Literature List – Red Blood Cells

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General

**Jongbloed W et al. (2018)**
Unstable haemoglobin variant Hb Leiden is detected on Sysmex XN-Series analysers
Clin Chem Lab Med; 56(9): e249

What we see as the essence: Case study with an abnormal WDF scattergram observed on the XN-9000 causing the flag "WBC abnormal scattergram" which highlighted the inability to separate the white blood cell population. Further genetic analysis confirmed an unstable haemoglobin.

**Arbiol-Roca A et al. (2018)**
Reference intervals for a complete blood count on an automated haematology analyser Sysmex XN in healthy adults from the southern metropolitan area of Barcelona.
EJIFCC; 29(1): 48
Free online: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5949618/

What we see as the essence: The aim of the study was to establish reference intervals for CBC, DIFF and reticulocytes for a Spanish population. Significant gender differences were found for RBC, PLT, HCT and HGB.

**La Gioia A et al. (2018)**
Short preheating at 41°C leads to a red blood cells count comparable to that in RET channel of Sysmex analysers in samples showing cold agglutination
J Clin Pathol; 71(8): 729
http://jcp.bmj.com/content/early/2018/03/13/jclinpath-2017-204954.long

What we see as the essence: Detailed evaluation of the Ret channel in terms of resolving cold-agglutinins. It provides comparable results like traditional manual 37°C/2h pretreatment and can therefore be recommended to shorten TAT.

**Huisjes R et al. (2017)**
Digital microscopy as a screening tool for the diagnosis of hereditary hemolytic anemia.
Int J Lab Hematol; 40(2): 159

What we see as the essence: Advanced RBC Morphology from CellaVision was used to characterise hereditary haemolytic anaemia. Cutoffs were reported for several RBC abnormalities and showed a high sensitivity and specificity for detection of different conditions.
**Tailor H et al. (2017)**
Evaluation of the Sysmex XN-550, a Novel Compact Haematology analyser from the XN-L® series, compared to the XN-20 system.
Int J Lab Hematol; 39(6):585


**What we see as the essence:** Samples from adult patients (N=202) were measured on the XN-550 and compared with an XN-20. Good correlations and low bias was observed for all parameters except for BASO%. PLT-O from the XN-550 showed no significant bias compared to PLT-F from the XN-20.

**Jo SY et al. (2017)**
Performance evaluation of recently launched Sysmex XN-550 Automatic Hematology Analyzer.
Int J Lab Hematol 39(1):e4


**What we see as the essence:** The XN-550 showed a good analytical performance and strong correlation with XE-2100 and XN-3000 analysers for routine CBC parameters.

**Cao J et al. (2017)**
Establishing a Stand-Alone Laboratory Dedicated to the Care of Patients With Ebola Virus Disease.
Lab Med; 48(2): 188

[https://doi.org/10.1093/labmed/lmw072](https://doi.org/10.1093/labmed/lmw072)

**What we see as the essence:** The pocH-100i was used in a laboratory dedicated to detection of Ebola virus disease. Its accuracy was verified by comparison with the XE-2100 in the main laboratory, and its precision and reportable range were also consistent with Sysmex's claims.

**Egele A et al. (2016)**
Classification of several morphological red blood cell abnormalities by DM96 digital imaging.
Int J Lab Hematol 38(5):e98


**What we see as the essence:** The authors report the cutoff values for most of the RBC abnormalities that can be detected by the Advanced RBC Morphology software.
Tamigniau A et al. (2017)
From XE-2100 to XN-9000, from SIS Standard to GFHC recommendations for slide review: potential impact on review rate and turnaround time.
Annales de biologie Clinique. 75(3): 285
http://www.jle.com/fr/revues/abc/e-docs/from_xe_2100_to_xn_9000_from_sis_standard_to_gfhc_recommendations_for_slide_review_potential_impact_on_review_rate_and_turnaround_time_309721/article.phtml

**What we see as the essence:** Changing from the XE-2100 to XN-9000 and implementing the Biomedical Validation ruleset led to a significant reduction in review rate (from 35.8% to 25.9%) and TAT. In this hospital this resulted in a cost reduction of 7000 Euros over 6 months.

