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As of 17 Mar. 2010 rm (Vers. 1.0)

Please use only the valid version of the package insert provided with the kit.

1 INTENDED USE

The Human FGF-19 ELISA is a sandwich enzyme immunoassay for the quantitative measurement of human fibroblast growth factor-19 (FGF-19).

Features

- It is intended for research use only
- The total assay time is less than 3.5 hours
- The kit measures FGF-19 in serum and plasma (EDTA, citrate, heparin)
- Assay format 96 wells
- Standard and Quality Controls are recombinant protein based
- Components of the kit are provided ready to use, concentrated or lyophilized

2 STORAGE, EXPIRATION

Store the complete kit at 2-8°C. Under this condition, the kit is stable until the expiration date (see the label on the box).

For stability of opened components see Chapter 9.

3 INTRODUCTION

Fibroblast growth factors (FGFs) are a large family of small (17-26 kDa) polypeptide growth factors found in organisms ranging from nematodes to humans. The FGF family has at least 22 members in vertebrates and share 13-71% amino acid identity. The initial characterization of these proteins focused on their ability to stimulate fibroblast proliferation. During embryonic development, FGFs have diverse roles in regulating cell proliferation, migration and differentiation. In the adult organism, FGFs are homeostatic factors and function in tissue repair and response to injury.

FGF signaling is mediated through one of four FGF receptors (FGFR1-FGFR4), a complex family of transmembrane receptor tyrosine kinases. FGFR5 has also been described, but lacks the kinase domain and signaling capability. FGFs have a high affinity for heparin sulfate proteoglycans and require heparin sulfate to activate FGF receptors. Although multiple FGFs interact with each of the four FGFRs, a novel fibroblast growth factor FGF-19 exhibits exclusive binding to only one of FGF receptors (FGFR4).

The normal function of FGF-19 has not been resolved, although its role in inner ear development has been suggested. It has been also found that hepatocyte expression of FGF-19 is induced by the transcription factor, farnesoid X receptor (FXR). FXR is a key regulator of cholesterol metabolism through suppression of the catabolic enzyme cyp7a, the first and rate-limiting step in the biosynthesis of bile acids (BA). A recent study found that, in humans, circulating FGF-19 has a diurnal rhythm controlled by the transintestinal BA flux, and that FGF-19 modulates hepatic BA synthesis. Through its systemic effects, circulating FGF-19 may also mediate other known BA-dependent effects on lipid and carbohydrate metabolism.





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In transgenic mice expressing human FGF-19, researchers found a significant increased metabolic rate as well as decreased body weight and adiposity. Additionally, resistance to both diet-induced obesity and insulin desensitization were found. Similar responses have been observed when recombinant FGF-19 was injected into the mice. However, it has been shown too, that FGF-19 transgenic mice develop hepatic adrenocarcinomas with age, and recombinant FGF-19 treated mice exhibit proliferation of hepatocytes.

Areas of investigation:

Cholesterol metabolism

Metabolic syndrome

4 TEST PRINCIPLE

In the Human FGF-19 ELISA, Standards, Quality Controls and samples are incubated in microtitration wells pre-coated with polyclonal anti-human FGF-19 antibody. After a 60 minute incubation followed by washing, biotin labelled polyclonal anti-human FGF-19 antibody is added and incubated with the captured FGF-19 for 60 minutes. After another washing, streptavidin-HRP conjugate is added. After 30 minutes incubation and the last washing step, the remaining conjugate is allowed to react with the substrate solution (TMB). The reaction is stopped by addition of acidic solution (0.2M H₂SO₄) and absorbance of the resulting yellow product is measured. The absorbance is proportional to the concentration of FGF-19. A standard curve is constructed by plotting absorbance values versus FGF-19 concentrations of Standards, and concentrations of unknown samples are determined using this standard curve.

5 PRECAUTION

For professional use only.

Wear gloves and laboratory coats when handling immunodiagnostic materials.

Do not drink, eat or smoke in the areas where immunodiagnostic materials are being handled.

This kit contains components of human origin. These materials were found non-reactive for hepatitis B surface antigen and for HIV antibody. However, these materials should be handled as potentially infectious, as no test can guarantee the complete absence of infectious agents.

Avoid contact with the acidic Stop Solution and Substrate (TMB) Solution, which contains hydrogen peroxide. Wear gloves and eye protection when handling these reagents. Stop and/or Substrate Solutions may cause skin/eyes irritation. In case of contact with the Stop Solution or the Substrate Solution, wash skin/eyes thoroughly with water and seek medical attention when necessary.

The materials must not be pipetted by mouth.

6 TECHNICAL HINTS

Reagents with different lot numbers should not be mixed.

Use thoroughly clean glassware.

Use deionized (distilled) water, stored in clean containers.

Avoid any contamination among samples and reagents. For this purpose, disposable tips should be used for each sample and reagent.

Substrate Solution should remain colourless until added to the plate. Keep Substrate Solution protected from light.







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Stop Solution should remain colourless until added to the plate. The colour developed in the wells will turn from blue to yellow immediately after the addition of the Stop Solution. Wells that are green in colour indicate that the Stop Solution has not mixed thoroughly with the Substrate Solution.

