

sebia

MINICAP HEMOGLOBIN(E)

Ref. 2207

Ref. 2227*

MINICAP FLEX-PIERCING

PHORESIS VS ≥ 9.15

IVD

CE

2020/07

MINICAP HEMOGLOBIN(E) USING THE MINICAP FLEX-PIERCING INSTRUMENT

INTENDED USE

The MINICAP HEMOGLOBIN(E) kit is designed for the separation of the normal hemoglobins (A, A2 and F) in human blood samples, and for the detection of the major hemoglobin variants (S, C, E and D), by capillary electrophoresis in alkaline buffer (pH 9.4) with the SEBIA MINICAP FLEX-PIERCING instrument. The MINICAP HEMOGLOBIN(E) kit is designed for laboratory use.

The MINICAP FLEX-PIERCING instrument is an automated analyzer which performs a complete hemoglobin profile for the quantitative analysis of the normal hemoglobin fractions A, A2 and F and for the detection of major hemoglobin variants S, C, E and D. The assay is performed on the hemolysate of whole blood samples collected in tubes containing K₂EDTA or K₃EDTA as anticoagulant.

For *In Vitro* Diagnostic Use.

PRINCIPLE OF THE TEST 1-20

Hemoglobin is a complex molecule composed of two pairs of polypeptide chains. Each chain is linked to the heme, a tetrapyrrolic nucleus (porphyrin) which chelates an iron atom. The heme part is common to all hemoglobins and their variants. The type of hemoglobin is determined by the protein part called globin. Polypeptide chains α , β , δ and γ constitute the normal human hemoglobins:

- hemoglobin A= $\alpha 2 \beta 2$
- hemoglobin A2= $\alpha 2 \delta 2$
- fetal hemoglobin F= $\alpha 2 \gamma 2$

The α -chain is common to these three hemoglobins.

The hemoglobin spatial structure and other molecular properties (like that of all proteins) depend on the nature and the sequence of the amino acids constituting the chains. Substitution of amino acids by mutation is responsible for formation of hemoglobin variants which have different surface charge and consequently different electrophoretic mobilities, which also depend on the pH and ionic strength of the buffer.

The resulting qualitative (or structural) abnormalities are called hemoglobinopathies^(9, 10, 13). Decreased synthesis of one of the hemoglobin chains leads to quantitative (or regulation) abnormalities, called thalassemias.

Hemoglobin electrophoresis is a well established technique routinely used in clinical laboratories for screening samples for hemoglobin abnormalities^(1, 2, 3, 4, 12). The MINICAP FLEX-PIERCING instrument has been developed to provide complete automation of this testing with fast separation and good resolution. In many aspects, the methodology can be considered as an intermediary type of technique between classical zone electrophoresis and liquid chromatography^(8, 11).

The MINICAP FLEX-PIERCING instrument uses the principle of capillary electrophoresis in free solution. With this technique, charged molecules are separated by their electrophoretic mobility in an alkaline buffer with a specific pH. Separation also occurs according to the electrolyte pH and electroosmotic flow⁽⁵⁾.

The MINICAP FLEX-PIERCING instrument has capillaries functioning in parallel allowing 2 simultaneous analyses for hemoglobin quantification. A sample dilution with hemolysing solution is prepared and injected by aspiration at the anodic end of the capillary. A high voltage protein separation is then performed and direct detection of the hemoglobins is made at 415 nm at the cathodic end of the capillary. Before each run, the capillaries are washed with a Wash Solution and prepared for the next analysis with buffer.

The hemoglobins, separated in silica capillaries, are directly and specifically detected at an absorbance wave length of 415 nm which is specific to hemoglobins. The resulting electrophoregrams are evaluated visually for pattern abnormalities.

Direct detection provides accurate relative quantification of individual hemoglobin fraction, with particular interest, such as A2 hemoglobin for thalassemia diagnostic. In addition, the high resolution of this procedure allows the identification of hemoglobin variants, in particular, differentiation of hemoglobins S from D, and E from C.

The hemoglobin A2 quantification can also be performed when hemoglobin E is present.

By using alkaline pH buffer, normal and abnormal (or variant) hemoglobins are detected in the following order, from cathode to anode: $\delta A'2$ (A2 variant), C, A2/O-Arab, E, S, D, G-Philadelphia, F, A, Hope, Bart's, J, N-Baltimore and H.

The carbonic anhydrase is not visualized on the hemoglobin electrophoretic patterns, this permits to identify hemoglobin A2 variants in this migration zone.

REAGENTS AND MATERIALS SUPPLIED IN THE MINICAP HEMOGLOBIN(E) KITS

WARNING: See the safety data sheets.

ITEMS	PN 2207	PN 2227*
Buffer (ready to use)	2 vials, 250 mL each	6 vials, 250 mL each
Hemolysing solution (ready to use)	1 vial, 225 mL	3 vials, 225 mL each
Wash solution (stock solution)	1 vial, 25 mL	3 vials, 25 mL each
Reagent cups	1 pack of 125	3 packs of 125 each
Filters	3 filters	3 filters
Bins for used cups	4 bins	12 bins
Hemolysing solution bar code labels	5 sheets of 4 labels	15 sheets of 4 labels

* MINICAP HEMOGLOBIN(E) MAXI-KIT

During transportation, the kit can be kept without refrigeration (15 to 30 °C) for 15 days without any adverse effects on performance.

FOR OPTIMAL MANAGEMENT OF TRACEABILITY : All reagents from the same kit must be used together.

TO OBTAIN THE EXPECTED PERFORMANCES : The package insert instructions must be observed.

WARNING : Do not use marketed deionized water, such as water for ironing for example (risk of important capillaries damage). Use only water with ultrapure quality, such as injection grade water.

1. BUFFER

Preparation

The buffer is ready to use. It contains : buffer solution pH 9.4 ± 0.5 ; additives, nonhazardous at concentrations used, necessary for optimum performance.

Use

Buffer for analysis of hemoglobins in MINICAP FLEX-PIERCING instrument.

Storage, stability and signs of deterioration

Store the buffer refrigerated (2 to 8 °C). It is stable until the expiration date indicated on the kit package or buffer vial labels. Avoid storage at room temperature for a long time or close to a window or to a heat source.

DO NOT FREEZE.

IMPORTANT : When stored at 2 – 8 °C and prior to use, it is necessary for the buffer to reach room temperature ; when it is full, let the buffer vial at room temperature for at least 3 hours prior to use. If this precaution is not respected, the performances of the procedure may be affected.

WARNING : Do not pre-heat the buffer in hot water.

Once the buffer vial has been opened and positioned on the MINICAP FLEX-PIERCING instrument, it is stable for a maximum of 1 month (accumulated) at room temperature (15 to 30 °C). After each use, the buffer must imperatively be stored refrigerated (between 2 and 8 °C) without any delay, it is then stable until the expiration date indicated on the buffer vial label.

IMPORTANT : The accumulated time of the buffer stored at room temperature must not exceed 1 month. This time of 1 month storage takes account of the time for the buffer to come to room temperature.

Discard buffer if it changes its appearance, e.g., becomes cloudy due to microbial contamination.

2. HEMOLYSING SOLUTION

Preparation

Hemolyzing solution is ready to use. It contains buffer solution pH 8.7 ± 0.5 ; additives, nonhazardous at concentrations used, necessary for optimum performance.

Use

To dilute and hemolyze red blood cells.

Storage, stability and signs of deterioration

Store Hemolysing Solution refrigerated (2 to 8 °C). It is stable until the expiration date indicated on the kit package or Hemolysing Solution vial label. Avoid storage at room temperature or close to a window or to a heat source. DO NOT FREEZE.

Discard Hemolysing Solution if it changes its appearance, e.g., becomes cloudy due to microbial contamination.

IMPORTANT : After each use, close immediately and tightly the hemolysing solution vial and store it refrigerated.

NOTE : Hemolysing solution may show a more or less marked yellow-orange colour without any adverse effects on its performance.

3. WASH SOLUTION

Preparation

The vial of the stock wash solution should be diluted up to 250 mL with distilled or deionized water.

After dilution, the wash solution contains an alkaline solution pH ≈ 12.

Use

For washing the capillaries before hemoglobin electrophoresis.

IMPORTANT : Before filling the wash solution container, it is recommended to wash the opening of the container, the connector and the tube with plenty of distilled or deionized water to avoid salts deposit.

Storage, stability and signs of deterioration

Store the stock and working wash solutions in closed containers at room temperature or refrigerated. The stock wash solution is stable until the expiration date indicated on the kit or wash solution vial label.

Working wash solution is stable for 3 months.

Discard working wash solution if it changes its appearance, e.g., becomes cloudy due to microbial contamination.

4. REAGENT CUPS

Use

Single use cups for the preparation of biological samples to analyze with the automated instrument. To be placed on the automated loading system for cups of MINICAP FLEX-PIERCING. One reagent cup is intended for the analysis of 2 samples.

WARNING : After use, reagent cups with biological samples have to be handled with care. When the analysis is completed, reagent cups must be discarded with biological waste products and they must NEVER be reused.

Storage

Before use, store the reagent cups in their sealed package in a clean and dry place and at a temperature comprised between 2 and 30 °C.

5. FILTERS

Use

Disposable filters for filtration of analysis buffer, working wash solution and distilled water (used for capillaries rinsing).

IMPORTANT : When kit replacement, change systematically all the filters. Wear clean gloves for handling and installation of filters.

Screw one filter at the connector situated at the extremity of each tube that plunges in the vials of buffer, wash solution and distilled or deionized water. When setting filters on MINICAP FLEX-PIERCING instrument, rinse the connectors and the tubes with distilled or deionized water.

Storage

Before use, store the filters in their sealed package in a dry place at room temperature or refrigerated.

6. BINS FOR USED CUPS

Use

Bins intended for automated collection of used reagent cups in MINICAP FLEX-PIERCING. To place in MINICAP FLEX-PIERCING at the location intended for this purpose.

WARNING : *Bins containing used reagent cups with biological samples have to be handled with care.*

7. HEMOLYSING SOLUTION BAR CODE LABELS

Use

Labels to identify the tube containing the hemolysing solution (HEMOGLOBIN(E) HEMOLYSING SOLUTION).

REAGENTS REQUIRED BUT NOT SUPPLIED

WARNING : *See the safety data sheets.*

1. NORMAL Hb A2 CONTROL

Composition

The Normal Hb A2 Control (SEBIA, PN 4778) is obtained from a pool of normal human blood samples. The Normal Hb A2 Control is in a stabilized lyophilised form.

Intended use

The Normal Hb A2 Control is designed for the migration control **before starting a new analysis sequence and after the analysis of a complete rotating sampler**, and for the quality control of human hemoglobin A2 quantification with the MINICAP HEMOGLOBIN(E) electrophoresis procedure performed with the MINICAP FLEX-PIERCING instrument.

Reconstitute each Normal Hb A2 Control vial with the volume of distilled or deionized water, as indicated in the package insert of the Normal Hb A2 Control. Allow to stand for 30 minutes and mix gently (avoid formation of foam).

NOTE : *The precision of the reconstitution volume to be maintained is ± 1.0 %.*

Migration control :

IMPORTANT : For optimal use of the Normal Hb A2 Control with the MINICAP FLEX-PIERCING instrument, it is necessary to use one specific tube designed for blood controls and its corresponding cap (see "EQUIPMENT AND ACCESSORIES REQUIRED", Tubes and caps for Controls) and to identify this tube with one Normal Hb A2 Control bar code label.

The Normal Hb A2 Control should be used as follows :

- Apply the reconstituted Normal Hb A2 Control in a tube designed for blood control.
- Close the tube with its cap.
- Place the tube (identified with the Normal Hb A2 Control bar code label), in position No. 28 on a MINICAP FLEX-PIERCING rotating sampler ("Control" position with centering ring).
- Pour 5 mL of MINICAP HEMOGLOBIN(E) hemolysing solution in a hemolysing tube (identified with the hemolysing solution bar code label) without introducing air bubbles and place it in position No. 27 on the rotating sampler ("Diluent / Solution" position without any centering ring) (A message will be displayed if the tube or the hemolysing solution is missing).

IMPORTANT : Ensure the absence of foam in the tube before placing it on the rotating sampler.

- Slide the rotating sampler into the MINICAP FLEX-PIERCING instrument.
- Close the doors of the MINICAP FLEX-PIERCING instrument, the analysis starts automatically.
- In the window which appears on the screen, select the number of analyses of the Normal Hb A2 Control to perform, according to the following indications, and validate :
 - 1 analysis of the Normal Hb A2 Control before starting a new analysis sequence,
 - 2 analyses after having changed the analysis buffer vial (even if the lot number is identical) or the technique, after a cleaning sequence with CAPICLEAN, after a software upgrade or after capillaries activation,
 - 3 analyses for the first use of the "HEMOGLOBIN(E)" analysis program with the MINICAP FLEX-PIERCING instrument or after a prolonged stoppage (over 1 week).

The results are then automatically considered by the software for the data analysis.

IMPORTANT : The hemoglobin A fraction of the Normal Hb A2 Control must show a minimal optical density (OD) of 0.10. Under this value, the recentering of the electrophoretic pattern will not occur correctly. When analysing samples, the identification of hemoglobin fractions, Hb A, Hb F, Hb A2 and Hb C and also the determination of the migration zone of other variants, may be impossible or wrong (see the paragraph RESULT ANALYSIS).

IMPORTANT : For optimal use of the Normal Hb A2 Control, it is necessary to use one bar code label intended to identify the tube for control which contains the Hb A2 Control (close the tube with its specific cap before using it).

After the installation of MINICAP FLEX-PIERCING instrument, during the first sequence of blood sample analysis, a red warning signal will appear if hemoglobin A is absent in one of the samples (and the recentering of the electrophoretic pattern will not be possible, see the paragraph « Result analysis »).

It is then recommended to analyze a blood sample with hemoglobin A on the concerned capillary and to analyze again the sample without hemoglobin A by placing it in a position corresponding to a capillary which has already detected hemoglobin A.

Quality control :

The Normal Hb A2 Control should be used as a normal human blood. After reconstitution, use directly the Normal Hb A2 Control as a blood sample to analyze or as a migration control (in position No. 28 on the rotating sampler, see paragraph before). It will be automatically diluted with hemolysing solution.

It is recommended to include one analysis of Normal Hb A2 Control. The values obtained must fall within the range provided with each batch of Hb A2 Control.

IMPORTANT : For optimal use of the Normal Hb A2 Control placed on the rotating sampler in positions No. 1 to 26, it is necessary to use one bar code label intended to identify the tube for control which contains the Hb A2 Control (close the tube with its specific cap before using it).

Storage, stability and signs of deterioration

See the Normal Hb A2 Control package insert.

IMPORTANT : After the analysis of the Normal Hb A2 Control with the MINICAP FLEX-PIERCING instrument (for migration control or for quality control), store the control at 2 - 8 °C or at - 18 / - 30 °C without any delay, according to the Normal Hb A2 Control package insert.

WARNING : No test method can provide an absolute assurance of the absence of HIV, hepatitis B and C or other infectious agents. Therefore, handle the Normal Hb A2 Control as a hazardous biological material.

This control was found negative on assays approved by FDA or EU equivalent regulatory agency :

- against hepatitis B surface antigen ;
- for antibody to HCV ;
- for antibody to HIV1 and HIV2.

2. DISTILLED OR DEIONIZED WATER**Use**

For rinsing capillaries in the SEBIA MINICAP FLEX-PIERCING automated instrument, for capillary electrophoresis.

It is recommended to use filtered distilled or deionized water (on a filter with a porosity $\leq 0.45 \mu\text{m}$) and with a conductivity lower than $3 \mu\text{S/cm}$, which corresponds to a resistivity higher than $0.33 \text{M}\Omega\cdot\text{cm}$.

To prevent microbial proliferation, change the water every day.

For optimal operation, add CLEAN PROTECT (SEBIA, PN 2059, 1 vial of 5 mL) in distilled or deionized water (see the instructions for use of CLEAN PROTECT) or use directly the ready to use CAP|protect* solution (SEBIA, PN 2061 : 2 containers of 5 L of distilled water with CLEAN PROTECT).

IMPORTANT : Before filling the rinse container, it is recommended to wash it with plenty of distilled or deionized water.

* *NOTE : The CAP|protect solution can also be used to dilute the stock wash solution. Then, in that case, the diluted wash solution may show a transient more or less marked yellow colour without any adverse effects on its performance.*

3. MINICAP FLEX-PIERCING CAPICLEAN**Composition**

The vial of CAPICLEAN concentrated solution (MINICAP FLEX-PIERCING CAPICLEAN, SEBIA, PN 2251, 25 mL) contains : proteolytic enzymes, surfactants and additives nonhazardous at concentrations used, necessary for optimum performances.

Use

For sample probe cleaning in the SEBIA MINICAP FLEX-PIERCING automated instrument, for capillary electrophoresis, during the CAPICLEAN cleaning sequence.

IMPORTANT : Launch a CAPICLEAN cleaning sequence at least once a week and at maximum once a day, or after every 500 analyses when performed within less than one week.

See the instruction sheets of MINICAP FLEX-PIERCING CAPICLEAN, SEBIA.

WARNING : Do not use a capped tube.

IMPORTANT : For optimal use of the CAPICLEAN solution with the MINICAP FLEX-PIERCING instrument, it is necessary to use one bar code label intended to identify the 100 mm tube which contains the diluted CAPICLEAN solution.

Storage, stability and signs of deterioration

Store CAPICLEAN refrigerated (2 - 8 °C). It is stable until the expiration date indicated on the vial label. DO NOT FREEZE.

Precipitate or combined particles in suspension (flocules) may be observed in the CAPICLEAN vial without any adverse effects on its utilization.

Do not dissolve this precipitate or these particles. It is recommended to collect only the supernatant.

For later use, store the tube containing the diluted solution at 2 - 8 °C. It must be used within the day.

4. SODIUM HYPOCHLORITE SOLUTION (for sample probe cleaning)**Preparation**

Prepare a sodium hypochlorite solution (2 % to 3 % chloride) by diluting 250 mL 9.6 % chloride concentrated solution to 1 liter with cold distilled or deionized water.

Use

For the sample probe cleaning in the MINICAP FLEX-PIERCING instrument (weekly maintenance in order to eliminate adsorbed proteins from the probe).

See the SEBIA MINICAP FLEX-PIERCING instruction manual.

WARNING : Do not use a capped tube.

- Apply in a 100 mm tube 2 mL of diluted chlorinated solution previously prepared.
- Place the 100 mm tube (identified with one bar code label specific to the sodium hypochlorite solution) on the rotating sampler of MINICAP FLEX-PIERCING.
- Position a new reagent cup on the automated loading system for cups of MINICAP FLEX-PIERCING (a message will be displayed if the reagent cup is missing).
- Slide the rotating sampler into the MINICAP FLEX-PIERCING instrument.
- Close the doors of the MINICAP FLEX-PIERCING instrument, the cleaning sequence starts automatically.

IMPORTANT : For optimal use of the sodium hypochlorite solution with the MINICAP FLEX-PIERCING instrument, it is necessary to use one bar code label intended to identify the 100 mm tube which contains the solution.

Storage, stability and signs of deterioration

Store the working chlorinated solution at room temperature in a closed container, it is stable for 3 months. Avoid storage in sunlight, close to heat and ignition source, and to acids and ammonia.

5. CAPILLARIES / MINICAP WASH SOLUTION

Preparation

Each vial of the stock CAPILLARIES / MINICAP Wash solution (SEBIA, PN 2052, 2 vials, 75 mL) should be diluted up to 750 mL with distilled or deionized water.

After dilution, the wash solution contains an alkaline solution pH \approx 12.

For MINICAP FLEX-PIERCING, it is convenient to dilute only 25 mL of the stock solution to 250 mL with distilled or deionized water.

Use

For washing the MINICAP FLEX-PIERCING capillaries.

IMPORTANT : Before filling the wash solution container, it is recommended to wash the opening of the container, the connector and the tube with plenty of distilled or deionized water to avoid salts deposit.

Storage, stability and signs of deterioration

Store the stock and working wash solutions in closed containers at room temperature or refrigerated. The stock wash solution is stable until the expiration date indicated on the kit or wash solution vial label.

Working wash solution is stable for 3 months.

Discard working wash solution if it changes its appearance, e.g., becomes cloudy due to microbial contamination.

6. SALINE

Preparation

Make 0.15 M (0.9 g/dL) NaCl solution in distilled or deionized water.

Use

To wash red blood cells before storage at - 70 / - 80 °C, if necessary.

Storage, stability and signs of deterioration

Store saline at room temperature or refrigerated.

Discard after 3 months or if it changes its appearance, e.g., becomes cloudy due to microbial contamination. For longer storage periods, add sodium azide, 0.1 g/dL.

OPTIONAL REAGENT BUT NOT SUPPLIED

WARNING: See the safety data sheet.

PATHOLOGICAL Hb A2 CONTROL

The Pathological Hb A2 Control, SEBIA, PN 4779, can be used for the migration control, in addition or as a replacement of the Normal Hb A2 Control. For its utilization for the migration control or quality control, the Pathological Hb A2 Control should be used like the Normal Hb A2 Control, see the previous paragraph "NORMAL Hb A2 CONTROL".

See the instructions for use of the Pathological Hb A2 Control for additional information.

NOTES :

The assays that were performed for the validation of reagents demonstrated that, for the different solutions and using an adapted equipment for the reconstitution volume, a variation of \pm 5 % on the final volume has no adverse effect on the analysis.

The distilled or deionized water used to reconstitute solutions, must be free of bacterial proliferation and mold (use a filter \leq 0.45 μ m) and have a conductivity lower than 3 μ S/cm, which corresponds to a resistivity higher than 0.33 M Ω .cm.

EQUIPMENT AND ACCESSORIES REQUIRED

- MINICAP FLEX-PIERCING instrument, SEBIA, PN 1232.
- Rotating sampler supplied with MINICAP FLEX-PIERCING, equipped with centering rings for MINICAP FLEX-PIERCING (specific for the MINICAP HEMOGLOBIN(E) procedure) for 13 mm diameter tubes.
- MINICAP FLEX-PIERCING centering rings for 11 mm diameter tubes, SEBIA (PN 1612) for 11 mm diameter tubes.
- Container Kit supplied with MINICAP FLEX-PIERCING : Rinse container (to fill with distilled or deionized water) and waste container.
- Collection tubes with 13 mm diameter and their corresponding caps (maximal length of tube with cap : 100 mm, maximal diameter of cap : 17 mm) : for example, BD Vacutainer, Terumo Venosafe 5 mL, Greiner Bio-one Vacuette 1, 2, 3 or 4 mL or Sarstedt S-Monovette 4 mL tubes (13 x 75 mm), or collection tubes with 11 mm diameter and their corresponding caps (maximal length of tube with cap : 100 mm, maximal diameter of cap : 17 mm) : for example, Sarstedt S-Monovette 2,7 mL or Kabe Labortechnik Primavette S 2,6 mL tubes (11 x 66 mm), or collection tubes with equivalent dimensions approved for clinical assays.
- Tubes and caps for Controls, SEBIA, PN 9205 : 500 conical tubes and their caps to analyze blood controls and particular samples with the MINICAP FLEX-PIERCING instrument.
- Hemolysing tubes (with 8 to 16 mm diameter and 50 to 100 mm length).