Van Dievoet MA et al. (2016)
Performance evaluation of the Sysmex® XP-300 in an oncology setting: evaluation and comparison of hematological parameters with the Sysmex® XN-3000
Int J Lab Hematol; 38(5): 490

**What we see as the essence:** The XP-300 showed very good precision and linearity results, comparable with the XN-3000 analyser.

Cornet E et al. (2016)
Evaluation and optimization of the extended information process unit (E-IPU) validation module integrating the sysmex flag systems and the recommendations of the French-speaking cellular hematology group (GFHC).

**What we see as the essence:** Using the biomedical validation criteria, 21.3% of samples triggered a smear review. Modification of four criteria reduced the number of smears from 21.3% to 15.0% without loss of clinical value.

Berda-Haddad Y et al. (2016)
Increased mean corpuscular haemoglobin concentration: artefact or pathological condition?
Int J Lab Hematol 39(1):32

**What we see as the essence:** The use of the optical RBC parameters from the XN-Series can save time and help in the determination of the cause of increased MCHC.
**Geara C et al. (2016)**
Comparative study of quantitative performances between the new Sysmex XN-L (XN-550) haematology analyser and the XN-9000 in a routine laboratory.
Int J Lab Hematol 38(1):e10


**What we see as the essence:** The XN-Series and XN-L Series were compared; correlations were good and the study showed that the XN-L Series provided the same high quality as the XN-Series.

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**Egele A et al. (2015)**
Automated detection and classification of teardrop cells by a novel RBC module using digital imaging/microscopy.
Int J Lab Hematol 37(6):e153


**What we see as the essence:** The authors report excellent detection of teardrop cells in samples from patients with myelofibrosis and MDS, using the Advanced RBC Morphology software.

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**Ferrero-Vacher C et al. (2015)**
Utilisation des paramètres érythrocytaires Sysmex dans un cas d’hémolyse sèvère (Erythrocytic parameters Sysmex in a case of severe haemolysis.
Annales de Biologie Clinique 73(6):729


**What we see as the essence:** Case report of severe haemolytic anaemia with cold agglutinins, identified by increased MCHC and qualitative alarms. The RBC-O and HGB-O parameters from the RET channel, and the RBC most frequent volume (R-MFV) allowed to report the correct results.

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**Arneth B et al. (2015)**
Technology and New Fluorescence Flow Cytometry Parameters in Hematological Analyzers.
J Clin Lab Anal 29(3):175


**What we see as the essence:** This paper gives a good overview of the technology behind the XE-Series and the benefits of flow cytometry and automatic cell counting. It shows that the XE-5000 delivers faster accurate results than older analysers.
Genevieve F et al. (2014)
Smear microscopy revision: propositions by the GFHC.
feuilles de Biologie VOL LVI N° 317
Free online: http://www.gfhc.fr/upload/smear-microscopic-revision.pdf

What we see as the essence: The GFHC reviewed in detail the criteria used within the CBC to generate blood smears and has decided on a number of minimum recommendations, defining threshold values and various situations in which the blood smear review is desirable.

Wang H et al. (2013)
Use of RBC-O and S-MCV Parameters of Sysmex XE-2100 in a Patient with RBC Cold Agglutination
Clin Lab.;59: 217
http://clinical-laboratory.de/article/1068

What we see as the essence: A combination of sample dilution and the use of RBC parameters from the RET channel on the XE-2100 is described to obtain accurate RBC parameters from samples with RBC cold agglutination without heating of the sample.

Godon A et al. (2012)
Anomalies et erreurs de détermination de l’hémogramme avec les automates d’hématologie cellulaire Partie 3. Hémoglobine, hématies, indices érythrocytaires, réticulocytes*.
Ann Biol Clin 2012; 70(2): 155
Article in French - Free online: http://www.jle.com/fr/revues/abc/e-docs/anomalies_et_erreurs_de_determination_de_lhemogramme_avec_les_automates_dhematologie_cellulaire_partie_3_hemoglobine_hematies_indices_erythrocytaires_reticulocytes_292317/article.phtml

What we see as the essence: A summary report about potential interferences of CBC parameters with focus on situations leading to abnormal HGB, RBC and MCV, resulting in abnormal calculated RBC indices, e.g. MCHC. Alternative strategies may support management of interferences.

