SIEMENS Healthineers

Alanine Aminotransferase P5P (ALTPLc)

Current Revision and Date[a]	Rev. 02, 2019-11	
Product Name	Atellica CH Alanine Aminotransferase P5P (ALTPLc)	REF 11097638 (2080 tests)
Abbreviated Product Name	Atellica CH ALTPLc	
Test Name/ID	ALTPLc	
Systems	Atellica CH Analyzer	
Materials Required but Not Provided	Atellica CH ENZ 2 CAL	REF 11099318
Specimen Types	Serum and plasma (lithium heparin)	
Sample Volume	25 µL	
Measuring Interval	9–1000 U/L	

[a] A vertical bar in the page margin indicates technical content that differs from the previous version.

CE

Intended Use

The Atellica® CH Alanine Aminotransferase P5P (ALTPLc) assay is for *in vitro* diagnostic use in the quantitative determination of alanine aminotransferase activity in human serum and plasma (lithium heparin) on the Atellica® CH Analyzer. This assay is used to aid in the diagnosis and treatment of certain types of liver disease.

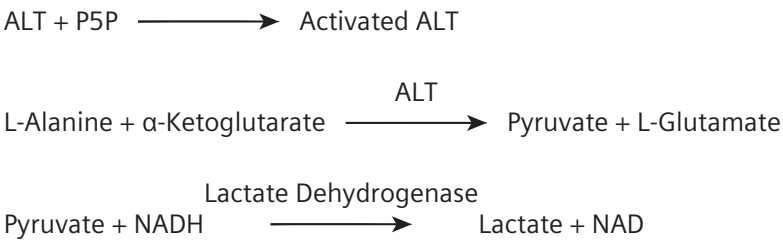
Summary and Explanation

The Atellica CH Alanine Aminotransferase P5P (ALTPLc) assay uses pyridoxal-5'-phosphate (P5P_L) added to Reagent 1 of the Atellica CH ALTPLc assay. The assay is based on work by Wroblewski and LaDue, which was later modified by H.U. Bergmeyer. The Atellica CH ALTPLc reagents are formulated in accordance with the IFCC recommendations.[1]

Principles of the Procedure

The reaction is initiated by the addition of α-ketoglutarate as a second reagent. The concentration of NADH is measured by its absorbance at 340/410 nm and the rate of absorbance decrease is proportional to the alanine aminotransferase activity.

Reaction Equation



Reagents

Material Description	Storage	Stability[a]
Atellica CH ALTPLc	Unopened at 2–8°C	Until expiration date on product
Pack 1 (P1)	Onboard per pack	19 days
Well 1 (W1) Reagent 1 (R1) 22.8 mL L-Alanine (1.22 mol/L); LD (pig heart) (≥ 2.4 kU/L); sodium azide (0.09%)		
Well 2 (W2) Reagent 1 (R1) 22.8 mL L-Alanine (1.22 mol/L); LD (pig heart) (≥ 2.4 kU/L); sodium azide (0.09%)		
Pack 2 (P2)		
Well 1 (W1) Reagent 2 (R2) 16.2 mL α-Ketoglutarate (93 mmol/L); reduced NADH (1.41 mmol/L); sodium azide (0.09%)		
Well 2 (W2) Reagent 2 (R2) 16.2 mL α-Ketoglutarate (93 mmol/L); reduced NADH (1.41 mmol/L); sodium azide (0.09%)		
Vial (P5P_L)	Unopened at 2–8°C	Until expiration date on product
0.475 mL Pyridoxal-5'-Phosphate (P5P) (30 mmol/L) sodium azide (0.09%)		

[a] Refer to *Storage and Stability*.

Warnings and Precautions

For *in vitro* diagnostic use.

For Professional Use.

CAUTION

Federal (USA) law restricts this device to sale by or on the order of a licensed healthcare professional.

Safety data sheets (SDS) available on siemens.com/healthineers.

CAUTION

This device contains material of animal origin and should be handled as a potential carrier and transmitter of disease.