8. "AUTOMATIC LOW VOLUME" bar code labels, SEBIA (20 units), PN 9208 : labels intended to identify samples with volume less than 1 mL for an analysis using the "automatic dilution" mode (VS \geq 8.61).
9. "MANUAL LOW VOLUME" bar code labels, SEBIA (20 units), PN 9209 : labels intended to identify samples with volume less than 1 mL for an analysis using the "manual dilution" mode (VS \geq 8.61).
10. MINICAP Reagent cups / 125 (3), SEBIA, PN 2281.
11. Lids for bins for used reagent cups, SEBIA (12 units), PN 2286 : lids to close the bins containing used cups.
12. Pump filter for MINICAP FLEX-PIERCING, SEBIA (2 units), PN 1620.

SAMPLES FOR ANALYSIS

Sample collection and storage

Fresh anticoagulated whole blood samples collected in tubes containing K_2 EDTA or K_3 EDTA as anticoagulant are recommended for analysis. Avoid anticoagulants containing iodoacetate. Blood must be collected according to established procedures used in clinical laboratory testing.

Samples may be stored for up to 7 days between 2 and 8 °C.

NOTE : Samples should not be stored at room temperature.

Progressive hemoglobins (Hb) degradation may occur for samples stored between 2 to 8 °C.

When the blood sample is stored for more than 7 days at 2 - 8 °C :

- a weak fraction, corresponding to methemoglobin, appears in the Hb S migration zone,
- when Hb C is present, a fraction corresponding to degraded Hb C appears more anodic than Hb A2 which does not interfere with it (Z(E) zone, see the table in paragraph "Interpretation"),
- when Hb O-Arab is present, a fraction corresponding to degraded Hb O-Arab appears in the Hb S migration zone (Z(S) zone, see the table in paragraph "Interpretation"),
- when Hb E is present, a fraction corresponding to degraded Hb E appears in the Z(D) zone (see the table in paragraph "Interpretation"),
- when Hb S is present, a fraction corresponding to degraded Hb S appears in the Hb F migration zone (Z(F) zone, see the table in paragraph "Interpretation"),
- when Hb A is present, a fraction corresponding to degraded Hb A ("aging fraction" of Hb A) appears more anodic (Z11 zone, see the table in paragraph "Interpretation"),
- when Hb F is present (in blood samples from newborn babies), a fraction appears in the Hb A migration zone (Z(A) zone, see the table in paragraph "Interpretation") due to the sample degradation.

When stored for more than 10 days, viscous aggregates in red blood cells are observed ; it is necessary to discard them before the analysis.

For longer storage, whole blood samples (without any preparation) or washed red blood cells can be frozen at - 70 / - 80 °C within 8 hours of collection. Frozen whole blood samples and red blood cells are stable for 3 months maximum at - 70 / - 80 °C.

IMPORTANT : For optimal storage of samples, store them at - 70 / - 80 °C. Do not store at - 20 °C (see BIBLIOGRAPHY, J. Bardakdjian-Michau *et al*, 2003).

Preparation of red blood cells for storage at - 70 / - 80 °C :

Wash red blood cells according to the following procedure: Centrifuge anticoagulated blood to obtain a red blood cells pellet ; discard the plasma ; wash the red blood cells (RBC) 2 times with 10 volumes of saline (centrifuge after each washing step) ; discard the excess of saline over the red blood cells pellet and vortex them before freezing.

Sample preparation

- Use directly whole blood samples.
- Check that all the tubes contain at least 1 mL of blood and are perfectly closed.
- **Vortex for 5 seconds blood samples stored at 2 - 8 °C for one week or stored at - 70 / - 80 °C.**

WARNING : *The tubes must be closed with their corresponding caps designed for the MINICAP HEMOGLOBIN(E) procedure with the MINICAP FLEX-PIERCING instrument (see EQUIPMENT AND ACCESSORIES REQUIRED).*

Particular cases:

IMPORTANT : The following samples must be identified with the "LOW VOLUME AUTO" or "LOW VOLUME MANUAL" bar code label ; these labels allow the instrument to apply respectively the automatic or manual dilution mode according to the sample preparation procedure. These samples must be placed on the rotating sampler at the beginning of an analysis series and the bar code of each tube must be visible in the opening of the rotating sampler. Without any bar code label, the analysis should be affected.

Analysis of samples without any Hb A and with Hb F < 3 % or without any Hb A2 (these samples are perfectly quantified but not identified by zones). To identify hemoglobin fractions in a sample without any Hb A and with Hb F < 3 % or without any Hb A2, it is recommended to prepare this sample according to one of the two following procedures:

- Vortex for 5 seconds the whole blood sample.
 - In a conical tube for control (identified with a "LOW VOLUME AUTO" bar code label), mix one volume (100 μ L) of whole blood to analyze with one volume (100 μ L) of Normal Hb A2 Control and cap the tube.
 - Vortex for 5 seconds.
 - Place the tube on the rotating sampler of the MINICAP FLEX-PIERCING instrument at the beginning of an analysis series and slide the rotating sampler into the instrument to run the analysis without any delay.
 - Close the doors of the MINICAP FLEX-PIERCING instrument, the analysis starts automatically.
- The results are then automatically considered by the software for the data analysis.

IMPORTANT : For a sample without any Hb A, Hb F or Hb A2 prepared according to this procedure, the result obtained with the mixed sample will enable presumptive variant identification due to the positioning of the hemoglobins fractions in the appropriate identification zones. Do not report the relative quantification from the mixed sample result.

The relative quantification of hemoglobins should be reported utilizing the initial, unmixed sample result (without any dilution in the blood control).

Analysis of samples stored at -70 / - 80 °C :

Whole blood :

Vortex for 5 seconds the tube with thawed whole blood and perform the analysis according to the standard procedure.

Red blood cells :

To analyze a red blood cells sample stored at -70 / - 80 °C, prepare it according to the following procedure :

- Vortex for 5 seconds the tube with thawed red blood cells.
 - In a conical tube for control (identified with a "LOW VOLUME MANUAL" bar code label), mix one volume (50 μ L) of red blood cells with 8 volumes (400 μ L) of MINICAP HEMOGLOBIN(E) hemolysing solution and cap the tube.
 - Vortex for 5 seconds.
 - Place the tube on the rotating sampler of the MINICAP FLEX-PIERCING instrument at the beginning of an analysis series and slide the rotating sampler into the instrument to run the analysis without any delay.
 - Close the doors of the MINICAP FLEX-PIERCING instrument, the analysis starts automatically.
- The results are then automatically considered by the software for the data analysis.

Analysis of samples with volume below 1 mL :

- Vortex for 5 seconds the whole blood sample and prepare the sample according to its volume following one of the 2 protocols.

1. Volume of sample comprised between 200 μ L and 1 mL :

- Apply at least 200 μ L of whole blood to analyze in a conical tube for control (identified with a "LOW VOLUME AUTO" bar code label) and cap the tube.
- Place the tube on the rotating sampler of the MINICAP FLEX-PIERCING instrument at the beginning of an analysis series and slide the rotating sampler into the instrument to run the analysis without any delay.
- Close the doors of the MINICAP FLEX-PIERCING instrument, the analysis starts automatically.

or

2. Volume of sample below 200 μ L :

- Apply 50 μ L of whole blood to analyze in a conical tube for control (identified with a "LOW VOLUME MANUAL" bar code label) and 250 μ L of MINICAP HEMOGLOBIN(E) hemolysing solution and cap the tube.
- Vortex for 5 seconds.
- Place the tube on the rotating sampler of the MINICAP FLEX-PIERCING instrument at the beginning of an analysis series and slide the rotating sampler into the instrument to run the analysis without any delay.
- Close the doors of the MINICAP FLEX-PIERCING instrument, the analysis starts automatically.

The results are then automatically considered by the software for the data analysis.

NOTES : The minimal volume of sample to analyze in a conical tube is 200 μ L. Without any bar code label on the conical tube, the sample cannot be identified and the dilution mode (automatic or manual according to the sample preparation procedure) cannot be applied.

Samples to avoid

- Avoid coagulated blood samples.
 - Avoid aged, improperly stored blood samples ; the automated hemolysis of samples may be disturbed by viscous aggregates in red blood cells. Then, degradation products (as artefacts) may affect the electrophoretic pattern.
- In these 2 cases, aggregates in red blood cells may affect the collection of the sample by the probe.
- Do not analyze directly tubes containing less than 1 mL of blood sample, the analysis should be affected (see particular cases).

NOTE : Collection tubes and centrifugation parameters for biological samples are described in the available documentation on pre-analytical phase for bio-medical analysis (data provided by the tube manufacturers, guides and recommendations on biological sample collection...). Without any indication in the instructions for use on the type of tube to use or on the centrifugation, please refer to this documentation and for the dimensions of tube to use, refer to the SEBIA document "Characteristics of tubes to use according to the instrument". The pre-analytical phase must be performed according to the state of art, the different recommendations, including those provided by the tube manufacturers, and applicable regulations.

PROCEDURE

The MINICAP FLEX-PIERCING instrument is a multiparameter instrument for hemoglobins analysis on parallel capillaries. The hemoglobins assay uses 2 capillaries to run the samples.

The sequence of automated steps is as follows :

- Bar code reading of rotating sampler and sample tubes (for up to 26 tubes) ;
- Mixing of blood samples before analysis ;
- Sample hemolysis and dilution from primary tubes into reagent cups ;
- Capillary washing ;
- Injection of hemolyzed samples ;
- Hemoglobin separation and direct detection of the separated hemoglobins on capillaries.

The manual steps include :

- Placement of sample tubes (with caps) in rotating sampler in positions 1 to 26 ;
- Placement of the uncapped hemolysing solution tube in rotating sampler in position 27 ;
- **Set up the Normal Hb A2 Control or the Pathological Hb A2 Control in rotating sampler in position 28;**
- Placement of the rotating sampler in the MINICAP FLEX-PIERCING instrument ;
- Removal of the sample tubes after analysis ;
- Removal and closing of the bins for used cups.

PLEASE CAREFULLY READ THE MINICAP FLEX-PIERCING INSTRUCTION MANUAL.

I. PREPARATION OF MINICAP ANALYSIS

1. Switch on MINICAP FLEX-PIERCING instrument and computer.
2. In order to start the instrument, position at least one new reagent cup on the automated loading system for cups of MINICAP FLEX-PIERCING (a message will be displayed if a reagent cup is missing).
3. Set up the software, the instrument automatically starts.
4. The MINICAP HEMOGLOBIN(E) kit is intended to run with "HEMOGLOBIN(E)" analysis program from the MINICAP FLEX-PIERCING instrument. To select "HEMOGLOBIN(E)" analysis program and place the MINICAP HEMOGLOBIN(E) buffer vial in position "B2" in the instrument, please read carefully the MINICAP FLEX-PIERCING instruction manual and follow the instructions displayed on the screen.
5. Position new reagent cups on the automated loading system for cups of MINICAP FLEX-PIERCING (a message will be displayed if the reagent cups are missing).
6. **Position a bin for used cups in MINICAP FLEX-PIERCING at the location intended for this purpose.**
7. Check the fill level of the reagent vials, add reagent if necessary and empty the waste container. In the window "Check reagent levels", update the software by moving the cursor buttons. Replace the filter of the rinsing system, if necessary.
8. The rotating sampler contains 28 positions for sample tubes:
 - Place up to 26 capped sample tubes with whole blood on the rotating sampler with specific centering rings (**positions No. 1 to 26**), the bar code of each tube must be visible in the opening of the rotating sampler.
 - Pour the MINICAP HEMOGLOBIN(E) hemolysing solution in an uncapped hemolysing tube, identified with the hemolysing solution bar code label, without introducing air bubbles: 2 mL for the analysis of 1 or 2 samples, 5 mL for the analysis of 12 samples. Place this tube in **position No. 27** without any centering ring on the rotating sampler ("Diluent / Solution" position).

NOTE : 5 mL of hemolysing solution applied in a 13 x 75 mm tube will allow to perform 12 analyses ; 10 mL of hemolysing solution applied in a 16 x 100 mm tube will allow to perform 24 analyses.

IMPORTANT : Ensure the absence of foam in the tube before placing it on the rotating sampler.

- **Position the Normal Hb A2 Control or the Pathological Hb A2 Control in position No. 28 on the rotating sampler ("Control" position) and select the number of analyses of the control to perform according to the indications previously described, see "Normal Hb A2 Control" paragraph, part "Migration control".**

IMPORTANT : If a tube is missing in position No. 1 (sample tube), in position No. 27 (hemolysing solution tube) and in position No. 28 (blood control tube), the analysis can not start and a message will be displayed.

NOTE : When using a control blood, it is necessary to use the specific bar code label.

9. Slide the rotating sampler into the MINICAP FLEX-PIERCING instrument.
10. Close the doors of the MINICAP FLEX-PIERCING instrument, the analysis starts automatically.
11. After the analysis, remove the rotating sampler with analyzed sample tubes.
12. If necessary, take off carefully the bin containing used reagent cups, close it tightly with the corresponding lid and discard it.

WARNING : Bins containing used reagent cups with biological samples have to be handled with care.

DILUTION - MIGRATION - DESCRIPTION OF THE AUTOMATED STEPS

1. Bar codes are read on both rotating sampler and sample tubes.
2. Mixing of tubes.
3. Samples are diluted in hemolysing solution and the sample probe is rinsed after each sample.
4. Capillaries are washed.
5. Diluted samples are injected into capillaries.
6. Migration is carried out under constant voltage for about 8 minutes and the temperature is controlled by Peltier effect.
7. Hemoglobins are detected directly by scanning at 415 nm and an electrophoretic profile appears on the screen of the system.

NOTE : These automated steps are described for the two first analyzed sample tubes. The electrophoretic patterns appear after about 20 minutes from the start of the analysis. For the following sample tubes, the first three steps (bar code reading, mixing and hemolysis of blood samples) are performed during the analysis of the 2 previous samples.

II. RESULT ANALYSIS

At the end of the analysis, relative quantification of individual hemoglobin fractions is automatically performed and profiles can be analyzed. The hemoglobin fractions Hb A, Hb F, Hb A2 and Hb C are automatically identified. The Hb A fraction is centered in the middle of the review window and Hb A2 is adjusted at a fixed position against that of Hb A. In the absence of Hb A and when Hb F is present ($\geq 3\%$), the recentering of the pattern is made with Hb F and Hb A2 peaks that are placed at fixed positions.

The resulting electrophoregrams are evaluated visually for pattern abnormalities.

The electrophoretic patterns are colored :

- in Cyan when the number of fractions / peaks is that which is configured by default for the procedure (2 fractions for HEMOGLOBIN(E) procedure, for example),
- in Magenta when the number of fractions / peaks is not that which is configured by default for the procedure.

With HEMOGLOBIN(E) procedure, the Hb F peak is orange (identified by «Hb F or variant») when the age of the patient is unknown and blue (identified by «Hb F») when the age of the patient is known and the fraction / peak is lower than 2 %.

The potential positions of the different hemoglobin variants (identified in zones called Z1 to Z15) are shown on the screen of the system and indicated on the result ticket. The table in paragraph "Interpretation" shows known variants which may be present in each corresponding zone. When the software identifies a hemoglobin fraction in a defined zone, the name of this zone is framed.

Patterns are automatically adjusted with regard to Hb A and Hb A2 fractions, or with regard to Hb F and Hb A2 fractions as the case may be, to facilitate their interpretation:

- when Hb A and / or Hb A2 fractions are not detected on an electrophoretic pattern and / or when Hb F (with no Hb A) is not detected or is at a level $< 3\%$,
 - a yellow warning signal appears,
 - the adjustment of the pattern is performed using the position of the Hb A fraction on the two previous patterns obtained with the same capillary,
 - no fraction is identified (except when Hb C is detected: in this case, Hb A2 and Hb C fractions are identified),
 - the different migration zones (Z1 to Z15) do not appear neither on the screen of the system, nor on the ticket result.
- when Hb F is detected at a level $\geq 3.0\%$, without any detection of Hb A (no Hb A or Hb A at a low level) on an electrophoretic pattern,
 - the adjustment of the pattern is performed using the position of the Hb F and Hb A2 fractions,
 - Hb F and Hb A2 fractions are placed at fixed positions,
 - Hb F and Hb A2 fractions are identified,
 - the different migration zones (Z1 to Z15) are indicated on the screen of the system and on the ticket result by the same way of patterns with Hb A,
 - abnormal fractions are grey-dashed and identified using their migration zone (a fraction detected in Z(D) zone is called "Z(D) zone" for example),
 - when a rare variant migrates in the Hb A2 migration zone, the different migration zones (Z1 to Z15) do not appear neither on the screen of the system, nor on the ticket result.
- when the adjustment is not possible,
 - a red warning signal appears,
 - no fraction is identified,
 - the different migration zones (Z1 to Z15) do not appear neither on the screen of the system, nor on the ticket result.
 Call SEBIA.
- when optical density (OD) is insufficient on a migration control electrophoretic pattern (obtained with the Normal Hb A2 Control or the Pathological Hb A2 Control, identified with its bar code label and analyzed in position No. 28 in the rotating sampler),
 - a warning message is displayed in order to consider or remove this analysis for the determination of Hb A fraction position,
 - a purple warning signal appears on the review window,
 - Hb A and Hb A2 fractions are not identified (except when the analysis is considered by the operator),
 - the different migration zones (Z1 to Z15) do not appear neither on the screen of the system, nor on the ticket result (except when the analysis is considered by the operator).

In all these cases, the different migration zones (Z1 to Z15) do not appear neither on the screen of the system, nor on the ticket result.

On the electrophoretic pattern, the curves of Hb A2 and Hb C fractions, are calculated and redrawn by fitting with adjustment (or fitted) and are overlaid with the native curve. This display allows the Hb A2 fraction quantification if Hb C is present in the sample.

WARNING : In some cases of hemoglobin C (homozygous) or after a technical problem, the hemoglobins A2 and C are not fitted ; these fractions are then under-quantified. It is then recommended to quantify the Hb A2 fraction by using another technique.

PLEASE CAREFULLY READ THE MINICAP FLEX-PIERCING INSTRUCTION MANUAL.

III. END OF ANALYSIS SEQUENCE

At the end of each analysis sequence, the operator must initiate the "shut down" procedure of the MINICAP FLEX-PIERCING in order to store capillaries in optimal conditions.

IMPORTANT : Position at least one new reagent cup on the automated loading system for cups of MINICAP FLEX-PIERCING (a message will be displayed if a reagent cup is missing).

IV. FILLING OF REAGENT CONTAINERS

The MINICAP FLEX-PIERCING instrument has a reagent automatic control.

IMPORTANT :

- Please refer to the instructions for replacement of reagent containers respecting color code for vials and connectors.
- Replace the filter of the rinsing system **every 500 analyses**, with a pump filter for MINICAP FLEX-PIERCING (see paragraph "EQUIPMENT AND ACCESSORIES REQUIRED").

A message will be displayed when it is necessary to perform one of the following tasks :

- Place a new buffer vial and / or ;
- Fill the container with working wash solution and / or ;
- Fill the container with filtered distilled or deionized water for rinsing capillaries and / or ;
- Empty the waste container and / or ;
- Replace the filter of the rinsing system.

WARNING: Do not use marketed deionized water, such as water for ironing for example (risk of important capillaries damage). Use only water with ultrapure quality, such as injection grade water.

IMPORTANT : Before filling the rinse container, it is recommended to wash it with plenty of distilled or deionized water.

PLEASE CAREFULLY READ THE MINICAP FLEX-PIERCING INSTRUCTION MANUAL.

QUALITY CONTROL

It is advised to include into each run of samples, an assayed control blood (for example, a blood sample containing hemoglobins A, F, C and S, such as Hb AFSC Control, SEBIA, PN 4792, or a normal blood sample, the Normal Hb A2 Control, SEBIA, PN 4778 or the Pathological Hb A2 Control, SEBIA, PN 4779).

RESULTS

Values

Direct detection at 415 nm in capillaries yields relative concentrations (percentages) of individual hemoglobin zones.

Reference values for individual major electrophoretic hemoglobin zones in the MINICAP FLEX-PIERCING system have been established from a healthy population of 113 adults (men and women), whose hemoglobin values (as established by HPLC) were normal :

Hemoglobin A : comprised between 96.8 and 97.8 %

Hemoglobin F : < 0.5 % (*)

Hemoglobin A2 : comprised between 2.2 and 3.2 %

(*) See *Interference and limitations*

It is recommended that each laboratory establish its own threshold values.

NOTE : Normal values have been established using the standard parameters of the PHORESIS software (smoothing 0 and hemoglobin fractions automatic quantification with HEMOGLOBIN(E) analysis program).

WARNING : Normal (reference) values must be considered only when hemoglobin variants are absent.

Interpretation

See *ELECTROPHORETIC PATTERNS*, figures 1 – 15.

The different migration zones of hemoglobin variants (called Z1 to Z15) are shown on the screen of the system and on the result ticket. Passing the mouse cursor over a zone name displays icon information containing possible hemoglobin variants that could be seen in this zone.

For each fraction, the position of the top of the peak (position of the maximum) defines the migration zone.

See the table showing the potential variants located in each zone.

With PHORESIS VS ≥ 9.15 , this table lists 525 different hemoglobin variants. Due to the history of their discovery, some variants may have many names. A second name is added between brackets close to the main name (for example, in zone Z(D), Hb Korle-Bu (G-Accra)). Other names are not listed in this table.

In zone Z(A), variants are listed in alphabetical order.

For other zones, variants are sorted in main fractions and minor fractions and presented by migration order from most cathodic variants to most anodic variants.

For variants with a main fraction that migrates in zone Z(A), their minor fractions which migrate in zone Z(A2) are not indicated.