RET/ IRF

Twari A et al. (2018)
Applying newer parameter Ret-He (reticulocyte haemoglobin equivalent) to assess latent iron deficiency (LID) in blood donors-study at a tertiary care hospital in India.
Vox Sang. 2018; 113(7): 639

What we see as the essence: RET-He can be used as a routine screening test to detect latent iron deficiency in blood donors. This could provide an opportunity to make appropriate and timely interventions like dietary changes or drug supplementation.
Buoro S et al. (2017)
Innovative haematological parameters for early diagnosis of sepsis in adult patients admitted in intensive care unit.
J Clin Pathol.; 71(4): 330
http://jcp.bmj.com/content/71/4/330.long

What we see as the essence: The combination of an increased value of IPF# and a decreased value of RET% 24 hours before the onset of sepsis in ICU patients may be considered an early, rapid, inexpensive and widely available measure of sepsis prediction.

Morkis IVC et al. (2015)
Assessment of immature platelet fraction and immature reticulocyte fraction as predictors of engraftment after hematopoietic stem cell transplantation.

What we see as the essence: Both IRF% and IPF% can be used to predict neutrophil and platelet recovery, respectively. Work was done on XE-5000.

Yesmin S et al. (2011)
Immature reticulocyte fraction as a predictor of bone marrow recovery in children with acute lymphoblastic leukaemia on remission induction phase.
Bangladesh Med Res Council Bull 37(2): 57-60

What we see as the essence: In 52% of paediatric ALL patients, IRF% values rose before NEUT# values during recovery after chemotherapy. Therefore, monitoring of both parameters may be beneficial.

Gonçalo AP et al. (2011)
Predictive value of immature reticulocyte and platelet fractions in hematopoietic recovery of allograft patients.
Transplant Proc 43: 24
http://www.transplantation-proceedings.org/article/S0041-1345(10)01945-7/abstract

What we see as the essence: The immaturity fractions IRF and IPF offer an easy and early evaluation method of posttransplantational recovery of the bone marrow.
Torres Gomez A et al. (2003)
Utility of reticulocyte maturation parameters in the differential diagnosis of macrocytic anemias.

What we see as the essence: Reticulocyte maturation parameters (measured here on ABX Pentra) can support differential diagnosis of macrocytic anaemias.

RET-He / RBC-He

Wirawan R et al. (2017)
Concordance between Reticulocyte Hemoglobin Equivalent and Reticulocyte Hemoglobin Content in CKD Patients Undergoing Hemodialysis.
Acta Med Indones; 49(1):34
Free online: http://www.actamedindones.org/index.php/ijim/article/view/316/pdf

What we see as the essence: A very strong correlation (r=0.91) and a good concordance was found between RET-He and CHr with a mean bias of 0.5 pg in chronic kidney disease patients undergoing haemodialysis. It indicates that RET-He and CHr can both be used for assessing iron status.

Toki Y et al. (2017)
Evaluation of the hypochromic erythrocyte and reticulocyte hemoglobin content provided by the Sysmex XE-5000 analyzer in diagnosis of iron deficiency erythropoiesis.
Int J Hematol; 106(1): 116
http://rd.springer.com/article/10.1007/s12185-017-2212-6

What we see as the essence: RET-He was shown to be a clinically useful marker for determining iron deficiency in the general population and can also be used for the evaluation of the efficacy of iron administration.

Buttarello M et al. (2016)
Evaluation of the hypochromic erythrocyte and reticulocyte hemoglobin content provided by the Sysmex XE-5000 analyzer in diagnosis of iron deficiency erythropoiesis.
Clin Chem Lab Med; 54(12):1939

What we see as the essence: RET-He and %Hypo-He, measured on the XE-5000, allowed identification of patients with iron deficiency, especially those who had already developed anaemia. RET-He had a better sensitivity, presumably because it is more responsive to iron deficiency.
Weimann A et al. (2016)
Delta-He, Ret-He and a New Diagnostic Plot for Differential Diagnosis and Therapy Monitoring of Patients Suffering from Various Disease-Specific Types of Anemia. Clin Lab;62(4):667
http://clinical-laboratory.de/article/1982

What we see as the essence: A diagnostic plot using RET-He and Delta-He was developed based on differences between different patient groups suffering from anaemia. Several case examples show the clinical utility of this plot for therapy monitoring.