Contains sodium azide as a preservative. Sodium azide can react with copper or lead plumbing to form explosive metal azides. On disposal, flush reagents with a large volume of water to prevent buildup of azides. Disposal into drain systems must be in compliance with prevailing regulatory requirements.

Dispose of hazardous or biologically contaminated materials according to the practices of your institution. Discard all materials in a safe and acceptable manner and in compliance with prevailing regulatory requirements.

Note For information about reagent preparation, refer to *Preparing the Reagents* in the *Procedure* section.

Storage and Stability

Unopened reagents are stable until the expiration date on the product when stored at 2–8°C.

Do not use products beyond the expiration date printed on the product labeling.

Onboard Stability

Reagents are stable onboard the system for 19 days per pack. Discard reagents at the end of the onboard stability interval. Do not use products beyond the expiration date printed on the product labeling.

Specimen Collection and Handling

Serum and plasma (lithium heparin) are the recommended sample types for this assay.

Collecting the Specimen

- Observe universal precautions when collecting specimens. Handle all specimens as if they are capable of transmitting disease.[2]
- Follow recommended procedures for collection of diagnostic blood specimens by venipuncture.[3]
- Follow the instructions provided with your specimen collection device for use and processing.[4]
- Allow blood specimens to clot completely before centrifugation.[5]
- Keep tubes capped at all times.[5]

Storing the Specimen

Separated specimens may be stored for up to 7 days at 2–8°C or stored frozen for up to 30 days at -20°C. Avoid repeated freezing and thawing.[6]

The handling and storage information provided here is based on data or references maintained by the manufacturer. It is the responsibility of the individual laboratory to use all available references and/or its own studies when establishing alternate stability criteria to meet specific needs.

Transporting the Specimen

Package and label specimens for shipment in compliance with applicable federal and international regulations covering the transport of clinical specimens and etiological agents.

Preparing the Samples

This assay requires 25 µL of sample for a single determination. This volume does not include the unusable volume in the sample container or the additional volume required when performing duplicates or other tests on the same sample. For information about determining the minimum required volume, refer to the online help.

Note Do not use specimens with apparent contamination.

Before placing samples on the system, ensure that samples are free of:

- Bubbles or foam.
- Fibrin or other particulate matter.

Note Remove particulates by centrifugation according to CLSI guidance and the collection device manufacturer's recommendations.[5]

Note For a complete list of appropriate sample containers, refer to the online help.

Procedure

Materials Provided

The following materials are provided:

REF	Contents	Number of Tests
11097638	**Pack 1 (P1)** Well 1 (W1) 22.8 mL of Atellica CH ALTPLc Reagent 1 Well 2 (W2) 22.8 mL of Atellica CH ALTPLc Reagent 1 **Pack 2 (P2)** Well 1 (W1) 16.2 mL of Atellica CH ALTPLc Reagent 2 Well 2 (W2) 16.2 mL of Atellica CH ALTPLc Reagent 2 **Vial (P5P_L)** 4 x Vial (P5P_L) 0.475 mL of Atellica CH P5P_L	2 x 1040

Materials Required but Not Provided

The following materials are required to perform this assay, but are not provided:

REF	Description	
	Atellica CH Analyzer[a]	
11099318	Atellica CH ENZ 2 CAL (calibrator)	6 x 1.5 mL calibrator CAL Calibrator lot-specific value sheet CAL LOT VAL
	Commercially available quality control materials	

[a] Additional system fluids are required to operate the system: Atellica CH Diluent, Atellica CH Wash, Atellica CH Conditioner, Atellica CH Cleaner, Atellica CH Reagent Probe Cleaner 1, Atellica CH Reagent Probe Cleaner 2, Atellica CH Reagent Probe Cleaner 4, Atellica CH Lamp Coolant, and Atellica CH Water Bath Additive. For system fluid instructions for use, refer to the Document Library.