Dispose of consumable materials and unused contents in accordance with applicable national regulatory requirements.

7 REAGENT SUPPLIED

Kit Components	State	Quantity
Antibody Coated Microtiter Strips	ready to use	96 wells
Biotin Labelled Antibody Conc. (50x)	concentrated	0.3 ml
Streptavidin-HRP Conjugate	ready to use	13 ml
Master Standard	lyophilized	2 vials
Quality Control HIGH	lyophilized	2 vials
Quality Control LOW	lyophilized	2 vials
Dilution Buffer	ready to use	20 ml
Biotin-Ab Diluent	ready to use	13 ml
Wash Solution Conc. (10x)	concentrated	100 ml
Substrate Solution	ready to use	13 ml
Stop Solution	ready to use	13 ml
Product Data Sheet + Certificate of Analysis	-	1 pc

8 MATERIAL REQUIRED BUT NOT SUPPLIED

Deionized (distilled) water

Test tubes for samples dilution

Glassware (graduated cylinder and bottle) for Wash Solution

Absorbent material (e.g. paper towels) for blotting the microtiter plate after washing

Vortex mixer

Precision pipettes to deliver 10-1000 µl with disposable tips

Multichannel pipette to deliver 100 µl with disposable tips

Orbital microplate shaker capable of approximately 300 rpm

Microplate washer (optional). Manual washing is possible but not preferable

Microplate reader with $450 \pm \Box 10$ nm filter, preferably with reference wavelength 630 nm (alternatively another one from the interval 550-650nm)

Software package facilitating data generation and analysis (optional)







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9 PREPARATION OF REAGENTS

- All reagents need to be brought to room temperature prior to use.
- Always prepare only the appropriate quantity of reagents for your test.
- Do not use components after the expiration date marked on their label.

9.1 Assay reagents supplied ready-to-use:

Antibody Coated Microtiter Strips

Stability and storage:

Return the unused strips to the provided aluminium zip-sealed bag with desiccant and seal carefully. Remaining Microtiter Strips are stable 3 months when stored at 2-8°C and protected from the moisture.

Dilution Buffer
Biotin-Ab Diluent
Streptavidin-HRP Conjugate
Substrate Solution
Stop Solution
Stability and storage:

Opened reagents are stable 3 month when stored at 2-8°C.

9.2 Assay reagents supplied concentrated or lyophilized:

Human FGF-19 Master Standard

Refer to Certificate of Analysis for current volume of Dilution Buffer needed for reconstitution of standard!!!

Reconstitute the lyophilized Master Standard with Dilution Buffer just prior to the assay. Let it dissolve at least 15 minutes with occasionally gently shaking (not to foam). The resulting concentration of the human FGF-19 in the stock solution is 800 pg/ml.

Prepare set of standards using Dilution Buffer as follows:

Volume of Standard	Dilution Buffer	Concentration
Stock		800 pg/ml
300 µl of stock	300 μΙ	400 ng/ml
300 µl of 400 pg/ml	300 μΙ	200 ng/ml
300 µl of 200 pg/ml	300 μΙ	100 ng/ml
300 µl of 100 pg/ml	300 μΙ	50 ng/ml
300 µl of 50 pg/ml	300 μΙ	25 ng/ml
300 µl of 25 pg/ml	300 μΙ	12.5 ng/ml





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Prepared Standards are ready to use, do not dilute them.

Stability and storage:

Do not store the standard stock solution and the set of standards.

Quality Controls High, Low

Refer to the Certificate of Analysis for current volume of Dilution Buffer needed for reconstitution and for current Quality Controls concentrations!!!

Reconstitute each Quality Control (High and Low) with Dilution Buffer just prior to the assay. Let it dissolve at least 15 minutes occasionally gently shaking (not to foam).

The reconstituted Quality Controls are ready to use, do not dilute them.

Stability and storage:

Do not store the reconstituted Quality Controls.

Biotin Labelled Antibody Concentrate (50x)

Prepare the working Biotin Labelled Antibody solution by adding 1 part concentrated Biotin Labelled Antibody Concentrate (50x) with 49 parts Biotin-Ab Diluent.

Example: 20 µl of Biotin Labelled Antibody Concentrate (50x) + 980 µl of Biotin-Ab Diluent for 1 strip (8 wells).

Stability and storage:

Opened Biotin Labelled Antibody Concentrate (50x) is stable 3 months when stored at 2-8°C.

Do not store the diluted Biotin Labelled Antibody solution.

Wash Solution Concentrate (10x)

Dilute Wash Solution Concentrate (10x) ten-fold in distilled water to prepare a 1x working solution.

Example: 100 ml of Wash Solution Concentrate (10x) + 900 ml of distilled water for use of all 96-wells.

Stability and storage:

The diluted Wash Solution is stable 1 month when stored at 2-8°C. Opened Wash Solution Concentrate (10x) is stable 3 months when stored at 2-8°C.

10 PREPARATION OF SAMPLES

The kit measures human FGF-19 in serum and plasma.