Mehta S et al. (2016)
Reticulocyte Hemoglobin vis-a-vis Serum Ferritin as a Marker of Bone Marrow Iron Store in Iron Deficiency Anemia. J Assoc Physicians India 64(11):38.
Free online: http://japi.org/november_2016/pdf/06_oa_Reticulocyte.pdf

What we see as the essence: This study showed that RET-He is a better predictor of bone marrow iron stores in patients with severe anaemia than serum ferritin.

Urrechaga E et al. (2016)

What we see as the essence: HYPO-He and RET-He are reliable parameters for the study of erythropoiesis status in hemodyalisis patients.

Al-Ghananim RT et al. (2016)
Reticulocyte Hemoglobin Content During the First Month of Life in Critically Ill Very Low Birth Weight Neonates Differs From Term Infants, Children, and Adults. J Clin Lab Anal. 30(4):326

What we see as the essence: "RET-He values from the XE-2100 were lower in very low birth weight infants than in term infants, children and adults. RET-He was 31.8 pg within 24 hr after birth and subsequently declined to a steady-state level of 28.4 pg."
Archer NM et al. (2015)
Diagnosis of iron-deficient states.  
http://www.tandfonline.com/doi/pdf/10.3109/10408363.2015.1038744

**What we see as the essence:** This review gives an overview of the haematological, biochemical and genetic markers for identifying iron deficiency. RBC-He, RET-He, Delta-He, HYPO-He and MicroR are mentioned besides the standard RBC indices.

Peerschke E et al. (2014)
Using the Hemoglobin Content of Reticulocytes (RET-He) to Evaluate Anemia in Patients With Cancer.  
Am J Clin Pathol 142:506-512  
http://ajcp.ascpjournals.org/content/142/4/506.long

**What we see as the essence:** RET-He values above 31 or 32 pg could be used to rule out iron deficiency in cancer patients. In the present study the use of RET-He would have reduced the number of biochemical iron studies by 66% (from 209 to 70).

Schoorl M et al. (2012)
Effects of iron supplementation on red blood cell hemoglobin content in pregnancy.  
Hematology Reports 4: e24.  
Free online: http://www.pagepress.org/journals/index.php/hr/article/view/hr.2012.e24

**What we see as the essence:** Ret-He and Ret-He/RBC-He ratio are sensitive markers for screening when a decrease in red blood cell haemoglobin content is observed and for monitoring short-term effects of iron supplementation. The authors recommend integrating these parameters into the protocol for anaemia screening and monitoring during pregnancy.

Schoorl M et al. (2012)
Temporary impairment of reticulocyte haemoglobin content in subjects with community-acquired pneumonia.  

**What we see as the essence:** In patients with community-acquired pneumonia, acute inflammation results in decreased RET-He values at an early stage, reflecting acute erythropoietic dysfunction.
Urrechaga E et al. (2013)
Erythrocyte and reticulocyte indices in the assessment of erythropoiesis activity and iron availability. Int J Lab Hematol 35: 144–149.

What we see as the essence: RET-He and %HYPO-He are helpful in assessing erythropoiesis and iron status.

Maier-Redelsperger M et al. (2010)

What we see as the essence: A special algorithm combining RBC-He, RET-He and lactate dehydrogenase bears the potential as a marker of haemolysis strongly correlated with albuminuria in sickle cell anemia patients.

Jonckheere S et al. (2010)
http://www.karger.com/Article/FullText/313785

What we see as the essence: Due to fluctuations of iron status parameters, a fixed time point should be used for iron status monitoring during erythropoietin therapy.

Leers MP et al. (2010)
The value of the Thomas-plot in the diagnostic work up of anemic patients referred by general practitioners. Int J Lab Hematol 32: 572–81.