Assay Procedure

The system automatically performs the following steps:

1. For serum/plasma, dispenses 50 µL of primary sample and 200 µL of Atellica CH Diluent into a dilution cuvette.
2. Dispenses 40 µL of Reagent 1 and 40 µL of special reagent water into a reaction cuvette.
3. Dispenses 25 µL of pre-diluted sample into a reaction cuvette.
4. Measures the absorbance after sample addition.
5. Dispenses 16 µL of Reagent 2 into a reaction cuvette.
6. Mixes and incubates the mixture at 37°C.
7. Measures the absorbance rate after Reagent 2 addition.
8. Reports results.

Note For information about special reagent water requirements, refer to the online help.

Test Duration: 10 minutes

Preparing the Reagents

The Atellica CH P5P_L reagent is supplied ready for use, it may be added directly into the Atellica CH ALTPLc R1 reagent:

1. Carefully open one vial of the Atellica CH P5P_L and well 1 of the Atellica CH ALTPLc P1 pack.
2. The P5P_L may be poured or pippetted into well 1 of the ALTPLc P1 pack.
   - While holding the P1 pack steady on the bench, carefully pour the contents of the P5P_L vial into well 1 of the ALTPLc P1 pack.

     Note 300 µL of P5P_L will be added to the P1 pack

     Note Do not shake the excess P5P_L out of the vial.

     or
   - Using a volumetric pipette, withdraw 300 µL of P5P_L from the vial and carefully dispense it into well 1 of the ALTPLc P1 pack.
3. Carefuly close well 1 of the ALTPLc P1 pack
4. To ensure homogeneity, mix the contents by gently swirling the P1 pack.

   Note Avoid formation of foam.

   Note Do not shake.
5. Repeat steps 1–4 for well 2 of the P1 pack with a fresh P5P_L vial.

The ALTPLc R2 reagent is liquid and ready to use.

Preparing the System

Ensure that the system has sufficient reagent packs loaded in the reagent compartment. For information about loading reagent packs, refer to the online help.

Performing Calibration

For calibration of the Atellica CH ALTPLc assay, use Atellica CH ENZ 2 CAL. Use the calibrators in accordance with the calibrator instructions for use.

Calibration Frequency

Perform a calibration if one or more of the following conditions exist:

- When changing lot numbers of primary reagent packs.
- At the end of the lot calibration interval, for a specified lot of calibrated reagent on the system.
- At the end of the pack calibration interval, for calibrated reagent packs on the system.
- When indicated by quality control results.
- After major maintenance or service, if indicated by quality control results.

At the end of the onboard stability interval, replace the reagent pack on the system with a new reagent pack. Recalibration is not required, unless the lot calibration interval is exceeded.

Stability Interval	Days
Lot Calibration	130
Pack Calibration	4
Reagent Onboard Stability	19

For information about lot calibration and pack calibration intervals, refer to the online help.

Follow government regulations or accreditation requirements for calibration frequency. Individual laboratory quality control programs and procedures may require more frequent calibration.

Performing Quality Control

For quality control of the Atellica CH ALTPLc assay, use at least two levels (low and high) of the appropriate quality control material of known analyte concentration. Use the quality control material in accordance with the quality control instructions for use.

For the assigned values, refer to the lot-specific value sheet provided. A satisfactory level of performance is achieved when the analyte values obtained are within the expected control range for the system or within your range, as determined by an appropriate internal laboratory quality control scheme. Follow your laboratory's quality control procedures if the results obtained do not fall within the acceptable limits. For information about entering quality control definitions, refer to the online help.

Follow government regulations or accreditation requirements for quality control frequency. Individual laboratory quality control programs and procedures may require more frequent quality control testing.

Taking Corrective Action

If the quality control results do not fall within the assigned values, do not report results. Perform corrective actions in accordance with established laboratory protocol. For suggested protocol, refer to the online help.

Results

Calculation of Results

The system determines the result using the calculation scheme described in the online help. The system reports results in U/L.

For information about results outside the specified measuring interval, refer to *Measuring Interval*.

Interpretation of Results

Results of this assay should always be interpreted in conjunction with the patient's medical history, clinical presentation, and other findings.