- The symbol "[]" indicates a hidden or partially hidden peak due to similar migration to normal Hb A or Hb A2 fraction. A partially hidden fraction corresponds to a more or less important shoulder of the normal fraction.
- The symbol "##" indicates the display in icon information of several visible fractions from the same variant, generally present in different zones (for example, alpha-chain variant with a second visible peak as Hb Q-India, or unstable variant as Hb Sabine and Hb Köln). Not concerned: beta chain variants except unstable variants, gamma and delta chain variants and delta-beta hybrids, alpha chain variants without second peak visible on the electrophoretic pattern.
- The symbol "!!!" alerts the potential risk of a migration zone shift for a rare variant located in a zone boundary. Additionally, the migration variation of a variant (± 1 point) depends on its percentage. For example, Hb Willamette, located on the far right of zone Z(F), may migrate in zone Z(D) when its percentage has decreased in case of an associated thalassemia.

These symbols are explained in the "Captions" icon information located in the upper left side of the review window.

1. Qualitative abnormalities: Hemoglobinopathies

Most hemoglobinopathies are due to substitution by mutation of a single amino acid in one of the four types of polypeptide chains (1, 2, 4, 9, 12). The clinical significance of such a change depends on the type of amino acid and the site involved (13). In clinically significant disease, either the α -chain or the β -chain is affected.

More than 1400 variants of adult hemoglobin have been described (6, 14). The first abnormal hemoglobins studied and the most frequent have an altered net electric charge, leading to an easy detection by electrophoresis.

There are five main abnormal hemoglobins which present a particular clinical interest: S, C, E, O-Arab and D.

The MINICAP HEMOGLOBIN(E) kit is intended for the identification of hemoglobinopathies and thalassemias.

Hemoglobin S

Hemoglobin S is the most frequent hemoglobinopathy. It is due to the replacement of one glutamic acid (an acidic amino acid No. 6) of the β -chain by valine (a neutral amino acid): when compared to Hb A, its isoelectric point is elevated and its total negative charge decreased with the analysis pH. Its electrophoretic mobility is therefore increased in the capillary and this hemoglobin migrates faster than A fraction.

With alkaline buffered MINICAP HEMOGLOBIN(E) procedure, hemoglobin S migrates between A and A2 fractions, next to Hb A2.

Hemoglobin C

One glutamic acid of the β -chain is replaced by lysine (a basic amino acid No. 6): its mobility is strongly reduced. When compared to Hb A, its isoelectric point is highly elevated and its total negative charge decreased with the analysis pH. Its electrophoretic mobility is therefore increased in the capillary and this hemoglobin is faster than A fraction which allows its differentiation. Hemoglobins C, E and O-Arab are not superimposed on the electrophoretic pattern and are easily identified.

Hemoglobin E

One glutamic acid of the β -chain (No. 26) is replaced by lysine. With MINICAP HEMOGLOBIN(E) procedure, hemoglobin E migrates just anodically behind hemoglobin A2 and is totally separated from it. Then, when hemoglobin E is present, A2 fraction can be measured to detect β thalassemia.

Hemoglobin O-Arab

One glutamic acid of the β -chain (No. 121) is replaced by lysine. With MINICAP HEMOGLOBIN(E) procedure, hemoglobin O-Arab migrates exactly like hemoglobin A2. In such a case, hemoglobin A2 can not be quantified. When this fraction is $> 10.5\%$, hemoglobin O-Arab must be suspected. Note that Hb O-Arab migrates separately from hemoglobins C and E.

Hemoglobin D (-Los Angeles)

One glutamic acid of the β -chain (No. 121) is replaced by glutamine. With MINICAP HEMOGLOBIN(E) procedure, hemoglobin D (called D-Punjab, D-Los Angeles, D-Chicago or D-Portugal) migrates more anodically than hemoglobin S, this property allows to differentiate hemoglobins S and D.

2. Quantitative abnormalities : Thalassemias

Thalassemias constitute a quite heterogeneous group of genetic disorders characterized by decreased synthesis of one type of the globin chains. The molecular mechanism of this decrease has not been fully described.

There are two types of thalassemia syndromes :

Alpha-thalassemias

They are characterized by the decrease of synthesis of the α -chains, consequently affecting the synthesis of all normal hemoglobins.

The excess of synthesis of the β - and γ -chains in relation to α -chains induces the formation of tetramers without any α -chain:

- hemoglobin Bart's = γ_4 ,
- hemoglobin H = β_4 .

Hemoglobin H presents a low isoelectric point ; with MINICAP HEMOGLOBIN(E) procedure, it migrates more anodically than hemoglobin A (and may appear as one or several fractions).

Beta-thalassemias

They are characterized by the decrease of synthesis of the β -chains. Only hemoglobin A synthesis is affected.

Therefore hemoglobin F and hemoglobin A2 percentages are increased with respect to hemoglobin A.

With MINICAP HEMOGLOBIN(E) procedure, values obtained for different normal hemoglobin fractions allow the detection of beta-thalassemias.

3. Particular cases

- When there is no hemoglobin A in the sample, a small fraction may be observed in anodic position compared with Hb F (in the Z8 zone when migration zones are displayed on the electrophoretic pattern) ; this fraction may be acetylated hemoglobin F which represents about 15 to 25 % of hemoglobin F. The MINICAP FLEX-PIERCING instrument can identify this acetylated hemoglobin separately from the hemoglobin A without any confusion.
- When a small fraction (about 0.5 to 3 %) migrates between hemoglobins F and δA^2 (A2 variant), a hemoglobin A2 variant may be suspected. The percentages of the normal hemoglobin A2 and of the variant are approximately identical for heterozygous patients.
- When a hemoglobin A2 variant is detected (δA^2 or any other A2 variant), it is recommended to add its percentage to hemoglobin A2 for a better beta-thalassemia diagnostic.
- Some hemoglobin variants (such as Hb Camperdown and Hb Okayama) migrate close to Hb A and may not be separated from this hemoglobin.
- Some hemoglobin variants (such as Hb Pörtl-Alegre or degraded Hb S, for example) including homozygous variants such as Hb Q-Thailand, migrate close to Hb F. In the absence of Hb A, the adjustment of the pattern using Hb F and Hb A2 peaks and the display of migration zones prevents any confusion of these variants with Hb F.
- **PHORESIS VS < 9.15 :** Weak hemoglobin fractions which migrate in zone Z12 are sometimes quantified with imprecision (too asymmetric Hb Bart's, for example). In such case, it is necessary to delete automatic quantification and then to quantify the fraction manually. **PHORESIS VS ≥ 9.15 :** In zone Z12, the curve of Hb Bart is calculated and redrawn by fitting with adjustment (or fitted). Fitted fractions are then called "Hb Bart suspected". Narrow fractions with low percentage are not Hb Bart's, they are identified "Z12 zone". Wide fractions with elevated percentage, suspected to be hemoglobin variants, are identified "Hb Bart zone".
- In the MINICAP HEMOGLOBIN(E) technique, for diabetic patients with elevated HbA_{1c} (over 10 %), a small fraction is observed and eventually identified as a peak in Z10 zone.
- The software version (≥ 9.15) allows a specific identification of Hb H in Z15 zone. Fractions with a width over 10 points and a percentage between 0.3 and 32 % are called "Hb H suspected". Fractions with a width below 10 points are not Hb H and are identified "Z15 zone". Wide fractions with a percentage between 10 and 58 %, suspected to be hemoglobin variants, are identified "Abnormal Hb".
- When analysing blood samples from newborn babies, Hb A from samples containing Hb F at high concentrations may be disturbed, especially due to the presence of degraded Hb F in its migration zone. The Hb A percentage indicated by the software may be overestimated. In addition, when hemoglobin variants ($> 4\%$, such as Hb S, Hb C, Hb E or Hb D-Punjab) are present in blood samples containing high Hb F levels ($> 60\%$), it is necessary to perform complementary analyses in order to confirm the presence of Hb A.
- For newborn babies until 6 – 9 months old, it is recommended to analyze different blood samples (collected monthly, for example) in order to check the Hb F concentration. It will allow to verify the decrease of Hb F concentration and the potential presence of a variant. In case of uncertainty, it is advised to confirm by using complementary studies and to analyze parents' blood samples.
- When analyzing blood samples from patients with sickle cell disease before transfusion, a variation of Hb S fraction may be observed for the analyses of the same patient due to the inhomogeneity of this type of sample. It is therefore recommended to homogenize this type of blood sample before the analysis.
- Hb S fraction may appear in a very anodic position in Z(S) zone (in far left of this zone) for the following cases:
 - blood sample with low Hb A level ($< 10\%$) and high Hb S level (for example, blood sample from transfused patient with sickle cell disease, or from patient with S beta-thalassemia) for which the pattern is adjusted with Hb A and Hb A2 peaks, and,
 - blood sample without any Hb A and with high Hb S level for which the pattern is adjusted with Hb F and Hb A2 peaks.

This migration zone, which corresponds to an intermediary zone comprised within Z(S) zone, allows to automatically detect Hb S fractions with modified migration. It is called "Shifted Hb S area" and is indicated by a dash located on the left side of the name Z(S) in the upper section of the Z(S) migration zone. A variant which migrates in this position is identified as "Borderline variant" but not "Z(S) zone" like any other peak which migrates in the rest of the Z(S) zone.

Mix the sample with the Normal Hb A2 Control according to the procedure described in paragraph "SAMPLES FOR ANALYSIS", section "Particular cases", in order to confirm the position of the variant in the Z(S) migration zone. It is necessary to analyze the hematologic state and to perform complementary studies to check the presence of Hb S.

• Examples with increased hemoglobin F (Hb F) (except for newborn babies):

- pregnancy ;
- patients with sickle cell disease, more than 2 years old, with a Hydrexa® (hydroxyurea) treatment and/or transfused and/or producing naturally Hb F increased by compensation ;
- patients, more than 2 years old, with HPFH (hereditary persistence of fetal hemoglobin exhibiting 15 to 35 % Hb F for heterozygous patients) ;
- patients, more than 2 years old, with leukemia (of any type), hereditary hemolytic anemia, diabetes, thyroid disease, hyperactivity of bone marrow, multiple myeloma, cancer with metastases.

Interference and Limitations

- See SAMPLES FOR ANALYSIS.
- Analyze only blood samples contained in collection tubes indicated in the paragraph "EQUIPMENT AND ACCESSORIES REQUIRED" or tubes with equivalent dimensions approved for clinical assays. Call SEBIA technical service for further information on these devices.
- Do not analyze directly tubes containing less than 1 mL of blood sample.
- Avoid aged, improperly stored blood samples ; degradation products (or artefacts) may affect the electrophoretic pattern after 7 days storage.
- After 10 days storage, viscous aggregates composed in red blood cells may appear, they must be discarded before analysis.
- When an abnormal hemoglobin is detected, use other means of identification (e.g., globin chain electrophoresis), or consult or send sample to a specialized laboratory.

IMPORTANT : The analysis of the hematologic state is also necessary, as complementary results.

- The migration of a hemoglobin variant close to Hb A involves an underestimation of Hb A fraction and that of the variant and consequently, an overestimation of Hb A2 fraction. In order to quantify Hb A2 with precision, it is necessary to delete the separate integration of both variants and Hb A, and to quantify these fractions together.
- Some homozygous "S" subjects receive a "Hydrexa® (hydroxyurea) treatment that can induce synthesis of fetal hemoglobin. With MINICAP HEMOGLOBIN(E) procedure, the mobility of the induced hemoglobin F is not different from the physiological hemoglobin F.
- Due to the resolution and sensitivity limits of zone electrophoresis, it is possible that some hemoglobin variants may not be detected with this method.
- In the case of patients with hyperleukocytosis, the migration speed of the sample may be accelerated causing a shift of the profile that may result in a non-recognition of the zones.

Hemoglobin variants observed with Hb A1c and / or HEMOGLOBIN(E) procedures :

Due to the different composition of Hb A1c and HEMOGLOBIN(E) buffers, the electrophoretic mobility of some hemoglobin variants may be different.

The common interfering factors with the MINICAP HEMOGLOBIN(E) procedure performed with the MINICAP FLEX-PIERCING instrument (triglycerides and bilirubin) for hemoglobin fractions analysis were evaluated in studies based on the Clinical Laboratory Standards Institute (CLSI - USA) EP7-A2 guideline "Interference Testing in Clinical Chemistry".

The results are summarized below:

- No qualitative or quantitative interference with the MINICAP HEMOGLOBIN(E) procedure performed with the MINICAP FLEX-PIERCING instrument was detected if triglycerides concentration is equal to or less than 15.57 g/L.
- No qualitative or quantitative interference with the MINICAP HEMOGLOBIN(E) procedure performed with the MINICAP FLEX-PIERCING instrument was detected if bilirubin concentration is equal to or less than 27.7 mg/dL, or 473 µmol/L.

Troubleshooting

Call SEBIA Technical Service of the supplier when the test fails to perform while the instruction for the preparation and storage of materials, and for the procedure were carefully followed.

Kit reagent Safety Data Sheets and information on cleaning and waste disposal, labeling and safety rules applied by SEBIA, packaging for the transportation of biological samples, and instruments cleaning are available on the SEBIA's extranet website : www.sebia.com.

PERFORMANCE DATA

Precision

The precision of the MINICAP HEMOGLOBIN(E) procedure performed with the MINICAP FLEX-PIERCING instrument was evaluated in a study based on the Clinical Laboratory Standards Institute (CLSI - USA) EP5-A2 guideline "Evaluation of Precision Performance of Quantitative Measurements Methods".

The means, standard deviations and coefficients of variation (CV's %) (n = 80) were calculated for percentages (%) of hemoglobin fractions for each sample, using statistical tools recommended by CLSI.

The results obtained with the MINICAP HEMOGLOBIN(E) procedure indicate a very good reproducibility for quantitative analysis for each hemoglobin component. All electrophoregrams were also interpreted visually.

The results presented below have been obtained using the standard parameters of the PHORESIS software (smoothing 0 and hemoglobin fractions automatic quantification with HEMOGLOBIN(E) analysis program).

Reproducibility between capillaries from the same instrument

Seven (7) different blood samples were run using the MINICAP HEMOGLOBIN(E) procedure in both capillaries of the same MINICAP FLEX-PIERCING instrument and with 1 lot of MINICAP HEMOGLOBIN(E) kit. The analyzed blood samples included 2 samples with normal Hb A2 level (No. 1 and 5), 1 sample with low Hb A2 level (No. 2), 2 samples with increased Hb A2 level (No. 3 and 6), 1 pathological sample with Hb F and Hb S (No. 4) and 1 pathological sample with Hb F, Hb S and Hb C (No. 7). In this study, each blood sample was analyzed on both capillaries from the same instrument, including 40 runs over 20 working days (at 2 different times of the day). Within each run, samples were analyzed in duplicate. The results for Hb A, Hb A2, Hb F, Hb S and Hb C percentages are summarized in the following tables.

	Sample No. 1	Sample No. 2	Sample No. 3	Sample No. 4	Sample No. 5	Sample No. 6	Sample No. 7
Mean Hb A %	97.6	98.3	95.4	46.5	97.2	93.3	45.7
Within-run reproducibility (CV %)	0.0	0.1	0.1	0.2	0.0	0.1	0.5
Between-run reproducibility (CV %)	0.0	0.0	0.0	0.1	0.0	0.0	0.1
Between-day reproducibility (CV %)	0.0	0.0	0.0	0.1	0.0	0.0	0.1
Total (CV %)	0.0	0.1	0.1	0.3	0.1	0.1	0.5

	Sample No. 1	Sample No. 2	Sample No. 3	Sample No. 4	Sample No. 5	Sample No. 6	Sample No. 7
Mean Hb A2 %	2.4	1.8	4.6	2.8	2.8	6.7	2.8
Within-run reproducibility (CV %)	1.8	3.1	1.1	1.8	1.7	1.0	4.0
Between-run reproducibility (CV %)	0.0	0.0	0.0	0.0	0.7	0.2	0.0
Between-day reproducibility (CV %)	0.0	0.7	0.5	0.3	0.0	0.0	1.6
Total (CV %)	1.8	3.1	1.2	1.8	1.8	1.1	4.3

	Sample No. 1	Sample No. 2	Sample No. 3	Sample No. 4	Sample No. 5	Sample No. 6	Sample No. 7
Mean Hb F %	/	/	/	13.0	/	/	25.6
Within-run reproducibility (CV %)	/	/	/	0.8	/	/	0.4
Between-run reproducibility (CV %)	/	/	/	0.0	/	/	0.0
Between-day reproducibility (CV %)	/	/	/	0.3	/	/	0.1
Total (CV %)	/	/	/	0.9	/	/	0.4

	Sample No. 1	Sample No. 2	Sample No. 3	Sample No. 4	Sample No. 5	Sample No. 6	Sample No. 7
Mean Hb S %	/	/	/	37.7	/	/	18.9
Within-run reproducibility (CV %)	/	/	/	0.4	/	/	0.7
Between-run reproducibility (CV %)	/	/	/	0.0	/	/	0.0
Between-day reproducibility (CV %)	/	/	/	0.2	/	/	0.2
Total (CV %)	/	/	/	0.5	/	/	0.8

	Sample No. 1	Sample No. 2	Sample No. 3	Sample No. 4	Sample No. 5	Sample No. 6	Sample No. 7
Mean Hb C %	/	/	/	/	/	/	7.1
Within-run reproducibility (CV %)	/	/	/	/	/	/	1.5
Between-run reproducibility (CV %)	/	/	/	/	/	/	0.0
Between-day reproducibility (CV %)	/	/	/	/	/	/	0.5
Total (CV %)	/	/	/	/	/	/	1.6

In addition, none of the repeats showed false positive or false negative values :

- Samples with normal Hb A2 level : all values are normal ;
- Samples with decreased Hb A2 level : all values are decreased ;
- Samples with elevated Hb A2 level : all values are elevated ;
- Samples with hemoglobin variants : all hemoglobin variants are detected.

Linearity

The linearity of the MINICAP HEMOGLOBIN(E) procedure performed with the MINICAP FLEX-PIERCING instrument was evaluated in a study based on the Clinical Laboratory Standards Institute (CLSI - USA) EP6-A guideline "Evaluation of the Linearity of Quantitative Measurement Procedures: A statistical Approach".

The results for percentages (%) of hemoglobin fractions were analyzed using statistical tools recommended by CLSI.

Hb A2 linearity

One Hb A2 enriched blood sample (containing 13.4 g/dL total hemoglobin with 12.2 % of Hb A2) was mixed with a Hb A2 depleted blood sample (containing 12.8 g/dL total hemoglobin with 0.0 % of Hb A2) within different proportions and the dilutions were electrophoresed with MINICAP HEMOGLOBIN(E) procedure performed with MINICAP FLEX-PIERCING instrument.

The MINICAP HEMOGLOBIN(E) procedure performed with MINICAP FLEX-PIERCING instrument gave a good linearity for Hb A and Hb A2 within the entire range studied, with a maximum of about 1.6 g/dL for Hb A2 (between 0.0 and 12.2 % of Hb A2).

In addition, a blood sample (containing 17.9 g/dL total hemoglobin with 97.6 % of Hb A and 2.4 % of Hb A2) was concentrated and serially diluted in MINICAP HEMOGLOBIN(E) hemolysing solution and electrophoresed with the MINICAP HEMOGLOBIN(E) procedure performed with MINICAP FLEX-PIERCING instrument.

The tests were determined to be linear within the entire range studied from 1.79 to 45.90 g/dL total hemoglobin and Hb A and Hb A2 fractions percentages were not affected by the total hemoglobin concentration of the sample.

Hb F linearity

One umbilical cord blood sample (containing 15.8 g/dL total hemoglobin with 82.2 % of Hb F) was mixed with a normal blood sample (containing 11.6 g/dL total hemoglobin with 0.0 % of Hb F) within different proportions and the dilutions were electrophoresed with MINICAP HEMOGLOBIN(E) procedure performed with MINICAP FLEX-PIERCING instrument.

The MINICAP HEMOGLOBIN(E) procedure performed with MINICAP FLEX-PIERCING instrument gave a good linearity for Hb A and Hb F within the entire range studied, with a maximum of about 13.0 g/dL for Hb F (between 0.0 and 82.2 % of Hb F).

In addition, one umbilical cord blood sample (containing 17.9 g/dL total hemoglobin with 14.7 % of Hb A and 85.3 % of Hb F) was concentrated and serially diluted in MINICAP HEMOGLOBIN(E) hemolysing solution and electrophoresed with the MINICAP HEMOGLOBIN(E) procedure performed with MINICAP FLEX-PIERCING instrument.

The tests were determined to be linear within the entire range studied from 1.79 to 28.87 g/dL total hemoglobin and Hb A and Hb F fractions percentages were not affected by the total hemoglobin concentration of the sample.

Hb S linearity

One pathological blood sample with Hb S (containing 9.1 g/dL total hemoglobin with 86.2 % of Hb S and 0.0 % of Hb A) was mixed with a normal blood sample (containing 13.5 g/dL total hemoglobin with 0.0 % of Hb S and 97.4 % of Hb A) within different proportions and the dilutions were electrophoresed with MINICAP HEMOGLOBIN(E) procedure performed with MINICAP FLEX-PIERCING instrument.

The MINICAP HEMOGLOBIN(E) procedure performed with MINICAP FLEX-PIERCING instrument gave a good linearity for Hb S within the entire range studied, with a maximum of about 7.8 g/dL for Hb S (between 0.0 and 86.2 % of Hb S) and a good linearity for Hb A within the entire range studied, with a maximum of about 13.1 g/dL for Hb A (between 0.0 and 97.4 % of Hb A).

In addition, a pathological blood sample (containing 9.2 g/dL total hemoglobin with 79.1 % of Hb S) was concentrated and serially diluted in MINICAP HEMOGLOBIN(E) hemolysing solution and electrophoresed with the MINICAP HEMOGLOBIN(E) procedure performed with MINICAP FLEX-PIERCING instrument.

The tests were determined to be linear within the entire range studied from 0.92 to 31.72 g/dL total hemoglobin and Hb S fraction percentage was not affected by the total hemoglobin concentration of the sample.

Accuracy – Internal correlation

The internal concordance study of the MINICAP HEMOGLOBIN(E) procedure performed with the MINICAP FLEX-PIERCING instrument was evaluated in a study based on the Clinical Laboratory Standards Institute (CLSI - USA) EP09-A2 guideline "Method Comparison and Bias Estimation Using Patient Samples; Approved Guideline – Second Edition (Interim Revision)".

The results for percentages (%) of hemoglobin fractions were analyzed using statistical tools recommended by CLSI.

NOTE : The results presented below have been obtained from 1 internal accuracy study that has been performed in SEBIA facility. The analyzed blood samples were provided by 17 hospital laboratories in France, USA and Asia.

All the samples were exactly treated the same way with both techniques and followed the same guidelines in regards to sample integrity.