What we see as the essence: The Thomas-plot is helpful in diagnosing patients referred from general practitioners and differentiating functional iron deficiency from classical iron deficiency.
**Schoorl M et al. (2010)**
Changes in red blood cell hemoglobinization during pregnancy.
What we see as the essence: RET-He is a useful sensitive and early indicator of iron status in the second half of pregnancy and should ideally be measured in combination with zinc protoporphyrin (ZPP) and IRF.

**Van Wyck DB et al. (2010)**
Analytical and biological variation in measures of anemia and iron status in patients treated with maintenance hemodialysis.
What we see as the essence: RET-He could prove superior to transferrin saturation (TSAT) and ferritin in monitoring iron status of haemodialysis patients due to a lower biological variation.

**Maconi M et al. (2009)**
Erythrocyte and reticulocyte indices in iron deficiency in chronic kidney disease: comparison of two methods.
What we see as the essence: RET-He and CHr correlate and agree well in evaluating CKD patients needing iron support.

**Miwa N et al. (2009)**
Usefulness of measuring reticulocyte hemoglobin equivalent in the management of haemodialysis patients with iron deficiency.
Int J Lab Hematol 32: 248
What we see as the essence: RET-He is equivalent to CHr and useful in managing haemodialysis patients with iron deficiency as it responds more rapidly than HGB.
Mast A et al. (2008)
Reticulocyte hemoglobin content.
Am J Hematol 83: 307
What we see as the essence: Reticulocyte haemoglobin can be used to differentiate iron deficiency from other causes of anaemia and as an early marker to monitor the therapy.

Thomas C et al. (2006)
The diagnostic plot: a concept for identifying different states of iron deficiency and monitoring the response to epoetin therapy.
What we see as the essence: The Thomas-plot incl. RET-He can be used for the differential diagnosis of anaemia and also gives therapy options.

Brugnara C et al. (2006)
Reticulocyte hemoglobin equivalent (Ret He) and assessment of iron-deficient states.
Free online: http://www.ncbi.nlm.nih.gov/pmc/articles/PMC1618805/pdf/clh0028-0303.pdf
What we see as the essence: RET-He is a reliable marker of cellular haemoglobin content and can be used to identify iron-deficient states, particularly in dialysis patients. RET-He and CHr are in good agreement.

Schoorl M et al. (2006)
Erythropoiesis activity, iron availability and reticulocyte hemoglobinization during treatment with hemodialysis and in subjects with uremia.
http://www.clin-lab-publications.com/abstracts/160.html
What we see as the essence: Biochemical parameters reflecting functional iron availability and haematological parameters reflecting haemoglobinisation are interdependent.
Thomas L et al. (2005)
Reticulocyte hemoglobin measurement -- comparison of two methods in the diagnosis of iron-restricted erythropoiesis.

**What we see as the essence:** RET-He can replace CHr in the diagnostic Thomas-plot without loss of sensitivity or specificity.

Canals C et al. (2005)
Clinical utility of the new Sysmex XE 2100 parameter - reticulocyte hemoglobin equivalent – in the diagnosis of anemia.
Haematologica 90: 1133–1134.
Free online: http://www.haematologica.org/content/90/8/1133.long

**What we see as the essence:** RET-He is useful for the differential diagnosis of iron deficiency anaemia vs anaemia of chronic disease and could also be helpful in the identification of thalassaemia patients.

Buttarello M et al. (2004)
The new reticulocyte parameter (RET-Y) of the Sysmex XE 2100: its use in the diagnosis and monitoring of posttreatment sideropenic anemia.
Free online: http://ajcp.ascpjournals.org/content/121/4/489.long

**What we see as the essence:** RET-Y closely correlates with CHr and can be used for diagnosis and early monitoring after the administration of intravenous iron.

HYPO-He / HYPER-He / MicroR / MacroR

Schoorl M et al. (2012)
Am J Clin Pathol 138: 300
http://ajcp.ascpjournals.org/content/138/2/300.abstract

**What we see as the essence:** The authors conclude that the advanced algorithms, derived from extended RBC parameters provided by the Sysmex XE-5000 analyzer, are useful as laboratory devices for anaemia screening.
Persijn L et al. (2012)
Screening for hereditary spherocytosis in routine practice: evaluation of a diagnostic algorithm with focus on non-splenectomised patients.