Limitations

The Atellica CH ALTPLc assay is limited to the detection of alanine aminotransferase in human serum and plasma (lithium heparin).

Sulfasalazine concentrations in serum or plasma above 50 mg/L may cause falsely depressed results with the Atellica CH ALTPLc assay.[7]

Expected Values

Reference Interval

A reference interval for healthy adults was established in accordance with CLSI Document EP28-A3c and verified on the Atellica CH Analyzer.[8]

The reference interval for alanine aminotransferase is 7–40 U/L for adults.[9]

As with all *in vitro* diagnostic assays, each laboratory should determine its own reference interval for the diagnostic evaluation of patient results. Consider these values as guidance only.[8]

Performance Characteristics

Measuring Interval

The Atellica CH ALTPLc assay provides results from 9–1000 U/L. The system flags all values that are outside the specified measuring interval.

Extended Measuring Interval

An automatic repeat condition for this assay extends the measuring interval to 6000 U/L for serum and plasma. You may configure the system to trigger an automatic repeat. Automatic repeat results will be flagged **Autorepeat**.

Detection Capability

Detection capability was determined in accordance with CLSI Document EP17-A2.[10] The assay is designed to have a limit of blank (LoB) ≤ 8 U/L and limit of detection (LoD) ≤ 8 U/L.

The LoD corresponds to the lowest concentration of alanine aminotransferase that can be detected with a probability of 95%. The LoD for the Atellica CH ALTPLc assay is 3 U/L and the LoB is 2 U/L; and they were determined using 180 determinations, with 60 blank and 60 low level replicates.

The Limit of Quantitation (LoQ) corresponds to the lowest amount of analyte in a sample at which the total error is ≤ 8 U/L. For serum and plasma, the assay is designed to have and LoQ ≤ 16 U/L. The LoQ for the Atellica CH ALTPLc assay is 5 U/L based on 180 determinations.

Assay results obtained at individual laboratories may vary from the data presented.

Precision

Precision was determined in accordance with CLSI Document EP05-A3.[11] The assay was designed to have serum/plasma Within-Lab precision of ≤ 7.0 %CV from 30–100 U/L, and ≤ 5.0 %CV from 101–1000 U/L alanine aminotransferase. Samples were assayed on an Atellica CH Analyzer in duplicate in 2 runs per day for 20 days (N ≥ 80 for each sample). The following results were obtained:

			Repeatability		Within-Lab Precision	
Sample Type	N	Mean U/L	SD[a] U/L	CV[b] (%)	SD U/L	CV (%)
Serum QC	80	36	0.7	2.0	1.0	2.8
Serum QC	80	102	1.8	1.8	2.7	2.7
Serum QC	80	218	1.4	0.6	2.9	1.3
Plasma	80	541	1.8	0.3	3.8	0.7
Serum	80	898	3.2	0.4	8.9	1.0

[a] Standard deviation.
[b] Coefficient of variation.

Assay results obtained at individual laboratories may vary from the data presented.

Assay Comparison

The Atellica CH ALTPLc assay is designed to have a correlation coefficient of ≥ 0.90 and a slope of 1.00 ± 0.10 compared to ADVIA® Chemistry 1800 ALTPLc. Assay comparison was determined using the Deming linear regression model in accordance with CLSI Document EP09-A3.[12] The following results were obtained:

Specimen	Comparative Assay (x)	Regression Equation	Sample Interval	N[a]	r[b]
Serum	ADVIA Chemistry 1800 ALTPLc	y = 1.02x - 0 U/L	10–995 U/L	103	0.999

[a] Number of samples tested.
[b] Correlation coefficient.

The agreement of the assay may vary depending on the study design, comparative assay, and sample population. Assay results obtained at individual laboratories may vary from the data presented.

Specimen Equivalency

Specimen equivalency was determined using the Deming linear regression model in accordance with CLSI Document EP09-A3.[12] The following results were obtained:

Specimen (y)	Reference Specimen (x)	Regression Equation	Sample Interval	N[a]	r[b]
Lithium heparin plasma	Serum	y = 0.96x + 0 U/L	9–936 U/L	56	0.999

[a] Number of samples tested.
[b] Correlation coefficient.