The levels of hemoglobin fractions were measured in 116 blood samples, including 54 samples with hemoglobin variants such as hemoglobins S, C, D and E, both by electrophoretic separations obtained with the MINICAP HEMOGLOBIN(E) procedure with the MINICAP FLEX-PIERCING instrument and a commercially available capillary electrophoresis technique for hemoglobins analysis.

The measured values of hemoglobin fractions from both procedures were analyzed by a linear regression statistical procedure. The results of linear regression analysis for Hb A, Hb A2, Hb F, Hb S, Hb C and Hb E are tabulated below (y = MINICAP HEMOGLOBIN(E) procedure with MINICAP FLEX-PIERCING instrument) :

Hb fraction	Number of samples	Correlation coefficient	y-Intercept	Slope	Range of Hb % values MINICAP HEMOGLOBIN(E) with MINICAP FLEX-PIERCING instrument
Hb A	110	1.000	1.175	0.990	12.1 – 98.6
Hb A2	115	0.997	- 0.030	0.980	0.2 – 6.6
Hb F	43	1.000	- 0.013	1.000	0.3 – 80.0
Hb S	27	1.000	- 0.486	0.996	7.8 – 90.8
Hb C	6	1.000	- 0.204	0.979	6.9 – 36.6
Hb E	6	0.991	- 0.408	0.981	21.7 – 25.7

This study demonstrated a perfect correlation between the 2 analysis procedures for the hemoglobin A2 quantitative determination in samples without any hemoglobin variant, with a 100.0 % sensibility and a 92.5 % specificity of MINICAP HEMOGLOBIN(E) procedure compared to the reference procedure, calculated using the recommended method (Wendling, 1986).

Additionally, this study demonstrated a perfect correlation between the 2 analysis procedures for the detection of abnormal hemoglobins, with a 100.0 % sensibility and a 100.0 % specificity of MINICAP HEMOGLOBIN(E) procedure compared to the reference procedure, calculated using the recommended method (Wendling, 1986). All abnormal hemoglobins (S, C, D, E and other variants) or abnormal levels of normal hemoglobins detected with the MINICAP HEMOGLOBIN(E) procedure performed with the MINICAP FLEX-PIERCING instrument were in agreement with the comparative capillary electrophoresis technique. There was no case observed of false positive, i.e., detection of an abnormal band or abnormal level of a normal band where no such abnormality existed.

BIBLIOGRAPHIE / BIBLIOGRAPHY

BIBLIOGRAFIE - BIBLIOGRAFIA - BIBLIOGRAFÍA - BIBLIOGRAFI - ΒΙΒΛΙΟΓΡΑΦΙΑ - BIBLIOGRAFIJU - BIBLIOGRAFIJA - КАРНАҚА - БИБЛІОГРАФІЯ - 参考书目 - БИБЛІОГРАФІЮ - 参考文献 - IZMANTOTĀ LITERĀTŪRA - BIBLIOGRAFIU - KIRJANDUS - DANH MỤC TÀI LIỆU THAM KHẢO

1. J. Bardakdjian-Michau, J.-L. Dhondt, R. Ducrocq, F. Galactéros, A. Guyard, F.-X. Huchet, A. Lahary, D. Lena-Russo, P. Maboudou, M.-L. North, C. Prelu, A.-M. Soummer, M. Verscheide, H. Wajzman (2003) Bonnes pratiques de l'étude de l'hémoglobine. *Ann. Biol. Clin.*, 61, 401-409.
2. V.F. Fairbanks, ed. (1980) Hemoglobinopathies and thalassemia: Laboratory methods and case studies. Brian C. Decker, New York.
3. F. Galacteros (1986) Thalassémie, drépanocytose et autres hémoglobinopathies. *Techniques et Biologie*, 3, 174-178.
4. JM Hempe, JN Granger and RD Craver (1997) Capillary isoelectric focusing of hemoglobin variants. *Electrophoresis*, 18, 1785-1795.
5. T.H.J. Huisman and J.H.P. Jonxis (1977) The hemoglobinopathies: techniques of identification. Marcel Dekker, New York.
6. Jellum E *et al.* Diagnostic applications of chromatography and capillary electrophoresis. *J. Chromatogr. B*, 689, 155-164 (1997).
7. Joutovsky A, Hadzi-Nesic J and Nardi MA (2004) HPLC retention time as a diagnostic tool for hemoglobin variants and hemoglobinopathies : a study of 60 000 samples in a clinical diagnostic laboratories. *Clin. Chem.*, 50, 10, 1736-1747.
8. J.S. Krauss, P.A. Drew, M.H. Jonah, M. Trinh, S. Shell, L. Black and C.R. Baisden (1986) Densitometry and microchromatography compared for determination of the hemoglobin C and A2 proportions in hemoglobin C and hemoglobin SC disease and in hemoglobin C trait. *Clin. Chem.* 32, 5, 860-863.
9. Landers JP. Clinical Capillary Electrophoresis. *Clin. Chem.*, 41, 495-509 (1995).
10. C. Livingstone (1986) The hemoglobinopathies. Edit. London.-
11. M. Maier-Redelsberger, R. Girot (1989) Diagnostic biologique des maladies de l'hémoglobine. *Feuilles de biologie*, 170.
12. Oda RP *et al.* Capillary electrophoresis as a clinical tool for the analysis of protein in serum and other body fluids. *Electrophoresis*, 18, 1715-1723 (1997).
13. R.G. Schneider (1978) Methods for detection of hemoglobin variants and hemoglobinopathies in the routine clinical laboratory. *CRC Crit. Rev. Clin. Lab. Sci.* 9, 243-271.
14. L. Vovan, D. Lara-Russo, A. Orsini (1985) Diagnostic biologique des hémoglobinoses. *Ann. Pédiat.* 32, 9, 780-789.
15. <http://globin.cse.psu.edu/hbvar/menu.html> : Hbvar : A Database of Human Hemoglobin Variants and Thalassemias.
16. <http://www.isns-neoscreening.org/agenda.htm>.
17. F Boemer, O Ketelslegers, JM Minon, V Bours, R Schoops (2008) Newborn screening for sickle cell disease using tandem mass spectrometry. *Clin. Chem.*, 54, 12, 2036-2041.
18. Lubin BH, Witkowska HE, Klemm K (1991) Laboratory diagnosis of hemoglobinopathies. *Clin. Biochem.*, 24, 363-374.
19. B Gulbis, B Fontaine, F Vertongen, F Cotton (2003) The place of capillary electrophoresis techniques in screening for haemoglobinopathies. *Ann. Clin. Biochem.*, 40, 659-662.
20. Aguilar-Martínez P *et al* (2010) Arbres décisionnels pour le diagnostic et la caractérisation moléculaire des hémoglobinopathies. *Ann. Biol. Clin.*, 68 (4) : 455-464.
21. Wendling A. Procédures de diagnostic ou de dépistage : Justification et validité d'un test de diagnostic ou de dépistage-sensibilité-spécificité. *Impact-Internat*, 1986 ; Sept : 93-97.
22. L. Guis, A. Chaumier, V. Le Gall, S. Havrez (Février 2013) Intégration du Capillarys 2 Flex Piercing (Sebia) dans un laboratoire de biologie médicale spécialisée. *Revue Francophone des Laboratoires*, 449, 47 – 56.
23. I. Agouti, F. Merono, N. Bonello-Palot, C. Badens (2013) Analytical evaluation of the Capillarys 2 Flex piercing for routine haemoglobinopathies diagnosis. *Int. Jnl. Lab. Hem.* 35, 217 – 221.
24. S. Altinier, M. Varagnolo, M. Zaninotto, M. Plebani. (2012) Identification and quantification of hemoglobins in whole blood: the analytical and organizational aspects of Capillarys 2 Flex Piercing compared with agarose electrophoresis and HPLC methods. *Clin. Chem. Lab. Med.* DOI 10.1515/ccml-2012-0061.
25. M. Angastiniotis, J.L. Vives Corrons, E.S. Soteriades, A. Eleftheriou (2013) The Impact of Migrations on the Health Services for Rare Diseases in Europe: The Example of Haemoglobin Disorders. *The Scientific World Journal* Volume 2013, Article ID 727905, 10 pages.
26. Nicole Borbely, Lorraine Phelan, Richard Szydio, *et al.* (2012) Capillary zone electrophoresis for haemoglobinopathy diagnosis. *J. Clin. Pathol.*, doi: 10.1136/jclinpath-2012-200946.
27. F. Cotton *et al.* (2009) Evaluation of an automated capillary electrophoresis system in the screening for hemoglobinopathies. *Clin. Lab.*, 55 : 217 – 221.
28. D. Greene, A.L. Pyle, J.S. Chang, C. Hoke, T. Lorey (2012) Comparison of Sebia Capillarys Flex capillary electrophoresis with the BioRad Variant II high pressure liquid chromatography in the evaluation of hemoglobinopathies. *Clin. Chim. Acta* 413, 1232 – 1238.
29. T. Higgins, M. Mack, A. Khajuria (2009) Comparison of two methods for the quantification and identification of hemoglobin variants. *Clin. Biochem.*, 42, 701 – 705.
30. D.F. Keren *et al* (2008) Comparison of Sebia Capillarys capillary electrophoresis with the Primus High-Pressure Liquid Chromatography in the evaluation of hemoglobinopathies. *Am. J. Clin. Pathol.*, 130 : 824 – 831.
31. D.F. Keren *et al* (2012) Expression of hemoglobin variant migration by capillary electrophoresis relative to Hemoglobin A2 improves precision. *Am. J. Clin. Pathol.*, 137 : 660 – 664.
32. Can Liao, Jian-Ying Zhou, Xing-Mei Xie, Jian Li, Ru Li, and Dong-Zhi Li (2010). Detection of Hb constant spring by a capillary electrophoresis method. *Hemoglobin*, 34 (2) : 175 – 178.
33. D.M. Mais *et al* (2009) The range of hemoglobin A2 in hemoglobin E heterozygotes as determined by capillary electrophoresis. *Am. J. Clin. Pathol.*, 132 : 34 – 38.
34. T. Munkongdee *et al* (2010) Quantitative analysis of Hb Bart's in cord blood by capillary electrophoresis system. *Ann. Hematol.* DOI 10.1007/s00277-010-1137-4.
35. R. Paleari, B. Gulbis, F. Cotton, A. Mosca (2012) Interlaboratory comparison of current high-performance methods for HbA2. *Int. Jnl. Lab. Hem.* 34, 362 – 368.
36. S. Sangkitporn *et al* (2011) Multicenter validation of fully automated capillary electrophoresis method for diagnosis of thalassemias and hemoglobinopathies in Thailand. *Southeast Asian J. Trop. Med. Public Health* 42 (5), 1224 – 1232.
37. F. Wolff, F. Cotton, B. Gulbis (2012) Screening for haemoglobinopathies on cord blood : laboratory and clinical experience. *J. Med. Screen.*, 19, 3, 116 - 122.

BIBLIOGRAPHIE / BIBLIOGRAPHY

BIBLIOGRAFIE - BIBLIOGRAFIA - BIBLIOGRAFÍA - BIBLIOGRAFÍ - ΒΙΒΛΙΟΓΡΑΦΙΑ - BIBLIOGRAFIJU - BIBLIOGRAFIJA - КАРНАҚСА - БИБЛИОГРАФИЯ - 参考书目 - БИБЛИОГРАФИЮ - 参考文献 - IZMANTOTĀ LĪTERĀTŪRA - BIBLIOGRAFIU - KIRJANDUS - DANH MỤC TÀI LIỆU THAM KHẢO

38. A.D. Stephens *et al* on behalf of the International Council for the Standardisation of Haematology (ICSH) (2012). ICSH recommendations for the measurement of Haemoglobin F. *Int. Jnl. Lab. Hem.*, 34, 14 – 20.
39. A. Mosca, R. Paleari, D. Leone, G. Ivaldi (2009) The relevance of hemoglobin F measurement in the diagnosis of thalassemias and related hemoglobinopathies. *Clinical Biochemistry*, 42, 1797–1801.
40. J. D. Hoyer, C. S. Penz, V. F. Fairbanks, C. A. Hanson and J.A. Katzmam (2002) Diagnostic Usefulness in the Distinction of Hereditary Persistence of Fetal Hemoglobin (HPFH) and Hemoglobin S–HPFH From Other Conditions With Elevated Levels of Hemoglobin F. *Am. J. Clin. Pathol.*, 117 : 857 – 863.
41. P. Van Delft *et al* (2009) Evaluating five dedicated automatic devices for haemoglobinopathy diagnostics in multi-ethnic populations. *Int. J. Lab. Hematol.*, 31 (5) : 484 – 95.
42. Prevention and diagnosis of haemoglobinopathies. A short guide for health professionals and laboratory scientists. Thalassaemia International Federation (TIF) publication No. 21.
43. A.J. Marengo-Rowe (2007) The thalassemias and related disorders. *Baylor University Medical Center Proceedings*, 20, 27 - 31.
44. Sae-ung *et al* (2012). Phenotypic expression of hemoglobins A2, E and F in various hemoglobin E related disorders. *Blood Cells, Molecules, and Diseases*, 48, 11–16.
45. A. Cao and R. Galanello (2010) Beta-thalassemia. *Genetics in Medicine*, 12 (2) : 61 – 76.
46. R.Z. Azma *et al* (2012) Co-inheritance of compound heterozygous Hb Constant Spring and a single – α 3.7 gene deletion with heterozygous $\delta\beta$ thalassaemia: A diagnostic challenge. *Malaysian J. Pathol.* ; 34(1) : 57 – 62.
47. C.L. Harteveld and D.R. Higgs (2010) α -thalassaemia. *Orphanet Journal of Rare Diseases*, 5, 13.

TABLEAU / TABLE

VARIANTS POTENTIELS PRÉSENTS DANS CHAQUE ZONE - POTENTIAL VARIANTS LOCATED IN EACH ZONE

Zone	Hémoglobines / Hemoglobins (Hb)
Z1	<p>Hb Santa Ana (pic mineur) #, Hb Mizuho (pic mineur) #, Hb delta A'2, Hb A2-Canebière, Hb A2-Lampang, Hb S-Oman, Hb A2-Turkish, Hb T-Cambodia, Hb Poissy (pic mineur) #, variant de Hb A2 "Chad" #, variant de Hb A2 "Savaria" #, variant de Hb A2 "Arya" #, variant de Hb A2 "Hasharon" #, variant de Hb A2 "Fort de France" #, variant de Hb A2 "Ottawa" #, variant de Hb A2 "Shimonoseki" #, variant de Hb A2 "Russ" (alpha 2) #, variant de Hb A2 "Russ" (alpha 1) #, variant de Hb A2 "Matsue-OkI" #, variant de Hb A2 "Reims" #, variant de Hb A2 "Mizushi" #, variant de Hb A2 "Stanleyville-II" #, variant de Hb A2 "O-Indonesia" #, variant de Hb A2 "San Antonio" #, variant de Hb A2 "G-Audhali" #, variant de Hb A2 "Handsworth" #, variant de Hb A2 "G-Philadelphia" #, variant de Hb A2 "Q-India" #, variant de Hb A2 "Memphis" #, variant de Hb A2 "Q-Iran" #, variant de Hb A2 "G-Waimanalo" #, variant de Hb A2 "Watts" #, variant de Hb A2 "Spanish Town" #, variant de Hb A2 "Montgomery" #, variant de Hb A2 "G-Norfolk" #, variant de Hb A2 "Inkster" #, variant de Hb A2 "Ube-4" #, variant de Hb A2 "G-Pest" #, variant de Hb A2 "Winnipeg" #, variant de Hb A2 "Queens" #, variant de Hb A2 "Etobicoke" #, variant de Hb A2 "Chapel Hill" #, variant de Hb A2 "Park Ridge" #, variant de Hb A2 "Q-Thailand" #, variant de Hb A2 "Delfzicht" # !!</p> <p><i>Hb Santa Ana (minor peak) #, Hb Mizuho (minor peak) #, Hb delta A'2, Hb A2-Canebière, Hb A2-Lampang, Hb S-Oman, Hb A2-Turkish, Hb T-Cambodia, Hb Poissy (minor peak) #, "Chad" Hb A2 variant #, "Savaria" Hb A2 variant #, "Arya" Hb A2 variant #, "Hasharon" Hb A2 variant #, "Fort de France" Hb A2 variant #, "Ottawa" Hb A2 variant #, "Shimonoseki" Hb A2 variant #, "Russ" Hb A2 variant (alpha 2) #, "Russ" Hb A2 variant (alpha 1) #, "Matsue-OkI" Hb A2 variant #, "Reims" Hb A2 variant #, "Mizushi" Hb A2 variant #, "Stanleyville-II" Hb A2 variant #, "O-Indonesia" Hb A2 variant #, "San Antonio" Hb A2 variant #, "G-Audhali" Hb A2 variant #, "Handsworth" Hb A2 variant #, "G-Philadelphia" Hb A2 variant #, "Q-India" Hb A2 variant #, "Memphis" Hb A2 variant #, "G-Waimanalo" Hb A2 variant #, "Watts" Hb A2 variant #, "Spanish Town" Hb A2 variant #, "Montgomery" Hb A2 variant #, "G-Norfolk" Hb A2 variant #, "Inkster" Hb A2 variant #, "Ube-4" Hb A2 variant #, "G-Pest" Hb A2 variant #, "Winnipeg" Hb A2 variant #, "Queens" Hb A2 variant #, "Etobicoke" Hb A2 variant #, "Chapel Hill" Hb A2 variant #, "Park Ridge" Hb A2 variant #, "Q-Thailand" Hb A2 variant #, "Delfzicht" Hb A2 variant # !!</i></p>
Z(C)	<p>Hb C-Ziguinchor !!, Hb F-Hull, Hb F-Texas-I, Hb Constant Spring, Hb Paksé, Hb C, Hb C-Harlem (C-Georgetown), variant de Hb A2 "Les Lilas" #, variant de Hb A2 "Boumerdes" #, variant de Hb A2 "Tarrant" #, variant de Hb A2 "Dunn" #, variant de Hb A2 "Bassett" #, variant de Hb A2 "Sassari" #, variant de Hb A2 "St. Luke's" #, variant de Hb A2 "Verdun" #, variant de Hb A2 "Manitoba-I" #, variant de Hb A2 "Setif" #, variant de Hb A2 "Sunshine Seth" #, variant de Hb A2 "Titusville" #, variant de Hb A2 "Swan River" #, variant de Hb A2 "Manitoba-II" #, variant de Hb A2 "Val de Marne" #</p> <p><i>Hb C-Ziguinchor !!, Hb F-Hull, Hb F-Texas-I, Hb Constant Spring, Hb Paksé, Hb C, Hb C-Harlem (C-Georgetown), "Les Lilas" Hb A2 variant #, "Boumerdes" Hb A2 variant #, "Tarrant" Hb A2 variant #, "Dunn" Hb A2 variant #, "Bassett" Hb A2 variant #, "Sassari" Hb A2 variant #, "St. Luke's" Hb A2 variant #, "Verdun" Hb A2 variant #, "Manitoba-I" Hb A2 variant #, "Setif" Hb A2 variant #, "Sunshine Seth" Hb A2 variant #, "Titusville" Hb A2 variant #, "Swan River" Hb A2 variant #, "Manitoba-II" Hb A2 variant #, "Val de Marne" Hb A2 variant #</i></p>
Z(A2)	<p>Hb A2, Hb Chad (E-Keelung) #, Hb A2-Madrid * #, Hb A2-Saint Denis * #, Hb A2-Saint-Etienne * #, Hb Hong Kong (cas anti-Lepore), Hb O-Tibesti, Hb Gun Hill, Hb O-Arab, Hb E-Saskatoon, Hb Shuangfeng, variant de Hb A2 "Charolles" #, variant de Hb A2 "Roubaix" #, variant de Hb A2 "El Escorial" * #, variant de Hb A2 "Dallas" * #, variant de Hb A2 "Barika" * #, variant de Hb A2 "Melusine" * #, variant de Hb A2 "Jura" #, variant de Hb A2 "Nouakchott" #, variant de Hb A2 "Pohnpei" #</p> <p><i>Hb A2, Hb Chad (E-Keelung) #, Hb A2-Madrid * #, Hb A2-Saint Denis * #, Hb A2-Saint-Etienne * #, Hb Hong Kong (anti-Lepore case), Hb O-Tibesti, Hb Gun Hill, Hb O-Arab, Hb E-Saskatoon, Hb Shuangfeng, "Charolles" Hb A2 variant #, "Roubaix" Hb A2 variant #, "El Escorial" Hb A2 variant * #, "Dallas" Hb A2 variant * #, "Barika" Hb A2 variant * #, "Melusine" Hb A2 variant * #, "Jura" Hb A2 variant #, "Nouakchott" Hb A2 variant #, "Pohnpei" Hb A2 variant #</i></p>
Z(E)	<p>Hb Hornchurch, Hb Seal Rock, Hb Köln (Ube-1) #, Hb Buenos Aires (pic mineur) #, Hb E, Hb Cleveland, Hb M-Saskatoon (pic mineur) #, Hb G-Siriraj, Hb A2-Babinga, Hb F-Moyen Orient, Hb O-Padova, Hb Agenogi, Hb Sabine #, Hb Santa Ana #, Hb Savaria # !!, Hb Djelfa (pic 3) # !!, variant de Hb A2 "M-Iwate" #, variant de Hb A2 "Saint Claude" #, variant de Hb A2 "Jackson" (alpha 2) #, Hb C dégradée</p> <p><i>Hb Hornchurch, Hb Seal Rock, Hb Köln (Ube-1) #, Hb Buenos Aires (minor peak) #, Hb E, Hb Cleveland, Hb M-Saskatoon (minor peak) #, Hb G-Siriraj, Hb A2-Babinga, Hb F-Moyen Orient, Hb O-Padova, Hb Agenogi, Hb Sabine #, Hb Santa Ana #, Hb Savaria # !!, Hb Djelfa (peak 3) # !!, "M-Iwate" Hb A2 variant #, "Saint Claude" Hb A2 variant #, "Jackson" Hb A2 variant (alpha 2) #, denatured Hb C</i></p>
Z(S)	<p>Hb Arya # !!, Hb Kenya (HPFH-7), Hb Hasharon (Sinai) #, Hb Dhofar (Yukuhashi), Hb Shimonoseki (Hikoshima) #, Hb O-Indonesia (Buginese-X) #, Hb Machida, Hb Vexin, Hb Corbeil, Hb Ottawa (Siam) #, Hb Fort de France #, Hb S, Hb G-Makassar, Hb Montgomery #, Hb G-Copenhague, Hb S-Antilles, Hb Handsworth #, Hb Lavagna, Hb Poissy #, Hb Hamadan, Hb Belfast, Hb Russ (alpha 1) #, Hb Russ (alpha 2) #, Hb Evanston, Hb Stanleyville-II # !!, Hb Cocody !!, Hb Reims # !!, variant de Hb A2 "Tokoname" #, variant de Hb A2 "Wayne" (pic 1) #, variant de Hb A2 "Pisa" #, variant de Hb A2 "J-Oxford" #, variant de Hb A2 "Lombard" #, variant de Hb A2 "Tatras" #, variant de Hb A2 "J-Cape Town" (alpha 2) #, variant de Hb A2 "Thionville" #, variant de Hb A2 "J-Cape Town" (alpha 1) #, variant de Hb A2 "Cemenelum" #, variant de Hb A2 "Nikaia" #, variant de Hb A2 "Hopkins-II" (alpha 1) #, variant de Hb A2 "Jackson" (alpha 1) #, variant de Hb A2 "Hopkins-II" (alpha 2) #, variant de Hb A2 "Singapore" # !!, Hb O-Arab dégradée</p> <p><i>Hb Arya # !!, Hb Kenya (HPFH-7), Hb Hasharon (Sinai) #, Hb Dhofar (Yukuhashi), Hb Shimonoseki (Hikoshima) #, Hb O-Indonesia (Buginese-X) #, Hb Machida, Hb Vexin, Hb Corbeil, Hb Ottawa (Siam) #, Hb Fort de France #, Hb S, Hb G-Makassar, Hb Montgomery #, Hb G-Copenhague, Hb S-Antilles, Hb Handsworth #, Hb Lavagna, Hb Poissy #, Hb Hamadan, Hb Belfast, Hb Russ (alpha 1) #, Hb Russ (alpha 2) #, Hb Evanston, Hb Stanleyville-II # !!, Hb Cocody !!, Hb Reims # !!, "Tokoname" Hb A2 variant #, "Wayne" Hb A2 variant (peak 1) #, "Pisa" Hb A2 variant #, "J-Oxford" Hb A2 variant #, "Lombard" Hb A2 variant #, "Tatras" Hb A2 variant #, "J-Cape Town" Hb A2 variant (alpha 2) #, "Thionville" Hb A2 variant #, "J-Cape Town" Hb A2 variant (alpha 1) #, "Cemenelum" Hb A2 variant #, "Nikaia" Hb A2 variant #, "Hopkins-II" Hb A2 variant (alpha 1) #, "Jackson" Hb A2 variant (alpha 1) #, "Hopkins-II" Hb A2 variant (alpha 2) #, "Singapore" Hb A2 variant # !!, denatured Hb O-Arab</i></p>

