or died before study day 42, compared to 55% patients in the placebo group. There was no significant difference in the two groups in mortality or adverse events.

## Discussion

The authors conclude that administration of rHuEPO to ICU patients resulted in a significant reduction in blood transfusion requirements. They also concluded that the ability of this ICU population to respond to rHuEPO may indicate that the anemia seen in this population is due to a blunted erythropoietic response, as well as an impaired ability to respond to endogenous erythropoietin. They suggested that further studies were needed to identify the potential benefits of avoiding RBC transfusion.

## References

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