What we see as the essence: The hereditary spherocytosis diagnostic tool by Mullier et al. is useful and works, but needs fine-tuning to the local patient population.

Mullier F et al. (2011)
Additional erythrocytic and reticulocytic parameters helpful for diagnosis of hereditary spherocytosis: results of a multicentre study.
Ann Hematol 90: 759–768.

What we see as the essence: Combining several RBC parameters allows to efficiently screen for hereditary spherocytosis even in mild cases.

Urrechaga E et al. (2011)
The role of automated measurement of RBC subpopulations in differential diagnosis of microcytic anemia and β-thalassemia screening.

What we see as the essence: Because of high sensitivity and specificity, the new index %MicroR-%HYPO-He was the most reliable index in the differential diagnosis of microcytic anaemias.

Urrechaga E. et al. (2011)
Erythrocyte and reticulocyte parameters in iron deficiency and thalassemia.

What we see as the essence: Beta-thalassaemia can be recognised through high RBC, small MCV, high %MicroR and moderately increased IRF, whereas iron deficiency shows high RDW and %HYPO-He.
Urrechaga E et al. (2011)
The role of automated measurement of red cell subpopulations on the Sysmex XE-5000 analyzer in the differential diagnosis of microcytic anemia.
Int J Lab Hematol 33: 30–36.

What we see as the essence: Because of high sensitivity and specificity, the new index %MicroR-%HYPO-He was the most reliable index in the differential diagnosis of microcytic anaemias.

Urrechaga E et al. (2009)

What we see as the essence: The new parameters %HYPO-He/%HYPER-He and %MicroR/%MacroR appear to be sensitive for detecting small changes in the number of red cells with inadequate haemoglobinisation and volume in order to distinguish beta-thalassaemia from iron deficiency anaemia.

NRBC

Menk M et al. (2018)
Nucleated red blood cells as predictors of mortality in patients with acute respiratory distress syndrome (ARDS): an observational study.
Ann Intensive Care. 8(1): 42
Free online: https://link.springer.com/content/pdf/10.1186%2Fs13613-018-0387-5.pdf

What we see as the essence: The study results confirmed previous findings in critically ill patients suggesting that NRBC are equally predictive of mortality in acute respiratory distress syndrome (ARDS). NRBC-positive patients were found to require longer treatment with mechanical ventilation, extra-corporal gas exchange and had prolonged ICU stay when compared with NRBC-negative patients.

Monteiro Junior JG et al. (2015)
Nucleated Red Blood Cells as Predictors of All-Cause Mortality in Cardiac Intensive Care Unit Patients: A Prospective Cohort Study.
PLoS One. 29;10(12):e0144259
Free online: http://www.ncbi.nlm.nih.gov/pmc/articles/PMC4695082/

What we see as the essence: "The presence of NRBC (XE-2100) was associated with a higher ICU mortality (49.4% vs 21.7%, P<0.001) as well as in-hospital mortality (61.4% vs 33.3%, p = 0.001)."
Cremer M et al. (2015)
Nucleated red blood cells as marker for an increased risk of unfavorable outcome and mortality in very low birth weight infants.
Early Hum Dev.; 91(10):559-563
http://www.earlyhumandevelopment.com/article/S0378-3782(15)00123-1/abstract

What we see as the essence: This study of 438 low birth weight infants indicates that an NRBC count obtained 24-120 h after birth can serve as a surrogate marker for later severe morbidity and mortality. The optimal cut-off value was 2x10^9/L with 83% sensitivity and 75% specificity.

Tantanate C et al. (2014)

What we see as the essence: NRBC counts from the XN-Series could replace manual counts: the precision of the XN-Series was superior and a small bias (manual counts slightly higher than NRBC counts from the XN-Series) was only observed for NRBC counts above 200/100 WBC.

Hotton J et al. (2013)
http://ajcp.ascpjournals.org/content/140/6/845.abstract

What we see as the essence: Repeatability, linearity and carryover was good for all tested analysers, and correlation between the analysers was good for HGB, MCV, PLT and WBC.
Quotes: "The XN showed a higher sensitivity than the SAPH and DxH for all flags of interest." "For the first time, we have decreased the slide review for our laboratory from 20% with the SAPH to 9.3% with the XN."