TABLEAU / TABLE

VARIANTS POTENTIELS PRÉSENTS DANS CHAQUE ZONE - POTENTIAL VARIANTS LOCATED IN EACH ZONE

Zone	Hémoglobines / Hemoglobins (Hb)
Z(D)	<p>Hb Memphis # !!, Hb G-Audhali # !!, Hb G-Szuhu (Gifu) !!, Hb Leiden !!, Hb Beograd (D-Camperdown), Hb Muravera, Hb D-Bushman, Hb Gavello, Hb Sogn, Hb Matsue-Oki #, Hb Osu Christiansborg, Hb D-Punjab (D-Los Angeles), Hb Watts #, Hb A2-Coburg, Hb G-Waimanalo (Aida) #, Hb Q-India #, Hb Muskegon, Hb Buenos Aires (pic mineur) #, Hb Lepore-BW, Hb Q-Iran #, Hb Akron, Hb Summer Hill, Hb G-Philadelphia #, Hb Karlskoga, Hb D-Ouled Rabah, Hb Aichi, Hb Oleander, Hb Yaizu, Hb Kenitra, Hb San Antonio #, Hb Aalborg, Hb Al-Hammadi Riyadh, Hb Ocho Rios, Hb Rocky Mountain (Paddington), Hb Lepore-Hollandia, Hb Quin-Hai, Hb Fort Worth, Hb Mizushi #, Hb G-Honolulu (G-Chinese), Hb Redondo (Isehara), Hb Lepore-Baltimore, Hb G-Ferrara, Hb Djelfa (pic 2) #, Hb G-Hsi-Tsou, Hb Hackney (Xu Chang), Hb Rothschild, Hb Spanish Town #, Hb Korle-Bu (G-Accra), Hb Khartoum, Hb Moabit, Hb Mobile, Hb Köln (Ube-1) #, Hb G-Norfolk #, Hb Ube-4 #, Hb Maputo, Hb Etobicoke #, Hb D-Iran, Hb Caribbean, Hb Okaloosa, Hb St. Luke's #, Hb G-Taipei, Hb G-Coushatta (G-Saskatoon), Hb Winnipeg #, Hb Canuts [A2], Hb Inkster #, Hb Zürich, Hb G-Pest #, Hb P-Galveston, Hb Queens (Ogi) #, Hb Canuts, Hb Aubenas, Hb Seif #, Hb P-Nilotic, Hb G-Galveston (G-Port Arthur), Hb Sunshine Seth # !!, Hb King's Mill !!, Hb Henri Mondor !!, Hb Titusville # !!, variant de Hb A2 "J-Sardegna" # !!, variant de Hb A2 "Suresnes" # !!, variant de Hb A2 "J-Meerut" (alpha 2) #, variant de Hb A2 "J-Broussais" (alpha 2) #, variant de Hb A2 "J-Rajappen" # !!, variant de Hb A2 "J-Anatolia" #, variant de Hb A2 "J-Meerut" (alpha 1) #, variant de Hb A2 "Ube-2" #, variant de Hb A2 "J-Broussais" (alpha 1) #, variant de Hb A2 "J-Abidjan" #, variant de Hb A2 "J-Toronto" (alpha 1) #, variant de Hb A2 "Mexico" (alpha 2) #, variant de Hb A2 "Thailand" #, variant de Hb A2 "Mexico" (alpha 1) #, variant de Hb A2 "J-Tongariki" #, variant de Hb A2 "Belliard" #, variant de Hb A2 "Neuilly-sur-Marne" #, variant de Hb A2 "J-Wenchang-Wuming" #, variant de Hb A2 "J-Paris-I" (alpha 2) #, variant de Hb A2 "J-Habana" #, variant de Hb A2 "J-Paris-I" (alpha 1) #, Hb E dégradée</p>
Z(F)	<p><i>Hb Memphis # !!, Hb G-Audhali # !!, Hb G-Szuhu (Gifu) !!, Hb Leiden !!, Hb Beograd (D-Camperdown), Hb Muravera, Hb D-Bushman, Hb Gavello, Hb Sogn, Hb Matsue-Oki #, Hb Osu Christiansborg, Hb D-Punjab (D-Los Angeles), Hb Watts #, Hb A2-Coburg, Hb G-Waimanalo (Aida) #, Hb Q-India #, Hb Muskegon, Hb D-Ibadan, Hb Buenos Aires (minor peak) #, Hb Lepore-BW, Hb Q-Iran #, Hb Akron, Hb Summer Hill, Hb G-Philadelphia #, Hb Karlskoga, Hb D-Ouled Rabah, Hb Aichi, Hb Oleander, Hb Yaizu, Hb Kenitra, Hb San Antonio #, Hb Aalborg, Hb Al-Hammadi Riyadh, Hb Ocho Rios, Hb Rocky Mountain (Paddington), Hb Lepore-Hollandia, Hb Quin-Hai, Hb Fort Worth, Hb Mizushi #, Hb G-Honolulu (G-Chinese), Hb Redondo (Isehara), Hb Lepore-Baltimore, Hb G-Ferrara, Hb Djelfa (peak 2) #, Hb G-Hsi-Tsou, Hb Hackney (Xu Chang), Hb Rothschild, Hb Spanish Town #, Hb Korle-Bu (G-Accra), Hb Khartoum, Hb Moabit, Hb Mobile, Hb Köln (Ube-1) #, Hb G-Norfolk #, Hb Ube-4 #, Hb Maputo, Hb Etobicoke #, Hb D-Iran, Hb Caribbean, Hb Okaloosa, Hb St. Luke's #, Hb G-Taipei, Hb G-Coushatta (G-Saskatoon), Hb Winnipeg #, Hb Canuts [A2], Hb Inkster #, Hb Zürich, Hb G-Pest #, Hb P-Galveston, Hb Queens (Ogi) #, Hb Canuts, Hb Aubenas, Hb Seif #, Hb P-Nilotic, Hb G-Galveston (G-Port Arthur), Hb Sunshine Seth # !!, Hb King's Mill !!, Hb Henri Mondor !!, Hb Titusville # !!, "J-Sardegna" Hb A2 variant # !!, "Suresnes" Hb A2 variant # !!, "J-Meerut" Hb A2 variant (alpha 2) #, "J-Broussais" Hb A2 variant (alpha 2) #, "J-Rajappen" Hb A2 variant #, "J-Anatolia" Hb A2 variant #, "J-Meerut" Hb A2 variant (alpha 1) #, "Ube-2" Hb A2 variant #, "J-Broussais" Hb A2 variant (alpha 1) #, "J-Abidjan" Hb A2 variant #, "J-Toronto" Hb A2 variant (alpha 1) #, "Mexico" Hb A2 variant (alpha 2) #, "Thailand" Hb A2 variant #, "Mexico" Hb A2 variant (alpha 1) #, "J-Tongariki" Hb A2 variant #, "Belliard" Hb A2 variant #, "Neuilly-sur-Marne" Hb A2 variant #, "J-Wenchang-Wuming" Hb A2 variant #, "J-Paris-I" Hb A2 variant (alpha 2) #, "J-Habana" Hb A2 variant #, "J-Paris-I" Hb A2 variant (alpha 1) #, denatured Hb E</i></p>
Z(F)	<p>Hb F, Hb Willamette !!, Hb Hoshida (Chaya) !!, Hb Languidic, Hb Chiapas, Hb P-India, Hb Tamano, Hb Sunnybrook, Hb Park Ridge #, Hb Delfzicht #, Hb Atago, Hb Deer Lodge, Hb Alabama, Hb Chapel Hill #, Hb Bunbury, Hb Tak, Hb Q-Thailand (G-Taichung), Hb Sabine #, Hb Bassett #, Hb Boyle Heights, Hb Les Lilas #, Hb Rampa, Hb Haaglanden, Hb G-Georgia, Hb Barcelona, Hb G-San José, Hb Denmark Hill, Hb Pôrto Alegre, Hb F-Sardinia, Hb Geldrop Santa Anna, Hb Ta-Li, Hb Chongqing, Hb Richmond, Hb Hirose, Hb Abruzzo, Hb Boumerdes #, Hb British Columbia, Hb Kansas, Hb Tarrant #, Hb Verdun #, Hb Swan River #, Hb Attleboro, Hb Sawara, Hb Burke, Hb Dunn #, Hb Manitoba-I #, Hb Manitoba-II #, Hb Sassari #, Hb Hazebrouck !!, Hb Port Phillip !!, Hb Vanderbilt !!, variant de Hb A2 "J-Rovigo" # !!, variant de Hb A2 "Wayne" (pic 2) # !!, Hb S dégradée, Hb D-Punjab dégradée</p> <p><i>Hb F, Hb Willamette !!, Hb Hoshida (Chaya) !!, Hb Languidic, Hb Chiapas, Hb P-India, Hb Tamano, Hb Sunnybrook, Hb Park Ridge #, Hb Delfzicht #, Hb Atago, Hb Deer Lodge, Hb Alabama, Hb Chapel Hill #, Hb Bunbury, Hb Tak, Hb Q-Thailand (G-Taichung), Hb Sabine #, Hb Bassett #, Hb Boyle Heights, Hb Les Lilas #, Hb Rampa, Hb Haaglanden, Hb G-Georgia, Hb Barcelona, Hb G-San José, Hb Denmark Hill, Hb Pôrto Alegre, Hb F-Sardinia, Hb Geldrop Santa Anna, Hb Ta-Li, Hb Chongqing, Hb Richmond, Hb Hirose, Hb Abruzzo, Hb Boumerdes #, Hb British Columbia, Hb Kansas, Hb Tarrant #, Hb Verdun #, Hb Swan River #, Hb Attleboro, Hb Sawara, Hb Burke, Hb Dunn #, Hb Manitoba-I #, Hb Manitoba-II #, Hb Sassari #, Hb Hazebrouck !!, Hb Port Phillip !!, Hb Vanderbilt !!, "J-Rovigo" Hb A2 variant # !!, "Wayne" Hb A2 variant (peak 2) # !!, denatured Hb S, denatured Hb D-Punjab</i></p>
Z8	<p>Hb F acétylée, Hb Grifton !!, Hb Lansing !!, Hb Hinsdale !!, Hb Ypsilanti (Ypsi - pic 1) # !!, Hb Auckland !!, Hb Roanne, Hb Southampton (Casper), Hb Yakima, Hb Saint Mandé, Hb Alberta, Hb Bruxelles, Hb Beth Israel, Hb Val de Marne (Footscray) #, Hb Kempsey, Hb Shelby (Leslie), Hb Atlanta, Hb Chemilly, Hb S-Clichy, Hb Sarrebourg, Hb Ypsilanti (Ypsi - pic 2) #, Hb Charolles #, Hb Athens-GA (Waco), Hb Debrousse, Hb Köln (Ube-1) #, Hb Aubagne, Hb Rainier</p> <p><i>Acetylated Hb F, Hb Grifton !!, Hb Lansing !!, Hb Hinsdale !!, Hb Ypsilanti (Ypsi - peak 1) # !!, Hb Auckland !!, Hb Roanne, Hb Southampton (Casper), Hb Yakima, Hb Saint Mandé, Hb Alberta, Hb Bruxelles, Hb Beth Israel, Hb Val de Marne (Footscray) #, Hb Kempsey, Hb Shelby (Leslie), Hb Atlanta, Hb Chemilly, Hb S-Clichy, Hb Sarrebourg, Hb Ypsilanti (Ypsi - peak 2) #, Hb Charolles #, Hb Athens-GA (Waco), Hb Debrousse, Hb Köln (Ube-1) #, Hb Aubagne, Hb Rainier</i></p>

TABLEAU / TABLE

VARIANTS POTENTIELS PRÉSENTS DANS CHAQUE ZONE - POTENTIAL VARIANTS LOCATED IN EACH ZONE

Zone	Hémoglobines / Hemoglobins (Hb)
	<p>Hb A, Hb Presbyterian *, Hb Roubaix (Poland) * #, Hb Silver Springs *, Hb El Escorial * #, Hb Dallas * #, Hb Phnom Penh *, Hb La Coruna *, Hb Bougardirey-Mail *, Hb Saint Nazaire *, Hb Barika * #, Hb Allentown *, Hb Allison Park *, Hb Alpertont *, Hb Aلدorf *, Hb Alzette *, Hb Anamosa *, Hb Antibes-Juan-Les-Pins *, Hb Arta (pic majeur) * #, Hb Aurillac *, Hb Austin *, Hb Aylesbury *, Hb Aztec *, Hb Bass Hill *, Hb Beirut *, Hb Belleville *, Hb Belluno *, Hb Bethesda *, Hb Bibba *, Hb Bladensburg *, Hb Boghé *, Hb Bonn *, Hb Brem-sur-Mer *, Hb Brest *, Hb Brigham *, Hb Brisbane (Great Lakes) *, Hb Broomhill *, Hb Brugg *, Hb Buenos Aires (Bryn Mawr - pic majeur) * #, Hb Buffalo (Reeuwijk) *, Hb Bushwick *, Hb Caen *, Hb Calvino *, Hb Cardarelli *, Hb Cheverly *, Hb Chicago *, Hb City of Hope *, Hb Coimbra (Ingelheim) *, Hb Columbia Missouri *, Hb Conakry *, Hb Cowtown *, Hb Crete *, Hb Dapu *, Hb Den Haag *, Hb Denver *, Hb Dhaka *, Hb Dhonburi (Neapolis) *, Hb Djelfa (pic 1) * #, Hb Ecuador *, Hb Evans *, Hb Flurlingen *, Hb Fontainebleau *, Hb Frankfurt *, Hb Fukuoka *, Hb Fukuyama *, Hb Geisinger *, Hb Genova (Hyogo) *, Hb Godavari *, Hb Gorwihl (Hinchingsbrooke) *, Hb Gouda *, Hb Grange Blanche *, Hb Groene Hart (Bernalda) *, Hb Grove City *, Hb Guanajuato *, Hb Haelen *, Hb Hamilton *, Hb Hammersmith (Chiba) *, Hb Heathrow *, Hb Hekinan *, Hb Hershey *, Hb Hyden *, Hb Inglewood *, Hb Iowa *, Hb Iraq-Halabja *, Hb Jabalpur *, Hb Jeddah *, Hb Johnstown *, Hb Kaiser West End *, Hb Kansas City *, Hb King Egbert *, Hb Knossos *, Hb Kokomo *, Hb Kosovo *, Hb La Desirade *, Hb Le Lamentin *, Hb Les Andelys *, Hb Linköping (Meilahti) *, Hb Lisbon *, Hb Little Rock *, Hb Louisville (Bucuresti) *, Hb Lulu Island *, Hb Lyon-Bron *, Hb M-Boston (M-Osaka) *, Hb M-Saskatoon (pic majeur) * #, Hb McKees Rocks *, Hb Malay *, Hb Malmö *, Hb Marijampolė *, Hb Marseille (Long Island) *, Hb Matsudo *, Hb Milledgeville *, Hb Minneapolis Laos *, Hb Mizuho * #, Hb Moriguchi *, Hb Mosella *, Hb Nakhon Ratchasima (Aberystwyth) *, Hb Nantes *, Hb Niguarda *, Hb Noko *, Hb Novara *, Hb Okayama *, Hb Olupona *, Hb Olympia *, Hb Owari *, Hb Ozieri *, Hb Parma *, Hb Part-Dieu *, Hb Perth (Abraham Lincoln) *, Hb Petit Bourg *, Hb Pierre-Bénite *, Hb Pittsburgh *, Hb Pohpei * #, Hb Port Huron *, Hb Potomac *, Hb Pressath *, Hb Princes Risborough *, Hb Puttelange *, Hb Raleigh *, Hb Ramona *, Hb Ravenscourt Park *, Hb Regina *, Hb Rhode Island (Southwark) *, Hb Riccarton *, Hb Rio Claro *, Hb Rotterdam *, Hb Rouen (Ethiopia) *, Hb Saclay *, Hb Saint-Clair *, Hb Saint-Jacques *, Hb St Joseph's *, Hb Saint-Marcellin *, Hb Saki *, Hb San Bruno *, Hb San Diego *, Hb San Martin *, Hb Santa Barnabas (Croxley Green) *, Hb Santa Juana (Serres) *, Hb Savannah *, Hb Saveh *, Hb Sendagi (Warsaw) *, Hb Sheffield *, Hb Sitia *, Hb Soderthalje *, Hb South Florida *, Hb South Milwaukee *, Hb South Yorkshire *, Hb Sydney *, Hb Taradale (Middlesbrough) *, Hb Taybe *, Hb Templeuve *, Hb Torino *, Hb Toulon *, Hb Twin Peaks *, Hb Ty Gard *, Hb Tyne *, Hb Utrecht *, Hb Uzes *, Hb Valletta *, Hb Valme *, Hb Venetia *, Hb Verona *, Hb Vientiane (Grey Lynn) *, Hb Vila Real *, Hb Villejuif *, Hb Villeparisis *, Hb Villeurbanne *, Hb Volga (Drenthe) *, Hb Voorhees *, Hb Washtenaw *, Hb Waterland *, Hb Weesp *, Hb Wembley *, Hb Westmead *, Hb Wiangpapao *, Hb William-Harvey *, Hb Wood *, Hb Worthing *, Hb Yaounde (Mataro) *, Hb Zoetermeer *, Hb Sinai-Baltimore *, Hb M-Milwaukee-I *, Hb Melusine * #, Hb Pitie-Salpetriere *, Hb Syracuse *, Hb Hounslow, Hb Fort Dodge, Hb Old Dominion (OD/BuT), Hb Camperdown, Hb Duarte !!, Hb Jura (Bamako) # !!</p>
Z(A)	<p>Hb A, Hb Presbyterian *, Hb Roubaix (Poland) * #, Hb Silver Springs *, Hb El Escorial * #, Hb Dallas * #, Hb Phnom Penh *, Hb La Coruna *, Hb Bougardirey-Mail *, Hb Saint Nazaire *, Hb Barika * #, Hb Allentown *, Hb Allison Park *, Hb Alpertont *, Hb Aلدorf *, Hb Alzette *, Hb Anamosa *, Hb Antibes-Juan-Les-Pins *, Hb Arta (main peak) * #, Hb Aurillac *, Hb Austin *, Hb Aylesbury *, Hb Aztec *, Hb Bass Hill *, Hb Beirut *, Hb Belleville *, Hb Belluno *, Hb Bethesda *, Hb Bibba *, Hb Bladensburg *, Hb Boghé *, Hb Bonn *, Hb Brem-sur-Mer *, Hb Brest *, Hb Brigham *, Hb Brisbane (Great Lakes) *, Hb Broomhill *, Hb Brugg *, Hb Buenos Aires (Bryn Mawr, major peak) * #, Hb Buffalo (Reeuwijk) *, Hb Bushwick *, Hb Caen *, Hb Calvino *, Hb Cardarelli *, Hb Cheverly *, Hb Chicago *, Hb City of Hope *, Hb Coimbra (Ingelheim) *, Hb Columbia Missouri *, Hb Conakry *, Hb Cowtown *, Hb Crete *, Hb Dapu *, Hb Den Haag *, Hb Denver *, Hb Dhaka *, Hb Dhonburi (Neapolis) *, Hb Djelfa (peak 1) * #, Hb Ecuador *, Hb Evans *, Hb Flurlingen *, Hb Fontainebleau *, Hb Frankfurt *, Hb Fukuoka *, Hb Fukuyama *, Hb Geisinger *, Hb Genova (Hyogo) *, Hb Godavari *, Hb Gorwihl (Hinchingsbrooke) *, Hb Gouda *, Hb Grange Blanche *, Hb Groene Hart (Bernalda) *, Hb Grove City *, Hb Guanajuato *, Hb Haelen *, Hb Hamilton *, Hb Hammersmith (Chiba) *, Hb Heathrow *, Hb Hekinan *, Hb Hershey *, Hb Hyden *, Hb Inglewood *, Hb Iowa *, Hb Iraq-Halabja *, Hb Jabalpur *, Hb Jeddah *, Hb Johnstown *, Hb Kaiser West End *, Hb Kansas City *, Hb King Egbert *, Hb Knossos *, Hb Kokomo *, Hb Kosovo *, Hb La Desirade *, Hb Le Lamentin *, Hb Les Andelys *, Hb Linköping (Meilahti) *, Hb Lisbon *, Hb Little Rock *, Hb Louisville (Bucuresti) *, Hb Lulu Island *, Hb Lyon-Bron *, Hb M-Boston (M-Osaka) *, Hb M-Saskatoon (main peak) * #, Hb McKees Rocks *, Hb Malay *, Hb Malmö *, Hb Marijampolė *, Hb Marseille (Long Island) *, Hb Matsudo *, Hb Milledgeville *, Hb Minneapolis Laos *, Hb Mizuho * #, Hb Moriguchi *, Hb Mosella *, Hb Nakhon Ratchasima (Aberystwyth) *, Hb Nantes *, Hb Niguarda *, Hb Noko *, Hb Novara *, Hb Okayama *, Hb Olupona *, Hb Olympia *, Hb Owari *, Hb Ozieri *, Hb Parma *, Hb Part-Dieu *, Hb Perth (Abraham Lincoln) *, Hb Petit Bourg *, Hb Pierre-Bénite *, Hb Pittsburgh *, Hb Pohpei * #, Hb Port Huron *, Hb Potomac *, Hb Pressath *, Hb Princes Risborough *, Hb Puttelange *, Hb Raleigh *, Hb Ramona *, Hb Ravenscourt Park *, Hb Regina *, Hb Rhode Island (Southwark) *, Hb Riccarton *, Hb Rio Claro *, Hb Rotterdam *, Hb Rouen (Ethiopia) *, Hb Saclay *, Hb Saint-Clair *, Hb Saint-Jacques *, Hb St Joseph's *, Hb Saint-Marcellin *, Hb Saki *, Hb San Bruno *, Hb San Diego *, Hb San Martin *, Hb Santa Barnabas (Croxley Green) *, Hb Santa Juana (Serres) *, Hb Savannah *, Hb Saveh *, Hb Sendagi (Warsaw) *, Hb Sheffield *, Hb Sitia *, Hb Soderthalje *, Hb South Florida *, Hb South Milwaukee *, Hb South Yorkshire *, Hb Sydney *, Hb Taradale (Middlesbrough) *, Hb Taybe *, Hb Templeuve *, Hb Torino *, Hb Toulon *, Hb Twin Peaks *, Hb Ty Gard *, Hb Tyne *, Hb Utrecht *, Hb Uzes *, Hb Valletta *, Hb Valme *, Hb Venetia *, Hb Verona *, Hb Vientiane (Grey Lynn) *, Hb Vila Real *, Hb Villejuif *, Hb Villeparisis *, Hb Villeurbanne *, Hb Volga (Drenthe) *, Hb Voorhees *, Hb Washtenaw *, Hb Waterland *, Hb Weesp *, Hb Wembley *, Hb Westmead *, Hb Wiangpapao *, Hb William-Harvey *, Hb Wood *, Hb Worthing *, Hb Yaounde (Mataro) *, Hb Zoetermeer *, Hb Sinai-Baltimore *, Hb M-Milwaukee-I *, Hb Melusine * #, Hb Pitie-Salpetriere *, Hb Syracuse *, Hb Hounslow, Hb Fort Dodge, Hb Old Dominion (OD/BuT), Hb Camperdown, Hb Duarte !!, Hb Jura (Bamako) # !!</p>