Agreement of the specimen types may vary depending on the study design and sample population used. Assay results obtained at individual laboratories may vary from the data presented.

Interferences

Hemolysis, Icterus, and Lipemia (HIL)

The Atellica CH ALTPLc assay is designed to have ≤ 10% interference from hemoglobin, bilirubin, and lipemia. Interfering substances at the levels indicated in the table below were tested in accordance with CLSI Document EP07-A2 using the Atellica CH ALTPLc assay.[13]

Bias is the difference in the results between the control sample (does not contain the interferent) and the test sample (contains the interferent) expressed in percent. Bias > 10% is considered interference. Analyte results should not be corrected based on this bias.

Substance	Substance Test Concentration Common Units (SI Units)	Analyte Concentration U/L	Percent Bias
Hemoglobin	200 mg/dL (0.12 mmol/L)	54	9
	200 mg/dL (0.12 mmol/L)	232	2
Bilirubin, conjugated	20 mg/dL (342 µmol/L)	49	4
	20 mg/dL (342 µmol/L)	273	-3
Bilirubin, unconjugated	20 mg/dL (342 µmol/L)	52	-2
	20 mg/dL (342 µmol/L)	279	-1
Lipemia (Intralipid®)[a]	400 mg/dL 4.5 mmol/L)	53	0
	400 mg/dL (4.5 mmol/L)	282	-2

[a] Percent bias based on Intralipid concentrations of 500 mg/dL

Assay results obtained at individual laboratories may vary from the data presented.

Non-Interfering Substances

The following substances do not interfere with the Atellica CH ALTPLc assay when present in serum and plasma at the concentrations indicated in the table below. Bias due to these substances is ≤ 10% at analyte concentrations indicated in the table below. These data were generated on the ADVIA Chemistry 1800 system with assay reaction conditions that are equivalent to those on the Atellica CH Analyzer.[7]

Substance	Substance Test Concentration Common Units (SI Units)	Analyte Concentration	Percent Bias
Sulfasalazine	50 mg/L (126 µmol/L)	65 U/L	≤ 10
	50 mg/L (126 µmol/L)	297 U/L	≤ 10
Sulfapyridine	299 mg/L (1.2 mmol/L)	62 U/L	≤ 10
	299 mg/L (1.2 mmol/L)	283 U/L	≤ 10

Assay results obtained at individual laboratories may vary from the data presented.

Standardization

The Atellica CH ALTPLc assay is traceable to IFCC reference method.

Assigned values for calibrators are traceable to this standardization.[7]

Technical Assistance

For customer support, contact your local technical support provider or distributor.