Parco S et al. (2013)
Free online: http://www.dovepress.com/getfile.php?fileID=15732

What we see as the essence: An excellent correlation was found between manual NRBC counts and NRBC counts from the XE-2100 (r2 = 0.94) in umbilical cord blood. This number may be used to correct the WBC count and thereby guarantee an adequate WBC concentration for blood banking of umbilical cord blood.
Gasparović V et al. (2012)
Nucleated red blood cells count as first prognostic marker for adverse neonatal outcome in severe preeclamptic pregnancies.
Free online: http://hrcak.srce.hr/index.php?show=clanak&id_clanak_jezik=133776

What we see as the essence: An increased count of nucleated red blood cells in preterm newborns born from pregnancies with severe preeclampsia seems to be the first significant marker for detecting adverse neonatal outcome.

Pipitone S et al. (2012)
Evaluation of automated nucleated red blood cells counting on Sysmex XE-5000 and Siemens ADVIA 2120.

What we see as the essence: The results show excellent analytical performances for the XE-5000, with high accuracy and precision. In agreement with previous studies, the authors also showed that despite similar performance in terms of analytical imprecision, the overall correlation with microscopy is higher for XE-5000 than for ADVIA 2120, i.e., correlation coefficient 0.97 vs. 0.67 and AUC 0.97 vs. 0.73, respectively.

Kuert S et al. (2011)
Association of nucleated red blood cells in blood and arterial oxygen partial tension.

What we see as the essence: The NRBC count is an independent risk indicator of poor prognosis and mortality, NRBC-positive patients required a longer stay in the intensive care unit.

Danise P et al. (2011)
Evaluation of nucleated red blood cells in the peripheral blood of hematological diseases.

What we see as the essence: NRBC are found in nearly all onco-haematological diseases at diagnosis and frequently during therapy. They are absent at remission.
**Danise P et al. (2009)**
Nucleated red blood cells and soluble transferrin receptor in thalassemia syndromes: relationship with global and ineffective erythropoiesis.


What we see as the essence: The NRBC count helps defining ineffective erythropoiesis in thalassaemia patients and supporting transfusion management.

**Stachon A et al. (2007)**
Nucleated red blood cells in the blood of medical intensive care patients indicate increased mortality risk: a prospective cohort study.
Crit Care 11: R62.
Free online: http://ccforum.com/content/pdf/cc5932.pdf

What we see as the essence: The NRBC count is one indicator of mortality — persistence (observed in daily screenings) and high concentration are both indicators for poor prognosis.

**Stachon A et al. (2006)**
Poor prognosis indicated by nucleated red blood cells in peripheral blood is not associated with organ failure of the liver or kidney.


What we see as the essence: The NRBC count is one indicator of mortality independent of other factors such as kidney or liver failure.

**Stachon A et al. (2006)**
Daily monitoring of nucleated red blood cells in the blood of surgical intensive care patients.

What we see as the essence: NRBC count is an early indicator of mortality – daily screening is recommended.
Otsubo H et al. (2005)
Persistent nucleated red blood cells in peripheral blood is a poor prognostic factor in patients undergoing stem cell transplantation.

What we see as the essence: Monitoring NRBC in stem cell transplantation patients provides useful clinical information - NRBC can be observed persistently in non-survivors.

Stachon A et al. (2004)
High in-hospital mortality of intensive care patients with nucleated red blood cells in blood.

What we see as the essence: The NRBC count is of high prognostic power regarding in-hospital mortality.

Wang FS et al. (2003)
Development and clinical application of nucleated red blood cell counting and staging on the automated haematology analyser XE-2100.

What we see as the essence: The NRBC count correlates well with flow cytometry.

Stachon A et al. (2002)
http://www.journals.elsevierhealth.com/periodicals/ymlc/article/S0022-2143(02)00104-X/abstract

What we see as the essence: NRBC are often an only transient observation, but they indicate a poor prognosis, whether transient or persistent.

Briggs C et al. (2000)
New quantitative parameters on a recently introduced automated blood cell counter - the XE 2100.