TABLEAU / TABLE

VARIANTS POTENTIELS PRÉSENTS DANS CHAQUE ZONE - POTENTIAL VARIANTS LOCATED IN EACH ZONE

Zone	Hémoglobines / Hemoglobins (Hb)
Z10	Hb Stockholm !!, Hb Créteil, Hb Nouakchott #, Hb M-Iwate (M-Kankakee) #, Hb Wayne (pic 1) #, Hb Complutense, Hb Camden (Tokuchi), Hb Hope
	<i>Hb Stockholm !!, Hb Créteil, Hb Nouakchott #, Hb M-Iwate (M-Kankakee) #, Hb Wayne (peak 1) #, Hb Complutense, Hb Camden (Tokuchi), Hb Hope</i>
Z11	Hb A dégradée, Hb Vaasa, Hb Tacoma, Hb Providence (pic X-Asn) #, Hb Yamagata, Hb Shepherds Bush, Hb Cook, Hb Corsica, Hb Pisa #, Hb K-Woolwich, Hb Lombard #, Hb J-Guantanamo, Hb Andrew Minneapolis, Hb J-Cape Town (alpha 1) #, Hb Kaohsiung (New York), Hb Fannin-Lubbock I, Hb Saint Claude #, Hb Thionville #, Hb Jackson (alpha 2) #, Hb J-Cape Town (alpha 2) #, Hb Strasbourg, Hb Osler (Fort Gordon), Hb Helsinki, Hb Doha, Hb Linwood, Hb J-Auckland, Hb Nancy, Hb Chesapeake, Hb Himeji, Hb Singapore #, Hb Jackson (alpha 1) #, Hb Cemenelum # !!, Hb Tatra # !!, variant de Hb A2 "I (I-Texas)" #
	<i>Denatured Hb A, Hb Vaasa, Hb Tacoma, Hb Providence (X-Asn peak) #, Hb Yamagata, Hb Shepherds Bush, Hb Cook, Hb Corsica, Hb Pisa #, Hb K-Woolwich, Hb Lombard #, Hb J-Guantanamo, Hb Andrew Minneapolis, Hb J-Cape Town (alpha 1) #, Hb Kaohsiung (New York), Hb Fannin-Lubbock I, Hb Saint Claude #, Hb Thionville #, Hb Jackson (alpha 2) #, Hb J-Cape Town (alpha 2) #, Hb Strasbourg, Hb Osler (Fort Gordon), Hb Helsinki, Hb Doha, Hb Linwood, Hb J-Auckland, Hb Nancy, Hb Chesapeake, Hb Himeji, Hb Singapore #, Hb Jackson (alpha 1) #, Hb Cemenelum # !!, Hb Tatra # !!, "I (I-Texas)" Hb A2 variant #</i>
Z12	Hb Bart, Hb Nikaia # !!, Hb Tokoname # !!, Hb J-Cubujuqui, Hb Hopkins-II (alpha 1) #, Hb J-Calabria (J-Bari), Hb J-Camagüey, Hb J-Tongariki #, Hb Wayne (pic 2) #, Hb J-Meerut (J-Birmingham - alpha 1) #, Hb Hopkins-II (alpha 2) #, Hb Zaire, Hb J-Meerut (J-Birmingham - alpha 2) #, Hb Trollhättan, Hb Pyrgos (Mizunami), Hb Providence (pic X-Asp) #, Hb Suresnes #, Hb J-Broussais (Tagawa-I - alpha 2) #, Hb Grady (Dakar - alpha 2), Hb Grady (Dakar - alpha 1), Hb Legnano, Hb Hikari, Hb J-Rajappen #, Hb J-Anatolia #, Hb J-Broussais (Tagawa-I - alpha 1) #, Hb J-Chicago, Hb J-Sardagna #, Hb J-Toronto (alpha 1) #, Hb J-Cordoba, Hb J-Meinung (J-Bangkok), Hb Ube-2 #, Hb Dagestan, Hb J-Cambridge (Rambam), Hb Hofu, Hb J-Abidjan #, Hb Ulm, Hb Belliard #, Hb J-Iran, Hb Riyadh (Karatsu), Hb Mexico (J-Paris-II - alpha 1) #, Hb Mexico (J-Paris-II - alpha 2) #, Hb Neuilly-sur-Marne #, Hb Pontoise (J-Pontoise), Hb Ankara, Hb J-Buda, Hb J-Medellin, Hb J-Paris-I (J-Aljezur - alpha 1) #, Hb Thailand #, Hb J-Habana #, Hb J-Baltimore (N-New Haven), Hb J-Wenchang-Wuming (Anantharaj) #, Hb J-Paris-I (J-Aljezur - alpha 2) #, Hb Beijing, Hb J-Oxford (I-Interlaken) #, Hb K-Ibadan !!
	<i>Hb Bart, Hb Nikaia # !!, Hb Tokoname # !!, Hb J-Cubujuqui, Hb Hopkins-II (alpha 1) #, Hb J-Calabria (J-Bari), Hb J-Camagüey, Hb J-Tongariki #, Hb Wayne (peak 2) #, Hb J-Meerut (J-Birmingham - alpha 1) #, Hb Hopkins-II (alpha 2) #, Hb Zaire, Hb J-Meerut (J-Birmingham - alpha 2) #, Hb Trollhättan, Hb Pyrgos (Mizunami), Hb Providence (X-Asp peak) #, Hb Suresnes #, Hb J-Broussais (Tagawa-I - alpha 2) #, Hb Grady (Dakar - alpha 2), Hb Grady (Dakar - alpha 1), Hb Legnano, Hb Hikari, Hb J-Rajappen #, Hb J-Anatolia #, Hb J-Broussais (Tagawa-I - alpha 1) #, Hb J-Chicago, Hb J-Sardagna #, Hb J-Toronto (alpha 1) #, Hb J-Cordoba, Hb J-Meinung (J-Bangkok), Hb Ube-2 #, Hb Dagestan, Hb J-Cambridge (Rambam), Hb Hofu, Hb J-Abidjan #, Hb Ulm, Hb Belliard #, Hb J-Iran, Hb Riyadh (Karatsu), Hb Mexico (J-Paris-II - alpha 1) #, Hb Mexico (J-Paris-II - alpha 2) #, Hb Neuilly-sur-Marne #, Hb Pontoise (J-Pontoise), Hb Ankara, Hb J-Buda, Hb J-Medellin, Hb J-Paris-I (J-Aljezur - alpha 1) #, Hb Thailand #, Hb J-Habana #, Hb J-Baltimore (N-New Haven), Hb J-Wenchang-Wuming (Anantharaj) #, Hb J-Paris-I (J-Aljezur - alpha 2) #, Hb Beijing, Hb J-Oxford (I-Interlaken) #, Hb K-Ibadan !!</i>
Z13	Hb Al-Ain Abu Dhabi, Hb J-Europa, Hb N-Baltimore (Hopkins-I), Hb J-Rovigo #, Hb J-Lome, Hb Arta (pic mineur) #, Hb J-Norfolk (Kagoshima), Hb Nigeria, Hb J-Kaohsiung (J-Honolulu)
	<i>Hb Al-Ain Abu Dhabi, Hb J-Europa, Hb N-Baltimore (Hopkins-I), Hb J-Rovigo #, Hb J-Lome, Hb Arta (minor peak) #, Hb J-Norfolk (Kagoshima), Hb Nigeria, Hb J-Kaohsiung (J-Honolulu)</i>
Z14	Hb N-Seattle, Hb J-Tashkuergan
	<i>Hb N-Seattle, Hb J-Tashkuergan</i>
Z15	Hb H, Hb I-Toulouse !!, Hb Sudbury, Hb Kurosaki (alpha 1), Poly A (A->G); AATAAA->AATAAG of the alpha2 gene alpha-Thal-2, Hb Kurosaki (alpha 2), Hb F-Emirates, Hb N-Timone, Hb I (I-Texas, I-Philadelphia) #, Hb Shaare Zedek
	<i>Hb H, Hb I-Toulouse !!, Hb Sudbury, Hb Kurosaki (alpha 1), Poly A (A->G); AATAAA->AATAAG of the alpha2 gene alpha-Thal-2, Hb Kurosaki (alpha 2), Hb F-Emirates, Hb N-Timone, Hb I (I-Texas, I-Philadelphia) #, Hb Shaare Zedek</i>

* Pic non ou peu visible car co-migrant de la fraction normale
Hidden or partially hidden peak due to similar migration time to normal fraction

Variant avec plusieurs fractions affichées (variant de la chaîne alpha ou variant instable ...)
Variant with several fractions displayed (alpha-chain or unstable variant ...)

!! Pic en bordure de zone (risque de saut de zone)
Peak in zone boundary (risk of zone shift)

Rappel : dans chaque zone les variants Hb sont listés selon leur temps de migration de droite vers la gauche
Reminder: in each zone Hb variants are sorted according to their migration time from right to left

TABLEAU / TABLE

FR : VARIANTI POTENTIELS PRÉSENTS DANS CHAQUE ZONE	Zone	Hémoglobines (Hb)	* Pic peu ou pas visible car migrant conjointement avec la fraction normale
GB : POTENTIAL VARIANTS LOCATED IN EACH ZONE	Zone	Hemoglobins (Hb)	* Hidden or partially hidden peak due to similar migration time to normal fraction
DE : POTENZIELLE VARIANTEN IN DEN EINZELNEN ZONEN	Zone	Hämoglobine (Hb)	* Versteckter oder teilweise versteckter Spitzenwert infolge ähnlicher Migrationszeit wie bei der normalen Fraktion
NL : POTENTIELE VARIANTEN IN ELKE ZONE	Zone	Hemoglobinen (Hb)	* Verborgen of gedeeltelijk verborgen pik vanwege migratietijd die vergelijkbaar is met normale fractie
IT : VARIANTI POTENZIALI PRESENTI IN CIASCUNA ZONA	Zona	Emoglobine (Hb)	* Picco invisibile o scarsamente visibile poiché migrante unitamente alla frazione normale
ES : VARIANTES POTENCIALES PRESENTES EN CADA ZONA	Zona	Hemoglobinas (Hb)	* Pico oculto o parcialmente oculto debido a migración conjunta con la fracción normal
PT : VARIANTES POTENCIAIS LOCALIZADAS EM CADA ZONA	Zona	Hemoglobinas (Hb)	* Pico oculto ou parcialmente oculto devido a tempo de migração similar para fração normal
SV : POTENTIALIA VARIANTER BELÄGNA I VARJE ZON	Zon	Hemoglobiner (Hb)	* Dold eller delvis dold topp på grund av liknande migratinstid till normal fraktion
GR : ΔΥΝΗΤΙΚΕΣ ΠΑΡΑΛΛΑΓΕΣ ΠΟΥ ΕΝΤΟΠΙΖΟΝΤΑΙ ΣΕ ΚΑΘΕ ΖΩΝΗ	Ζώνη	Αιμοσφαιρίνες (Hb)	* Αποκρύφει ή μερικώς αποκρύφει λόγω παρόμοιου χρόνου ηλεκτροφόρησης σε φυσιολογική κλάση
HR : POTENCIJALNE VARIJANTE U LOČNIM ZONAMA U SVAKOJ ZONI	Zona	Hemoglobini (Hb)	* Vrh je potpuno ili djelomično skriven zbog vremena migracije sličnog onome za normalnu frakciju
IT : GALIMI VARIANTATI, ESANTYI KIEVNIJEZOJE ZONJOJE	Zona	Hemoglobinas (Hb)	* Паспета арба іs далес паспета кірвіс, нс міграціос кілкс панасу і normalis фракціос міграціос кілкс
PL : POTENCJALNE ODMIANY ZŁOKAZIKOWANE W KAŻDEJ STREFIE	Strefa	Hemoglobiny (Hb)	* Ukryty lub częściowo ukryty pik ze względu na podobny czas migracji względem frakcji prawidłowej
RO : VARIANTE POTENTIALE SITUATE ÎN FIECARE ZONĂ	Zonă	Hemoglobine (Hb)	* Vârf ascuns sau parțial ascuns datorită timpului de migrare asemănător al fracției normale
CS : POTENCIJALNE VARIANTY KOJE SE NALAZE U SVAKOJ ZONI	Zona	Hemoglobini (Hb)	* Sakriveni ili delimično sakriveni vrh zbog sličnog vremena migracije u normalnu frakciju
HU : LEHETSÉGES VARIÁNSOK AZ EGYES ZÓNÁKBAN	Zóna	Hemoglobinek (Hb)	* A normál frakciónéhoz hasonló migrációs idő miatt rejtett vagy részben rejtett csúcs
TR : HER BİR BÖLGEDE YER ALAN VARYANTLAR	Bölge	Hemoglobiner (Hb)	* Normal fraksiyona benzer migrasyon süresine benzerliği gibi veya kısmen gibi pik
CZ : POTENCIJALNÍ VARIANTY HEMOGLOBINU UMÍSTĚNĚ V KAŽDĚ ZÓNĚ	Zóna	Hemoglobiny (Hb)	* Skrytý nebo částečně skrytý pik v porovnaní s časem migrace normální frakce
BG : ВЪЗМОЖНИ ВАРИАНТИ, РАЗПОЛОЖЕНИ ВЪВ ВСЯКА ЗОНА	Зона	Хемоглобини (Hb)	* Скрип или частично скрип пик поради време на миграция, подобно на това на нормалната фракция
NO : POTENSIELLE VARIANTER PÅLIGGER I HVER SONE	Sone	Hemoglobiner (Hb)	* Skjult eller delvis skjult topp på grunn av lignende migreringstid til normal fraksjon
DK : POTENTIELLE VARIANTER I HVER ZONE	Zone	Hæmoglobiner (Hb)	* Skjult eller delvis skjult topp på grund af lignende migratinstid til normal fraktion
CS : 在每个区中潜在的变种	区	血红蛋白 (Hb)	* 由于电泳时间与标准组分相似，因此峰被隐藏或部分隐藏
RU : ВОЗМОЖНЫЕ ВАРИАНТЫ, РАСПОЛОЖЕННЫЕ В КАЖДОЙ ЗОНЕ	Зона	Гемоглобин (Hb)	* Скрытый или частично скрытый пик по причине схожего времени миграции в нормальную фракцию
JP : ゼーンに位置する潜在的な変異体	ゾーン	ヘモグロビン (Hb)	* 正常フラクションへの移行時間が同一であるため、非表示または部分的に非表示のピーク
LV : POTENCIJALIE VARIANTI KATRĀ ZONĀ	Zona	Hemoglobīns (Hb)	* Slēpta vai daļēji slēpta maksimālā vērtība, ko izraisa migrācijas laiks, kas ir līdzīgs ar normālu frakciju
SK : VARIANTY POTENCIJALNE, KTORÉ SA NACHÁZAJÚ V KAŽDEJ ZÓNĚ	Zóna	Hemoglobiny (Hb)	* Skrytá alebo čiastočne skrytá špička v dôsledku podobného času migrácie ako pri normálnej frakcii
EE : POTENTSIJALSED VARIANDID IGAS TSOONIS	Tsoon	Hemoglobiindid (Hb)	* Varjatud või osaliselt varjatud pik normaalise fraktsiooniga sarnase migratsioonaja tõttu
VN : BIẾN THÌỀN ẨM TÀI MỖI VÙNG	Vùng	Hemoglobin (Hb)	* Đỉnh ẩn hoặc ẩn một phần vì thời gian di chuyển sang mang thông thường tương tự

FR : # Variant avec plusieurs fractions affichées (variant de la chaîne alpha ou variant instable ...)	!! Pic en bordure de zone (risque de changement de zone)
GB : # Variant with several fractions displayed (alpha-chain or unstable variant ...)	!! Peak in zone boundary (risk of zone shift)
DE : # Variante mit mehreren angezeigten Fraktionen (Alpha-Kette oder instabile Variante ...)	!! Spitzenwert im Zonenrenzgebiet (Risiko einer Zonenverschiebung)
NL : # Variant met verschillende zichtbare fracties (alketten of onstabiele variant ...)	!! Piek in zonegrens (gevaar voor zonevernoering)
IT : # Variante con più frazioni visualizzate (variante della catena alfa o variante instabile, ecc.)	!! Picco sul confine di zona (rischio di variazione di zona)
ES : # Variante que presenta varias fracciones (variante de cadena alfa o variante inestable ...)	!! Pico en el límite de la zona (riesgo de cambio de zona)
PT : # Variante com várias frações apresentadas (cadeia alfa ou variante instável, etc.)	!! Pico no limite da zona (risco de deslocação de zona)
SV : # Variant med flera fraktioner som visas (alfa-kedja eller instabil variant ...)	!! Topp i zongräns (risk för zonförskjutning)
GR : # Παραλλαγή με εμφάνιση πολλών κλάσμάτων (άλλα αλυσίδα ή ασταθής παραλλαγή ...)	!! Κορυφή σε όριο ζώνης (κίνδυνος μετατόπισης ζώνης)
HR : # Varijanta s nekoliko prikazanih frakcija (alfa-lanac ili nestabilna varijanta ...)	!! Vrh u granici zone (opasnost od pomaka zone)
IT : # Varianti su kelomis rodonomis frakcijomis nr. (alfa grandis arba nestabilus variantas ...)	!! Viršūnė ant zonos ribos (zonos paslinkimo pavojus)
PL : # Varianty z kilkoma wyświetlonymi frakcjami (alfa łańcuch lub odmianna niestabilna ...)	!! Pik w granicy strefy (ryzyko przesunięcia strefy)
RO : # Varianta cu mai multe fractii afisate (lant alfa sau varianta instabila ...)	!! Vârf în limita zonei (risc de schimbare a zonei)
CS : # Varianta sa prikazanih niekoľko frakcijs (alfa lanca ili nestabilna varijanta ...)	!! Vrh u hranici zóny (rizik od pomaka zone)
HU : # Több fragmentumot mutató variáns (alfa lánc vagy instabil variáns ...)	!! Csúcs a zóna határánál (zónaeltolódás kockázata)
TR : # Birden çok fraksiyonun görüldüğü varyant (alfa zincir veya kararsız varyant ...)	!! Bölge sınırında pik değeri (bölge kayması riski)
CZ : # Varianta s několika zobrazenými frakcemi (alfa řetězec nebo nestálá varianta ...)	!! Pik na hranici zóny (riziko posunu zóny)
BG : # Появеща се есариум с няколко фракции (алфа-верука или нестационарен есариум ...)	!! Пик на границата на зоната (опасност от преместване на зоната)
NO : # Variant med flere fraksjoner vises (Alpha-kjeden eller ustabil variant ...)	!! Topp i sonegrense (risiko for soneforskyvning)
DK : # Variant med flere fraktioner vist (alfakæde eller ustabil variant ...)	!! Top i zonegrænse (fare for zoneskiftning)
CN : # 显示了多个区带的变体 (α 链或不稳定变体 ...)	!! 区边界峰重叠 (有区位转移的风险)
RU : # Вариация с отображением нескольких фракций (альфа-цепь или неустойчивая вариация ...)	!! Пик на границе зоны (риск смещения зоны)
JP : # 数個のフラクションが表示される変異体 (α 鎖または不安定な変異体 ...)	!! ゼーン境界でのピーク (ゾーンソフトフォーカスがある)
LV : # Tiek parādīts variants ar vairākām frakcijām (alfa ķēde vai nestabils variants u.c.)	!! Maksimālā vērtība zona robežās (zonas maiņas risks)
SK : # Variant s viacerými zobrazenými frakciami (alfa-řetazec alebo nestabilný variant ...)	!! Špička v hranici zóny (riziko posunutia zóny)
EE : # Kuvatud on mitme fraktsiooniga variant (alfa-ahel või ebastabiilne variant ...)	!! Pikk tsoonipiir (tsoonide nihke risk)
VN : # Biến cố vài miền hiển thị (biến chuỗi alpha hay biến không ổn định ...)	!! Đỉnh nằm trong ranh giới vùng (nguy cơ thay đổi vùng)