siemens.com/healthineers

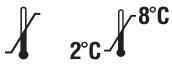
References

1. International Federal of Clinical Chemistry and Laboratory Medicine. IFCC primary reference procedures for the measurement of catalytic activity concentrations of enzymes at 37 degrees C. *Clin Chem Lab Med*. 2002;40(7):718–724.
2. Clinical and Laboratory Standards Institute. *Protection of Laboratory Workers from Occupationally Acquired Infections; Approved Guideline—Fourth Edition*. Wayne, PA: Clinical and Laboratory Standards Institute; 2014. CLSI Document M29-A4.
3. Clinical and Laboratory Standards Institute. *Procedures for the Collection of Diagnostic Blood Specimens by Venipuncture; Approved Standard—Sixth Edition*. Wayne, PA: Clinical and Laboratory Standards Institute; 2007. CLSI Document GP41-A6.
4. Clinical and Laboratory Standards Institute. *Tubes and Additives for Venous and Capillary Blood Specimen Collection; Approved Standard—Sixth Edition*. Wayne, PA: Clinical and Laboratory Standards Institute; 2010. CLSI Document GP39-A6.
5. Clinical and Laboratory Standards Institute. *Procedures for the Handling and Processing of Blood Specimens for Common Laboratory Tests; Approved Guideline—Fourth Edition*. Wayne, PA: Clinical and Laboratory Standards Institute; 2010. CLSI Document GP44-A4.
6. Young DS. *Effects of Preanalytical Variables on Clinical Laboratory Tests*. 2nd ed. Washington, DC: AACC Press; 1997:3-10–3-12.
7. Data on file at Siemens Healthcare Diagnostics.
8. Clinical and Laboratory Standards Institute. *Defining, Establishing, and Verifying Reference Intervals in the Clinical Laboratory; Approved Guideline—Third Edition*. Wayne, PA: Clinical and Laboratory Standards Institute; 2010. CLSI Document EP28-A3c.
9. Tietz NW. *Clinical Guide to Laboratory Tests*. 3rd ed. Philadelphia, PA: Saunders; 1995.
10. Clinical and Laboratory Standards Institute. *Evaluation of Detection Capability for Clinical Laboratory Measurement Procedures; Approved Guideline—Second Edition*. Wayne, PA: Clinical and Laboratory Standards Institute; 2012. CLSI Document EP17-A2.
11. Clinical and Laboratory Standards Institute. *Evaluation of Precision of Quantitative Measurement Procedures; Approved Guideline—Third Edition*. Wayne, PA: Clinical and Laboratory Standards Institute; 2014. CLSI Document EP05-A3.
12. Clinical and Laboratory Standards Institute. *Measurement Procedure Comparison and Bias Estimation Using Patient Samples; Approved Guideline—Third Edition*. Wayne, PA: Clinical and Laboratory Standards Institute; 2013. CLSI Document EP09-A3.
13. Clinical and Laboratory Standards Institute. *Interference Testing in Clinical Chemistry; Approved Guideline—Second Edition*. Wayne, PA: Clinical and Laboratory Standards Institute; 2005. CLSI Document EP07-A2.

Definition of Symbols

The following symbols may appear on the product labeling:

Symbol	Symbol Title and Description
	Consult instructions for use
	Version of instructions for use
	Internet URL address to access the electronic instructions for use
	Revision
	Caution Consult instructions for use or accompanying documents for cautionary information such as warnings and precautions that cannot, for a variety of reasons, be presented on the medical device.
	Biological risks Potential biological risks are associated with the medical device.
	Corrosive
	Dangerous to environment
	Irritant Oral, dermal, or inhalation hazard
	Inhalation hazard Respiratory or internal health
	Flammable Flammable to extremely flammable
	Oxidizing
	Explosive
	Toxic
	Compressed gas
	Keep away from sunlight Prevent exposure to sunlight and heat.

Symbol	Symbol Title and Description
	Up Store in an upright position.
	Do not freeze
	Temperature limit Upper and lower limits of temperature indicators are adjacent to the upper and lower horizontal lines.
	Handheld barcode scanner
	*In vitro* diagnostic medical device
	Contains sufficient for <n> tests Total number of IVD tests the system can perform with the IVD kit reagents appears adjacent to the symbol.
RxOnly	Prescription device (US only) Applies only to United States-registered IVD assays. CAUTION: Federal (USA) law restricts this device to sale by or on the order of a licensed healthcare professional.
	Mixing of substances Mix product before use.
	Reconstitute and mix lyophilized product before use.
	Target
	Interval
	Legal Manufacturer
	Authorized Representative in the European Community
	Use-by date Use by the designated date.
	Batch code
	Catalog number
	Recycle
	Printed with soy ink
	CE Mark

Symbol	Symbol Title and Description
CE 0088	CE Mark with notified body ID number Notified body ID number can vary.
YYYY-MM-DD	Date format (year-month-day)
CHECKSUM	Variable hexadecimal number that ensures the Master Curve and Calibrator definition values entered are valid.
UNITS C	Common Units
UNITS SI	International System of Units
MATERIAL	Material
MATERIAL ID	Unique material identification number
CONTROL NAME	Name of control
CONTROL TYPE	Type of control

Legal Information

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