What we see as the essence: The automated NRBC count was highly correlated with the manual reference count (r2=0.97) and thus eliminates the need for manual NRBC counts.
FRC

Hervent AS et al. (2015)
Evaluation of schistocyte analysis by a novel automated digital cell morphology application.
Int J Lab Hematol 37(5):588

What we see as the essence: This performance evaluation showed that the CELLAVISION Advanced RBC Software Application is easy to use and provides a sensitive and reproducible measurement of schistocytes in peripheral blood.

Lesesve JF et al. (2015)
Fragmented red cells reference range for the Sysmex XN®-series of automated blood cell counters.
Int J Lab Hematol 37(5):583.

What we see as the essence: Normal values were determined on the XN-Series for the percentage fragmented red blood cells, FRC%: 0.14 +/- 0.35% (mean 0%). It was also found that HYPO-He correlates with FRC%; so samples with both a high HYPO-He and FRC% should be interpreted with care.

Lesesve JF et al. (2012)
Fragmented red blood cells automated measurement is a useful parameter to exclude schistocytes on the blood film.
Int J Lab Hematol 34(6):566.

What we see as the essence: The automated FRC count offers a better degree of certainty than microscopy to exclude the presence of fragmented RBC.

Abe Y et al. (2009)
The effectiveness of measuring for fragmented red cells using an automated hematology analyser in patients with thrombotic microangiopathy.
http://cat.sagepub.com/content/15/3/257.full.pdf

What we see as the essence: In conclusion, the FRC level is a simple and useful marker for thrombotic microangiopathy (TMA), and an FRC level of 1.2% is recommended as the cutoff value for the diagnosis of TMA.
Imoto S et al. (2005)
Usefulness of sequential automated analysis of fragmented red blood cells for the differential diagnosis of TTP-hemolytic uremic syndrome following allogeneic hematopoietic cell transplantation. Lab Hematol 11: 131–136.
http://europepmc.org/abstract/med/16024337

What we see as the essence: Sequential monitoring of FRC% may be a reliable marker for a specific type of complication (TTP-HUS; thrombotic thrombocytopenic purpura haemolytic uraemic syndrome) after allogeneic haematopoietic precursor cell transplantation.

Banno S et al. (2005)

What we see as the essence: The determination of FRC% by the XE-2100 enables early diagnoses and monitoring of TTP (thrombotic thrombocytopenic purpura) or TMA (thrombotic microangiopathy) and will be useful in the clinical field.

Jiang M et al. (2001)

Initial study showing that the automated FRC% measurement is promising.

Malaria-infected RBC (MI-RBC)

Pillay E et al. (2019)
Evaluation of automated malaria diagnosis using the Sysmex XN-30 analyser in a clinical setting Malaria J 18(1): 15
Free online: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC6341646/

What we see as the essence: The novel technology of the Sysmex XN-30 provides a robust, rapid, automated and accurate platform for diagnosing malaria. It allows precise recognition and demonstrated a sensitivity and specificity of 100% for malaria parasitaemia detection.
**Post A et al. (2019)**

The XN-30 hematology analyzer for rapid sensitive detection of malaria: a diagnostic accuracy study

*BMC Medicine; 17(1):103*

*Free online: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC6543632/*

**What we see as the essence:** The novel technology of the Sysmex XN-30 provides a robust, rapid, automated and objective platform for diagnosing and quantifying malaria. The XN-30 ensures the prompt initiation of the malaria treatment and the malaria anaemia together with the reliable treatment monitoring.

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**ESR**

**Kratz A et al. (2017)**

ICSH recommendations for modified and alternate methods measuring the erythrocyte sedimentation rate.

*Int J Lab Hematol; 39(5):448.*

*Free online: https://onlinelibrary.wiley.com/doi/full/10.1111/ijlh.12693*

**What we see as the essence:** The gold standard for the determination of the erythrocyte sedimentation rate is still the Westergren method. Alternative methods should be accepted when they have been appropriately validated and their results are expressed by comparison with the gold standard.

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**Schapkaitz E et al. (2017)**

Evaluation of the InteRRliner automated erythrocyte sedimentation rate analyzer for a large academic laboratory.


**What we see as the essence:** Indicated by the high correlation coefficient of 0.96 the InteRRliner showed an excellent comparability to the HumaSed ESR method.