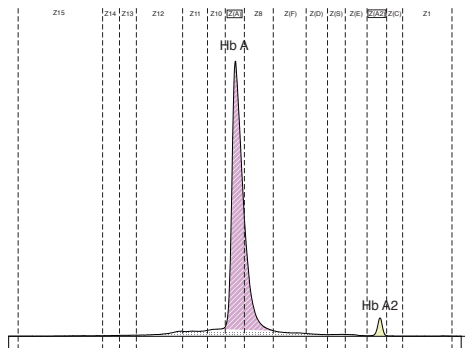
FR : Rappel : Dans chaque zone, les variants Hb sont listés selon leur temps de migration de la droite vers la gauche
GB : Reminder: In each zone, Hb variants are sorted according to their migration time from the right to the left
DE : Hinweis : Die Hb-Varianten werden in allen Zonen nach ihrer Migrationszeit von rechts nach links sortiert
NL : Herinnering : in elke zone worden Hb varianten op basis van hun migratietijd van rechts naar links gesorteerd
IT : NB : In ciascuna zona, le varianti Hb sono elencate in base al loro tempo di migrazione da destra verso sinistra
ES : NOTA : En cada zona, las variantes de la Hb son listadas según su tiempo de migración de derecha a izquierda
PT : Lembrete : Em cada zona, as variantes Hb são ordenadas de acordo com o seu tempo de migração da direita para a esquerda
SV : PÅMINNELSE : I varje zon, sorteras Hb-varianter enligt deras migratinstid från höger till vänster
GR : Υπενθύμιση : Σε κάθε ζώνη, οι παραλλαγές Hb ταξινομούνται ανάλογα με τον χρόνο ηλεκτροφόρησης τους από τα δεξιά προς τα αριστερά
HR : Podsjetnik : Hb varijante razvrstane su u svakoj zoni prema svom vremenu migracije s desne na lijevu stranu
IT : Pirmenname : kiekvienoje zonoje Hb variantai pagal migracijos laiką surūšiuoti iš dešinės į kairę.
PL : Przypomnienie : W każdej strefie odmiany Hb są sortowane według czasu migracji, od prawej do lewej
RO : Memento : În fiecare zonă, variantele Hb sunt sortate în funcție de timpul de migrare de la dreapta la stânga
CS : Podsetník : U svakoј zoni, Hb varijante se sortiraju prema svom vremenu migracija sa desna na levo
HU : Émlékeztető : A Hb-variánsok mindegyik zónában a migrációs idejüknek megfelelően rendeződnek jobbról balra
TR : Hatırlatma : Her bir bölgede, Hb varyantları sağdan sola migrasyon sürelerine göre sıralandırlır
CZ : Pripomínka : V každé zóně jsou varianty Hb rozříděny podle své doby migrace zprava doleva
BG : Напомняне : Hb вариантите във всяка зона се сортират от дясно наляво според времето им на миграция
NO : Påminnelse : I hver sone, er Hb-varianter sortert i henhold til deres migreringstid fra høyre til venstre
DK : Påmindelse : I hver zone sorteres Hb-varianter efter deres migratinstid fra højre til venstre
CN : 提示 : 在每个区中, 根据 Hb 变体从右到左的电泳时间进行排序
RU : Напоминание : вариации Hb сортируются по времени миграции в каждой зоне справа налево
JP : 注意 : 各ゾーンにおいて、Hb変異体は移動時間に従って右から左にソートされ (逆で替えられ) ます
LV : Atgādinājums! Katrā zonā Hb varianti tiek sakārtoti pēc to migrācijas laika no labās uz kreiso pusi
SK : Pripomienka : Varianty Hb sú v každej zóne usporiadané sprava doľava podľa času migrácie
EE : Meelespea : Igas tsoonis sorteeritakse Hb variandid vastavalt nende migratsiooniale paremalt vasakule.
VN : Xin nhắc lại : Ở mỗi vùng, biến thể Hb được sắp xếp theo thời gian di chuyển từ phải sang trái

SCHÉMAS / FIGURES

ABBILDUNGEN - FIGUREN - FIGURE - FIGURAS - BILDER - EIKONES - SLIKE - PAVEIKSLAI - RYSUNKI - FIGURI -
ÁBRÁK - ŞEKİLLER - OBRÁZKY - ΦΙΓΥΡΗ - FIGURER - 插图 - РИСУНКИ - 図 - CIPARI - JOONISED - SƠ ĐỒ

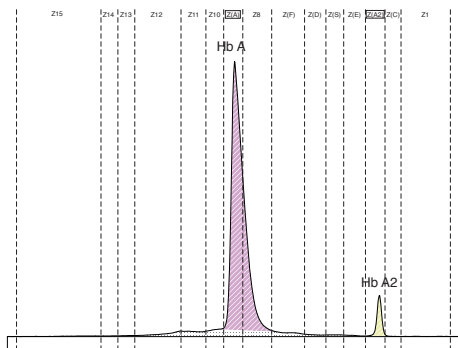
PROFILS ÉLECTROPHORÉTIQUES - ELECTROPHORETIC PATTERNS

1



Sang normal
Normal blood sample

2



Sang bêta-thalassémique
Blood sample with beta-thalassemia

FR : PROFILS ÉLECTROPHORÉTIQUES
GB : ELECTROPHORETIC PATTERNS
DE : ELEKTROPHORESEMUSTER
NL : ELEKTROPHORETISCHE PATRONEN
IT : PROFILI ELETTROFORETICI
ES : PERFILES ELECTROFORETICOS
PT : PADRÕES ELETTROFORETICOS
SV : ELEKTROFORETISKA MÖNSTER
GR : ΗΛΕΚΤΡΟΦΟΡΗΤΙΚΑ ΠΡΟΤΥΠΑ
HR : ELEKTROFORETSKI OBRASCI
LT : ELEKTROFOREZES ŠABLONAI
PL : OBRAZY ELEKTROFORETYCZNE
RO : TIPARE ELECTROFORETICE
CS : ELEKTROFORETSKÍ ŠABLONI
HU : ELEKTROFORETIKUS MINTÁZATOK
TR : ELEKTROFORETİK PATERNLER
CZ : ELEKTROFORETICKÉ TYPY
BG : ЕЛЕКТРОФОРΕΤИЧНИ МОДЕЛИ
NO : ELEKTROFORETISKE MONSTRE
DK : ELEKTROFORETISCHE MONSTRE
CN : 电泳图谱
RU : ЭЛЕКТРОФОРΕΤИЧЕСКИЕ ПРОФИЛИ
JP : 電気泳動パターン
LV : ELEKTROFORETISKIE SPEKTRI
SK : ELEKTROFOREZNE VZORY
EE : ELEKTROFORETILISED MUSTRID
VN : MÔ HÌNH ĐIỆN DI

Sang normal
Normal blood sample
Normalblutprobe
Normaal bloedmonster
Sangue normale
Sangre normal
Amostra de sangue normal
Normalt blodprov
Φυσιολογικό δείγμα αίματος
Normalan uzorak krvi
Normalus kraujo mėginys
Próbka krwi prawidłowej
Próbă normală de sânge
Normalan uzorak krvi
Normalí vérminta
Normal kan numenesi
Normalní vzorek krve
Normalna krvana proba
Normal blodprøve
Normal blodprøve
正常血液样品
Образец нормальной крови
正常血液サンプル
Normalis asins paraugs
Vzorka normalnej krvi
Normalne vereproov
Mẫu máu thông thường

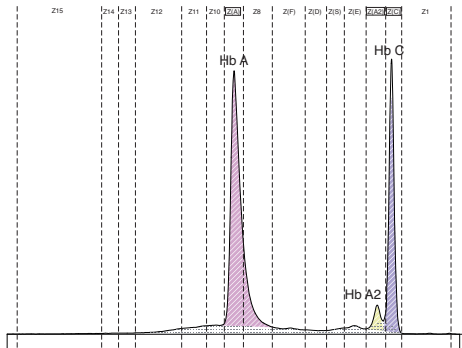
Sang bêta-thalassémique
Blood sample with beta-thalassemia
Blutprobe mit Beta-Thalassämie
Bloedmonster met bèta-thalassemie
Sangue beta-thalassémico
Sangre con beta talasemia
Amostra de sangue com beta-talassemia
Blodprov med beta-thalassemi
Δείγμα αίματος με βήτα-θαλασσαιμία
Uzorak krvi s beta-talasemijom
Pacientu, serganciu beta talasemija, kraujo mėginys
Próbka krwi z beta-talasemią
Próbă de sânge cu beta-talasemie
Uzorak krvi sa beta-talasemijom
Bêta-falasszémias vérminta
Beta-talassemi içeren kan numenesi
Vzorek krve s beta talasemii
Krvana proba s beta-talasemija
Blodprøve med beta-talassemi
Blodprøve med beta-talassemi
β-地中海貧血的血液样品
Образец крови с бета-талассемией
βサラセミアの血液サンプル
Asins paraugs ar beta talasēmiju
Vzorka krvi s beta-talassēmiju
Beeta-talassemiaga vereproov
Mẫu máu có beta-thalassemia

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PROFILS ÉLECTROPHORÉTIQUES - ELECTROPHORETIC PATTERNS

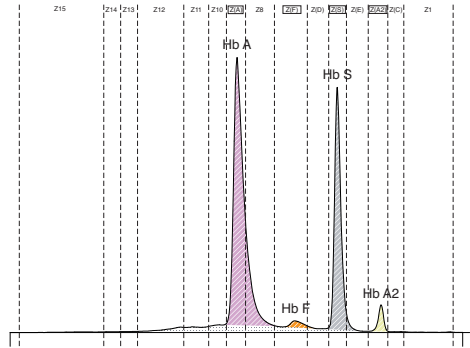
3



Sang de patient hétérozygote avec variant Hb C
Blood sample from heterozygous patient with Hb C variant

FR : Sang de patient hétérozygote avec variant Hb C
GB : Blood sample from heterozygous patient with Hb C variant
DE : Blutprobe eines heterozygoten Patienten mit Hb C-Variante
NL : Bloedmonster van heterozygote patiënt met Hb C variant
IT : Sangue di paziente eterozygote con variante Hb C
ES : Sangre de paciente heterocigoto con la variante Hb C
PT : Amostra de sangue de doente heterozigótico com variante Hb C
SV : Blodprov från heterozygot patient med Hb C-variant
GR : Δείγμα αίματος από ετερόζυγο ασθενή με παραλλαγή Hb C
HR : Uzorak krvi heterozigotnog bolesnika s varijantom Hb C
LT : Heterozigotinio paciento kraujo, kuriame yra Hb C varianto, kraujo mėginys
PL : Próba krwi od heterozygotycznego pacjenta z odmianą Hb C
RO : Probă de sânge de la pacient heterozigot cu varianta Hb C
CS : Uzorak krvi od heterozigotnog pacijenta s Hb C varijantom
HU : Heterozigóta beteg vérmintéje Hb C variánsal
TR : Hb C varyantı taşıyan heterozigot hastasına ait kan numunesi
CZ : Vzorek kve heterozigotního pacienta s variantou Hb C
BG : Кръвна проба от хетерозиготен пациент с Hb C вариант
NO : Blodprobe fra heterozygot pasient med Hb C-variant
DK : Blodprobe fra heterozygot patient med Hb C-variant
CN : 来自 Hb C 杂合患者的血液样品
RU : Образец крови от гетерозиготного пациента с вариацией Hb C
JP : Hb C変異体を含むヘテロ接合体患者からの血液サンプル
LV : Heterozigota pacienta asins paraugs ar Hb C variantu
SK : Vzorka krvi od heterozigotného pacienta s variantom Hb C
EE : Vereproov Hb C variantiga heterozigotselt patsiendilt
VN : Mẫu máu của bệnh nhân bị bệnh dị hợp tử với biến thể Hb C

4



Sang de patient hétérozygote avec variant Hb S
Blood sample from heterozygous patient with Hb S variant

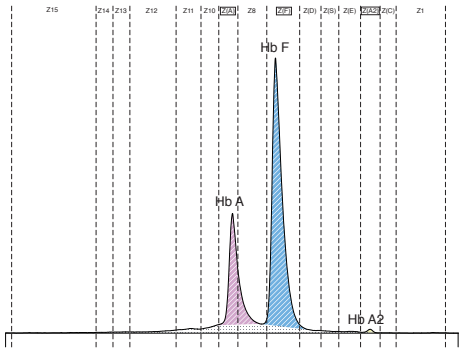
Sang de patient hétérozygote avec variant Hb S
Blood sample from heterozygous patient with Hb S variant
Blutprobe eines heterozygoten Patienten mit Hb S-Variante
Bloedmonster van heterozygote patiënt met Hb S variant
Sangue di paziente eterozygote con variante Hb S
Sangre de paciente heterocigoto con la variante Hb S
Amostra de sangue de doente heterozigótico com variante Hb S
Blodprov från heterozygot patient med Hb S-variant
Δείγμα αίματος από ετερόζυγο ασθενή με παραλλαγή Hb S
Uzorak krvi heterozigotnog bolesnika s varijantom Hb S
Heterozigotinio paciento kraujo, kuriame yra Hb S varianto, mėginys
Próba krwi od heterozygotycznego pacjenta z odmianą Hb S
Probă de sânge de la pacient heterozigot cu varianta Hb S
Uzorak krvi od heterozigotnog pacijenta sa Hb S varijantom
Heterozigóta beteg vérmintéje Hb S variánsal
TR : Hb S varyantı taşıyan heterozigot hastasına ait kan numunesi
CZ : Vzorek kve heterozigotního pacienta s variantou Hb S
BG : Кръвна проба от хетерозиготен пациент с Hb S вариант
Blodprobe fra heterozygot pasient med Hb S-variant
Blodprobe fra heterozygot patient med Hb S-variant
来自 Hb S 杂合患者的血液样品
RU : Образец крови от гетерозиготного пациента с вариацией Hb S
Hb S変異体を含むヘテロ接合体患者からの血液サンプル
Heterozigota pacienta asins paraugs ar Hb S variantu
Vzorka krvi od heterozigotného pacienta s variantom Hb S
Vereproov Hb S variantiga heterozigotselt patsiendilt
Mẫu máu của bệnh nhân bị bệnh dị hợp tử với biến thể Hb S

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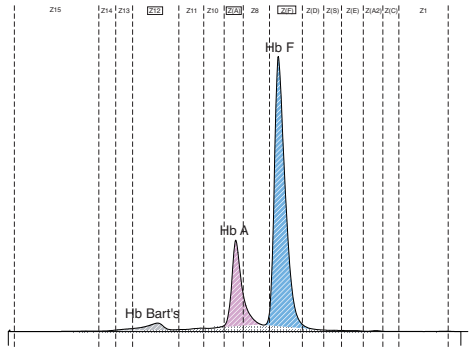
PROFILS ÉLECTROPHORÉTIQUES - ELECTROPHORETIC PATTERNS

5



Sang normal de bébé (âgé de 3 semaines)
 Normal blood sample from baby (3 weeks old)

6



Sang de bébé avec Hb Bart's
 Baby blood sample with Hb Bart's

FR : Sang normal de bébé (âgé de 3 semaines)
 GB : Normal blood sample from baby (3 weeks old)
 DE : Normalblutprobe eines Säuglings (Alter: 3 Wochen)
 NL : Normaal bloedmonster van baby (3 weken oud)
 IT : Sangue normale di neonato (età 3 settimane)
 ES : Sangre normal de bebé (3 semanas de edad)
 PT : Amostra de sangue normal de bebé (3 semanas de idade)
 SV : Normalt blodprov från baby (3 veckor gammal)
 GR : Φυσιολογικό δείγμα αίματος από βρέφος (ηλικίας 3 εβδομάδων)
 HR : Normalan uzorak krvi dojenčeta (u dobi od 3 tjedna)
 LT : Normalus kūdikio (3 savaičių amžiaus) kraujo mėginys
 PL : Próbk krwi prawidłowej od niemowlęcia (3-tygodniowego)
 RO : Probă normală de sânge de la bebeluș (trei săptămâni)
 CS : Normalan uzorak krvi od bebe (stare 3 nedelje)
 HU : 3 hetes csecsemő normál vérmintája
 TR : Bebekten alınan normal kan numunesi (3 haftalık)
 CZ : Normální vzorek krve malého dítěte (věk 3 týdnů)
 BG : Нормална кръвна проба от бебе (на възраст 3 седмици)
 NO : Normal blodprøve fra nyfødt barn (3 uker gammel)
 DK : Normalt blodprøve fra baby (3 uger gammel)
 CN : 来自婴儿 (3个月) 的正常血液样品
 RU : Образец нормальной крови младенца (возраст — 3 недели)
 JP : 新生児 (生後3週) からの正常血液サンプル
 LV : Mazuļa (3 nedēļas vecs) normāls asiņu paraugs
 SK : Vzorka normálnej krvi od novorodenca (vo veku 3 týždňov)
 EE : Normaalne vereproov imikul (3-nädalane)
 VN : Mẫu máu thông thường của trẻ sơ sinh (3 tuần tuổi)

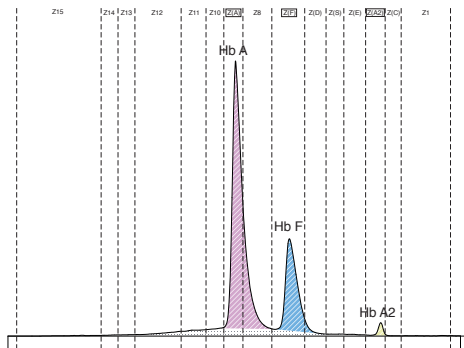
Sang de bébé avec Hb Bart
 Baby blood sample with Hb Bart's
 Blutprobe eines Säuglings mit Hb-Bart's
 Bloedmonster van baby met Hb Bart
 Sangue di neonato con Hb Bart
 Sangre de bebé con Hb Bart
 Amostra de sangue de bebé com Hb de Bart
 Blodprov från baby med Hb Bart's
 Δείγμα αίματος βρέφους με Hb Bart's
 Uzorak krvi dojenčeta s Bartovim Hb
 Kūdikio kraujo, kuriame yra Hb Bart, mėginys
 Próbk krwi niemowlęcia z Hb Barta
 Probă de sânge de la bebeluș cu Hb Bart's
 Uzorak krvi bebe sa Hb Bart's
 Csecsemő vérmintája Hb Barttal
 Hb Bart's taşınan bebeğin ait kan numunesi
 Vzorek krve malého dítěte s Bartovym Hb
 Кръвна проба от бебе с Hb на Bart
 Blodprøve fra nyfødt barn med Hb Bart's
 Babyblodprøve med Hb Bart's
 含 Hb Bart's 的血液样品
 Образец крови младенца с гемоглобином Барта
 Hb-バートを含む新生児の血液サンプル
 Mazuļa asiņu paraugs ar Hb Bart
 Vzorka krvi od novorodenca s Hb Bartovym Hb
 Imiku vereproov Hb Bartiga
 Mẫu máu của trẻ sơ sinh có Hb Bart's

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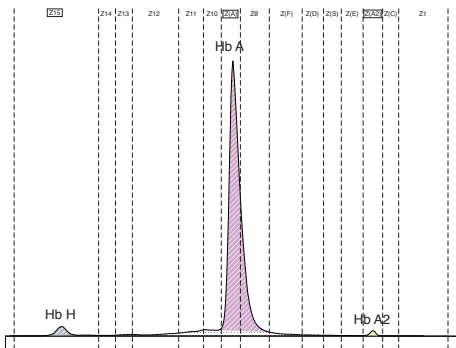
PROFILS ÉLECTROPHORÉTIQUES - ELECTROPHORETIC PATTERNS

7



Sang avec Hb F élevée (jeune enfant)
 Blood sample with elevated Hb F (young child)

8



Sang avec Hb H
 Blood sample with Hb H

FR : Sang avec Hb F élevée (jeune enfant)
 GB : Blood sample with elevated Hb F (young child)
 DE : Blutprobe mit erhöhtem Hb F (Kleinkind)
 NL : Bloedmonster met verhoogd Hb F niveau (klein kind)
 IT : Sangue con Hb F alta (bambino)
 ES : Sangre con Hb F elevada (niño de corta edad)
 PT : Amostra de sangue com Hb F elevada (criança pequena)
 SV : Blodprov med förhöjd Hb F (litt barn)
 GR : Δείγμα αίματος με αυξημένο επίπεδο Hb F (μικρό παιδί)
 HR : Uzorak krvi s povišenom vrijednošću Hb F (malo dijete)
 LT : Kraujo, kuriame padidėjęs Hb F kiekis, mėginys (maža vaiko)
 PL : Próba krwi o podwyższonym stężeniu Hb F (male dziecko)
 RO : Probă de sânge cu Hb F crescută (copil mic)
 CS : Uzorak krvi sa povišením Hb F (malo dete)
 HU : Vércinta emelkedett Hb F-tel (kisgyermek)
 TR : Yüksek Hb F içeren kan numunesi (genç çocuk)
 CZ : Vzorek krve se zvýšeným Hb F (dítě)
 BG : Кръвна проба с повишен Hb F (малко дете)
 NO : Blodprøve med forhøyet Hb F (små barn)
 DK : Blodprøve med forhøjet Hb F (lille barn)
 CN : Hb F 升高 (幼儿) 的血液样品
 RU : Образец крови с повышенным уровнем Hb F (ребенок младшего возраста)
 JP : Hb Fが高値の血液サンプル (幼児)
 LV : Asins paraugs ar paaugstinātu Hb F līmeni (mazs bērns)
 SK : Vzorka krvi so zvýšenou hladinou Hb F (dôčča)
 EE : Vereproov kõrgeunud Hb F-ga (noor laps)
 VN : Mẫu máu có Hb F gia tăng (trẻ nhỏ)

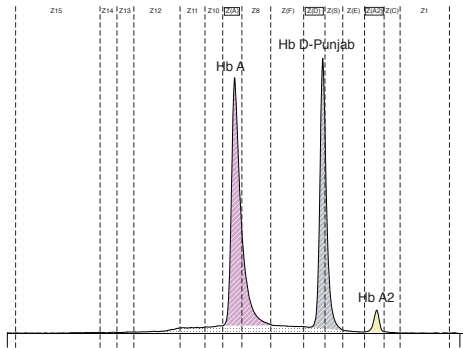
Sang avec Hb H
 Blood sample with Hb H
 Blutprobe mit Hb H
 Bloedmonster met Hb H
 Sangue con Hb H
 Sangre con Hb H
 Amostra de sangue com Hb H
 Blodprov med Hb H
 Δείγμα αίματος με Hb H
 Uzorak krvi s Hb H
 Kraujo, kuriame yra Hb H, mėginys
 Próba krwi z Hb H
 Probă de sânge cu Hb H
 Uzorak krvi sa Hb H
 Vércinta Hb H-val
 Hb H içeren kan numunesi
 Vzorek krve s Hb H
 Кръвна проба с Hb H
 Blodprøve med Hb H
 Blodprøve med Hb H
 含 Hb H 的血液样品
 Образец крови с Hb H
 Hb Hを含む血液サンプル
 Asins paraugs ar Hb H
 Vzorka krvi s Hb H
 Vereproov Hb H-ga
 Mẫu máu có Hb H

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PROFILS ÉLECTROPHORÉTIQUES - ELECTROPHORETIC PATTERNS

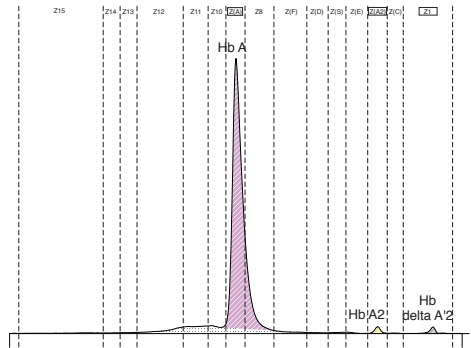
9



Sang de patient hétérozygote avec variant Hb D-Punjab
Blood sample from heterozygous patient with Hb D-Punjab variant

FR : Sang de patient hétérozygote avec variant Hb D-Punjab
GB : Blood sample from heterozygous patient with Hb D-Punjab variant
DE : Blutprobe eines heterozygoten Patienten mit Hb D-Punjab-Variante
NL : Bloedmonster van heterozygote patiënt met Hb D-Punjab variant
IT : Sangue di paziente eterozygote con variante Hb D-Punjab
ES : Sangre de paciente heterocigoto con la variante Hb D-Punjab
PT : Amostra de sangue de doente heterocigótico com variante Hb D-Punjab
SV : Blodprov från heterozygot patient med Hb D-Punjab variant
GR : Δείγμα αίματος από ετερόζυγο ασθενή με παραλλαγή Hb D-Punjab
HR : Uzorak krvi heterozigotnog bolesnika s varijantom Hb D-Punjab
LT : Heterozigotinio paciento kraujo, kuriame yra Hb D-Punjab varianto, mėginys
PL : Próba krwi od heterozygotycznego pacjenta z odmianą Hb D-Punjab
RO : Probă de sânge de la pacient heterozigot cu varianta Hb D-Punjab
CS : Uzorak krvi od heterozigotného pacienta sa Hb D-Punjab variantom
HU : Heterozigóta beteg vérmintája Hb D-Punjab variánsával
TR : Hb D-Punjab varyantı taşıyan heterozigot hastasına ait kan numunesi
CZ : Vzorek krve heterozigotního pacienta s variantou Hb D-Punjab
BG : Кръвна проба от хетерозиготен пациент с Hb D-Punjab вариант
NO : Blodprobe fra heterozygot pasient med Hb D-Punjab variant
DK : Blodprobe fra heterozygot patient med Hb D-Punjab-variant
CN : 来自 Hb D-Punjab 变体杂合患者的血液样品
RU : Образец крови от гетерозиготного пациента с вариацией Hb D-Punjab
JP : Hb D-Punjab変異体を含むヘテロ接合体患者からの血液サンプル
LV : Heterozigota pacienta asins paraugs ar Hb D-Punjab variantu
SK : Vzorka krvi od heterozigotného pacienta s variantom Hb D-Punjab
EE : Vereproov Hb D-Punjab variantidiga heterosügotselt patsiendilt
VN : Mẫu máu của bệnh nhân bị bệnh di hpp tú với biến thể Hb D-Punjab

10



Sang de patient hétérozygote avec variant delta Hb A2
Blood sample from heterozygous patient with delta Hb A2 variant

Sang de patient hétérozygote avec variant delta Hb A2
Blood sample from heterozygous patient with delta Hb A2 variant
Blutprobe eines heterozygoten Patienten mit Delta-Hb A2-Variante
Bloedmonster van heterozygote patiënt met delta Hb A2 variant
Sangue di paziente eterozygote con variante delta Hb A2
Sangre de paciente heterocigoto con variante delta Hb A2
Amostra de sangue de doente heterocigótico com variante delta Hb A2
Blodprov från heterozygot patient med delta Hb A2-variant
Δείγμα αίματος από ετερόζυγο ασθενή με παραλλαγή δέλτα Hb A2
Uzorak krvi heterozigotnog bolesnika s varijantom delta Hb A2
Heterozigotinio paciento kraujo, kuriame yra delta Hb A2 varianto, mėginys
Próba krwi od heterozygotycznego pacjenta z odmianą delta Hb A2
Probă de sânge de la pacient heterozigot cu varianta delta Hb A2
Uzorak krvi od heterozigotného pacienta sa delta Hb A2 varijantom
Heterozigóta beteg vérmintája delta Hb A2 variánsával
Delta Hb A2 varyantı taşıyan heterozigot hastasına ait kan numunesi
Vzorek krve heterozigotního pacienta s variantou delta Hb A2
Кръвна проба от хетерозиготен пациент с дельта Hb A2 вариант
Blodprobe fra heterozygot pasient med delta Hb A2 variant
Blodprobe fra heterozygot patient med delta Hb A2-variant
来自 Hb A2 变体杂合患者的血液样品
Образец крови от гетерозиготного пациента с вариацией delta Hb A2
デルタHb A2変異体を含むヘテロ接合体患者からの血液サンプル
Heterozigota pacienta asins paraugs ar delta Hb A2 variantu
Vzorka krvi od heterozigotného pacienta s variantom Hb A2
Vereproov delta Hb A2 variantidiga heterosügotselt patsiendilt
Mẫu máu của bệnh nhân bị bệnh di hpp tú với biến thể Hb A2 delta

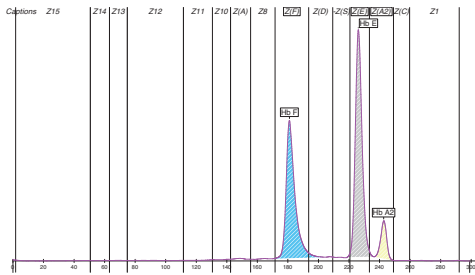
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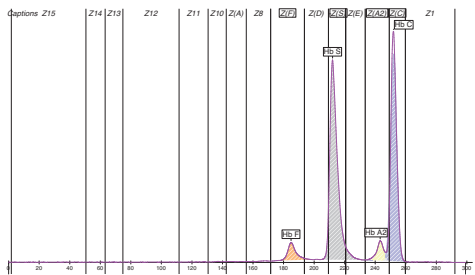
PROFILS ÉLECTROPHORÉTIQUES - ELECTROPHORETIC PATTERNS

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Sang de patient homozygote avec variant Hb E et fraction Hb F élevée
Blood sample from homozygous patient with Hb E variant and elevated Hb F



Sang de patient hétérozygote composite avec variants Hb S et Hb C
Blood sample from compound heterozygous patient with Hb S & Hb C variants

- FR : Sang de patient homozygote avec variant Hb E et fraction Hb F élevée
- GB : Blood sample from homozygous patient with Hb E variant and elevated Hb F
- DE : Blutprobe eines homozygoten Patienten mit Hb E-Variante und erhöhtem Hb F
- NL : Bloedmonster van homozygote patiënt met Hb E variant en verhoogd Hb F niveau
- IT : Sangue di paziente omozigote con variante Hb E e frazione Hb F alta
- ES : Sangre de paciente homocigoto con la variante Hb E y la fracción Hb F elevada
- PT : Amostra de sangue de doente heterozigótico com variante Hb E e Hb F elevada
- SV : Blodprov från heterozygot patient med Hb E variant och förhöjd Hb F
- GR : Δείγμα αίματος από ομόζυγο ασθενή με παραλλαγή Hb E και αυξημένο επίπεδο Hb F
- HR : Uzorak krvi homozigotnog bolesnika s varijantom Hb E i povišenom vrijednošću Hb F
- LT : Homozigotinio paciento kraujo, kuriame yra Hb E varianto ir padidėjęs Hb F kiekis, mėginys
- PL : Próbk krwi od homozygotycznego pacjenta z odmianną Hb E i podwyższonym stężeniem Hb F
- RO : Probă de sânge de la pacient heterozigot cu varianta Hb E și Hb F crescută
- CS : Uzorak krvi od heterozigotného pacienta sa Hb E variantom i povišením Hb F
- HU : Homozigóta beteg vérmintája Hb E variánsal és emelkedett Hb F-fel
- TR : Hb E variantı ve yüksek Hb F taşıyan homozigot hastasına ait kan numunesi
- CZ : Vzorek krve homozigotního pacienta s variantou Hb E a zvýšením Hb F
- BG : Кръвна проба от хомозиготен пациент с Hb E вариант и повишен Hb F
- NO : Blodprobe fra homozygot pasient med Hb E-variant og forhøyet Hb F
- DK : Blodprobe fra homozygot patient med Hb E-variant og forhøjet Hb F
- CN : 来自 Hb E 变体和 Hb F 升高杂合患者的血液样品
- RU : Образец крови от гомозиготного пациента с вариацией Hb E и повышенным уровнем Hb F
- JP : Hb E変異体および高値のHb Fを含むホモ接合患者からの血液サンプル
- LV : Homozigota pacienta asins paraugs ar Hb E variantu un paaugstinātu Hb F līmeni
- SK : Vzorka krvi od homozigotného pacienta s variantom Hb E a zvýšenou hladinou Hb F
- EE : Vereproov Hb E variandi ja kõrgenevad Hb F-iga heterosügootsest patsiendilt
- VN : Mẫu máu của bệnh nhân bị bệnh di hợp từ với biến thể Hb E và Hb F gia tăng

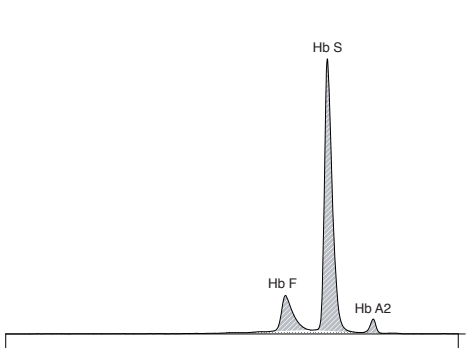
- Sang de patient hétérozygote composite avec variants Hb S et Hb C
- Blood sample from compound heterozygous patient with Hb S & Hb C variants
- Blutprobe eines compound-heterozygoten Patienten mit Hb S- und Hb C-Varianten
- Bloedmonster van samengestelde heterozygote patiënt met Hb S en Hb C varianten
- Sangue di paziente eterozigote composto con varianti Hb S e Hb C
- Sangre de paciente heterocigoto compuesto con las variantes Hb S y Hb C
- Amostra de sangue de doente heterozigótico composto com variantes Hb S e Hb C
- Blodprov från förenad heterozygot patient med Hb S & Hb C-varianter
- Δείγμα αίματος από σύνθετο ετερόζυγο ασθενή με παραλλαγές Hb S & Hb C
- Uzorak krvi složnog heterozigotnog bolesnika s varijantama Hb S i Hb C
- Paciento kraujo, kuriame yra heterozigotinių junginių ir Hb S bei Hb C variantų, mėginys
- Próbka krwi od heterozygotycznego pacjenta z jednoczesną obecnością odmianną Hb S oraz Hb C
- Probă de sânge de la pacient heterozigot compus cu variantele Hb S și Hb C
- Uzorak krvi od složnog heterozigotného pacienta sa Hb S & Hb C varijantama
- Összetett heterozigóta beteg vérmintája Hb S és Hb C variánsokkal
- Hb S ve Hb C varyantlarını taşıyan blegik heterozigot hastasına ait kan numunesi
- Vzorek krve sdruženého heterozigotního pacienta s variantami Hb S a Hb C
- Кръвна проба от пациент със съставна хетерозиготност с Hb S и Hb C варианти
- Blodprobe fra sammensatt heterozygot pasient med Hb S og Hb C-varianter
- Blodprobe fra heterozygot patient med Hb S- og Hb C-varianter
- 来自 Hb S & Hb C 变体杂合患者的血液样品
- Образец крови от компаунд-гетерозигота с вариациями Hb S и Hb C
- Hb SおよびHb C変異体を含む複合ヘテロ接合患者からの血液サンプル
- Kompaundta heterozigota pacienta asins paraugs ar Hb S un Hb C variantu
- Vzorka krvi od heterozigotného pacienta s variantami Hb S a Hb C
- Vereproov Hb S ja Hb C variantidega ühend-heterosügootsest patsiendilt
- Mẫu máu của bệnh nhân bị bệnh di hợp từ kép với biến thể Hb S & Hb C

SCHÉMAS / FIGURES

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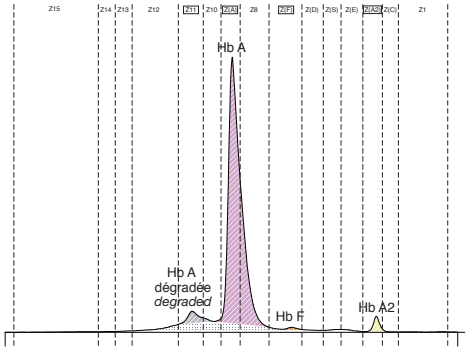
PROFILS ÉLECTROPHORÉTIQUES - ELECTROPHORETIC PATTERNS

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Sang de patient homozygote avec Hb F et variant Hb S
Blood sample from homozygous patient with Hb F and Hb S variant

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Sang avec Hb A dégradée (Hb A3) et Hb F faible
Blood sample with degraded Hb A (Hb A3) and faint Hb F

- FR : Sang de patient homozygote avec Hb F et variant Hb S
- GB : Blood sample from homozygous patient with Hb F and Hb S variant
- DE : Blutprobe eines homozygoten Patienten mit Hb F- und Hb S-Variante
- NL : Bloedmonster van homozygote patiënt met Hb F en Hb S variant
- IT : Sangue di paziente omozigote con Hb F e variante Hb S
- ES : Sangre de paciente homocigoto con Hb F y la variante Hb S
- PT : Amostra de sangue de doente homocigótico com variantes Hb F e Hb S
- SV : Blodprov från homozygot patient med Hb F- och Hb S-variant
- GR : Δείγμα αίματος από ομόζυγο ασθενή με παραλλαγή Hb F και Hb S
- HR : Uzorak krvi homocigotnog bolesnika s varijantama Hb F i Hb S
- LT : Homocigotinio paciento kraujo, kuriame yra Hb F bei Hb S variantų, mėginys
- PL : Próbk krwi od homocigotycznego pacjenta z odmianną Hb F i Hb S
- RO : Probă de sânge de la pacient heterocigot cu variantele Hb F și Hb S
- CS : Uzorak krvi od heterocigotnog pacijenta sa Hb F i Hb S varijantom
- HU : Homocigóta beteg vérmintája Hb F-fei és Hb S variánsal
- TR : Hb F ve Hb S varyantlarını taşıyan homozigot hastasına ait kan numunesi
- CZ : Vzorek krve homocigotního pacienta s variantami Hb F a Hb S
- BG : Кръвна проба от хомозиготен пациент с Hb F u Hb S вариант
- NO : Blodprobe fra homozygot pasient med Hb F og Hb S variant
- DK : Blodprobe fra heterozygot patient med Hb F- og Hb S-variant
- CN : 来自 Hb F 和 Hb S 杂合患者的血液样品
- RU : Образец крови от гомозиготного пациента с вариациями Hb F и Hb S
- JP : Hb FおよびHb S変異体を含むホモ接合患者からの血液サンプル
- LV : Heterocigota pacienta asins paraugs ar Hb F un Hb S variantu
- SK : Vzorka krvi od homozygotného pacienta s variantmi Hb F a Hb S
- EE : Vereproov Hb F ja Hb S variantidga homosiigotselt patsiidilt
- VN : Mẫu máu của bệnh nhân bị bệnh di hpp từ vôi biến Hb F và Hb S

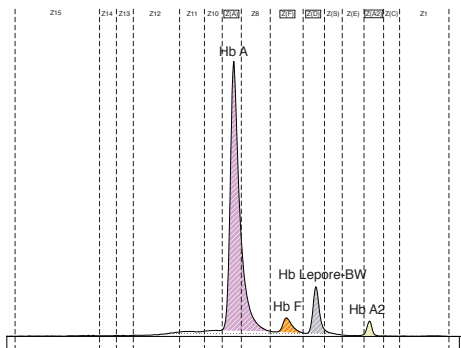
- Sang avec Hb A dégradée (Hb A3) et Hb F faible
- Blood sample with degraded Hb A (Hb A3) and faint Hb F
- Blutprobe mit degradiertem Hb A (Hb A3) und schwachem Hb F
- Bloedmonster met afgebroken Hb A (Hb A3) and nauwelijks waarneembare Hb F
- Sangue con Hb A degradata (Hb A3) e Hb F bassa
- Sangre con Hb A degradada (Hb A3) y Hb F débil
- Amostra de sangue com Hb A (Hb A3) degradada e Hb F baixa
- Blodprov med nedbrutet Hb A (Hb A3) och svagt Hb F
- Δείγμα αίματος με αποσυντεθειμένη Hb A (Hb A3) και αμυδρή Hb F
- Uzorak krvi s degradiranim Hb A (Hb A3) i slabim Hb F
- Kraujo, kuriame yra suskaidius Hb A (Hb A3) ir išblukusio Hb F mėginys
- Próbka krwi z rozłożoną Hb A (Hb A3) i śladową obecnością Hb F
- Probă de sânge cu Hb A degradată (Hb A3) și Hb F slabă
- Uzorak krvi sa degradiranim Hb A (Hb A3) i niskim Hb F
- Vérminta degradálódott Hb A-val (Hb A3) és halvány Hb F-fei
- İndirgenmiş Hb A (Hb A3) ve belirsizsayıf Hb F iperen kan numunesi
- Vzorek krve s degradovaným Hb A (Hb A3) a slabým Hb F
- Кръвна проба с разграден Hb A (Hb A3) и малко количество Hb F
- Blodprobe med degradert Hb A (Hb A3) og svakt Hb F
- Blodprobe med nedbrudt Hb A (Hb A3) og svag Hb F
- Hb A (Hb A3) 降低和Hb F虚弱的血液样品
- Образец крови с подвергнувшимся разложению Hb A (Hb A3) и низким Hb F
- 劣化したHb A (Hb A3) および僅少のHb Fを含む血液サンプル
- Asins paraugs ar noārdētu Hb A (Hb A3) un nelielu Hb F daudzumu
- Vzorka krvi s degradovaným Hb A (Hb A3) a nevýrazným Hb F
- Vereproov lagunenud Hb A (Hb A3) ja nõrga Hb F-ga
- Mẫu máu có Hb A (Hb A3) suy giảm và Hb F yếu

SCHÉMAS / FIGURES

ABBILDUNGEN - FIGUREN - FIGURE - FIGURAS - BILDER - EIKONES - SLIKE - PAVEIKSLAI - RYSUNKI - FIGURI -
 ÁBRÁK - ŞEKİLLER - OBRÁZKY - ΦΙΓΥΡΗ - FIGURER - 插图 - ΡΙΣΥΗΚΗ - 図 - CIPARI - JOONISED - SƠ ĐỒ

PROFILS ÉLECTROPHORÉTIQUES - ELECTROPHORETIC PATTERNS

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Sang de patient hétérozygote avec variant Hb Lepore-Boston-Washington
 Blood sample from heterozygous patient with Hb Lepore-Boston-Washington variant

FR : Sang de patient hétérozygote avec variant Hb Lepore-Boston-Washington
 GB : Blood sample from heterozygous patient with Hb Lepore-Boston-Washington variant
 DE : Blutprobe eines heterozygoten Patienten mit Hb Lepore-Boston-Washington-Variante
 NL : Bloedmonster van heterozygote patiënt met Hb Lepore-Boston-Washington variant
 IT : Sangue di paziente eterozygote con variante Hb Lepore-Boston-Washington
 ES : Sangre de paciente heterocigoto con la variante Hb Lepore-Boston-Washington
 PT : Amostra de sangue de doente heterozigótico com variante Hb Lepore-Boston-Washington
 SV : Blodprov från heterozygot patient med Hb Lepore-Boston-Washington-variant
 GR : Δείγμα αίματος από ετερόζυγο ασθενή με παραλλαγή Hb Lepore-Boston-Washington
 HR : Uzorak krvi heterozigotnog bolesnika s varijantom Hb Lepore-Boston-Washington
 LT : Heterozigotinio paciento kraujo, kuriame yra Hb Lepore-Boston-Washington variantu, kraujo mėginys
 PL : Próbk krwi od heterozygotycznego pacjenta z odmianą Hb Lepore-Boston-Washington
 RO : Probă de sânge de la pacient heterozigot cu varianta Hb Lepore-Boston-Washington
 CS : Uzorak krvi od heterozigotnog pacijenta sa Hb Lepore-Boston-Washington varijantom
 HU : Heterozigóta beteg vérmintája Hb Lepore-Boston-Washington variánsal
 TR : Hb Lepore-Boston-Washington varyanti taşıyan heterozigot hastasına ait kan numunesi
 CZ : Vzorek krve heterozygotního pacienta s variantou Hb Lepore-Boston-Washington
 BG : Кръвна проба от хетерозимотен пациент с Hb Lepore-Boston-Washington екарием
 NO : Blodprøve fra heterozygot pasient med Hb Lepore-Boston-Washington variant
 DK : Blodprøve fra heterozygot patient med Hb Lepore-Boston-Washington-variant
 CN : 杂合 Hb Lepore-Boston-Washington 变异杂合患者的血液样品
 RU : Образец крови от гетерозиготного пациента с вариацией Hb Lepore-Boston-Washington
 JP : Hb Lepore-Boston-Washington 変異体を含むヘテロ変異患者からの血液サンプル
 LV : Heterozigota pacienta asins paraugs ar Hb Lepore-Boston-Washington variantu
 SK : Vzorka krvi od heterozigotného pacienta s variantom Hb Lepore-Boston-Washington
 EE : Vereproov Hb Lepore-Boston-Washingtoni variantiga heterosügotselt patsiendilt
 VN : Mẫu máu của bệnh nhân bị bệnh dị hợp tử với biến thể Hb Lepore-Boston-Washington



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