Samples should be assayed immediately after collection or should be stored at -20°C or - 70°C. Thoroughly mix thawed samples just prior to the assay and avoid repeated freeze/thaw cycles, which may cause erroneous results. Avoid using hemolyzed or lipemic samples.





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Preparation of samples:

Dilute samples 3x with Dilution Buffer just prior to the assay, e.g. $50 \mu l$ of sample $+ 100 \mu l$ of Dilution Buffer for singlets or $80 \mu l$ of sample $+ 160 \mu l$ of Dilution Buffer for duplicates. **Mix well** (not to foam). Vortex is recommended.

Stability and storage:

Serum samples should be stored at -20°C, or preferably at -70°C for long-term storage. Avoid repeated freeze/ thaw cycles.

Do not store the diluted samples.

See Chapter 15 for stability of serum and plasma samples if stored at 2-8°C, effect of freezing/thawing and effect of sample matrix (serum/plasma) on the concentration of FGF-19.

Note: It is recommended to use a precision pipette and a careful technique to perform the dilution in order to get precise results!

11 ASSAY PROCEDURE

- 1. Pipet **100 μl** of Standards reconstituted Quality Control and diluted samples, preferably in duplicates, into the appropriate wells. See Figure 1 for example of work sheet.
- 2. Incubate the plate at room temperature (ca. 25°C) for **1 hour**, shaking at ca. 300 rpm on an orbital microplate shaker.
- 3. Wash the wells 3-times with Wash Solution (0.35 ml per well). After final wash, invert and tap the plate strongly against paper towel. This removes all liquid from wells.
- 4. Pipet 100 μl of Biotin Labelled Antibody Solution into each well.
- 5. Incubate the plate at room temperature (ca. 25°C) for 1 hour, shaking at ca. 300 rpm on an orbital microplate shaker.
- 6. Wash the wells 3-times with Wash Solution (0.35 ml per well). After final wash, invert and tap the plate strongly against paper towel.
- 7. Pipet 100 μl of Streptavidin-HRP Conjugate into each well.
- 8. Incubate the plate at room temperature (ca. 25°C) for **30 minutes**, shaking at ca. 300 rpm on an orbital microplate shaker.
- 9. Wash the wells 3-times with Wash Solution (0.35 ml per well). After final wash, invert and tap the plate strongly against paper towel.
- 10. Add **100** μl of Substrate Solution. (Avoid exposing the microtiter plate to direct sunlight. Covering the plate with e.g. aluminium foil is recommended.)
- 11. Incubate the plate for **10 minutes** at room temperature. (The incubation time may be extended [up to 20 minutes] if the reaction temperature is below than 20°C). No shaking!
- 12. Stop the colour development by adding 100 μl of Stop Solution.
- 13. Determine the absorbance of each well using a microplate reader set to 450 nm, preferably with the reference wavelength set to 630 nm (acceptable range: 550–650 nm). Subtract readings at 630 nm (550-650 nm) from the readings at 450 nm. The absorbance should be read within 5 minutes following step 12.







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Note: If some samples and standard/s have absorbances above the upper limit of your microplate reader, perform a second reading at 405 nm. A new standard curve, constructed using the values measured at 405 nm, is used to determine FGF-19 concentration of off-scale standards and samples. The readings at 405 nm should not replace the readings for samples that were "in range" at 450 nm.

Note 2: Manual washing: Aspirate wells and pipet 0.35 ml Wash Solution into each well. Aspirate wells and repeat twice. After final wash, invert and tap the plate strongly against paper towel. Make certain that Wash Solution has been removed entirely.

	strip 1+2	strip 3+4	strip 5+6	strip 7+8	strip 9+10	strip 11+12
Α	Standard 800	QC High	Sample 7	Sample 15	Sample 23	Sample 31
В	Standard 400	QC Low	Sample 8	Sample 16	Sample 24	Sample 32
С	Standard 200	Sample 1	Sample 9	Sample 17	Sample 25	Sample 33
D	Standard 100	Sample 2	Sample 10	Sample 18	Sample 26	Sample 34
E	Standard 50	Sample 3	Sample 11	Sample 19	Sample 27	Sample 35
F	Standard 25	Sample 4	Sample 12	Sample 20	Sample 28	Sample 36
G	Standard 12.5	Sample 5	Sample 13	Sample 21	Sample 29	Sample 37
Н	Blank	Sample 6	Sample 14	Sample 22	Sample 30	Sample 38

Figure 1: Example of a work sheet.



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12 CALCULATIONS

Most microtiter plate readers perform automatic calculations of analyte concentration. The Standards curve is constructed by plotting the absorbance (Y) of Standards against the known concentration (X) of Standards in logarithmic scale, using the four-parameter algorithm. Results are reported as concentration of FGF-19 (pg/ml) in samples.

Alternatively, the *logit log* function can be used to linearize the standard curve, i.e. *logit* of absorbance (Y) is plotted against *log* of the known concentration (X) of standards.

The measured concentration of samples calculated from the standard curve must be multiplied by their respective dilution factor, because samples have been diluted prior to the assay; e.g. 200 pg/ml (from standard curve) x 3 (dilution factor) = 600 pg/ml.

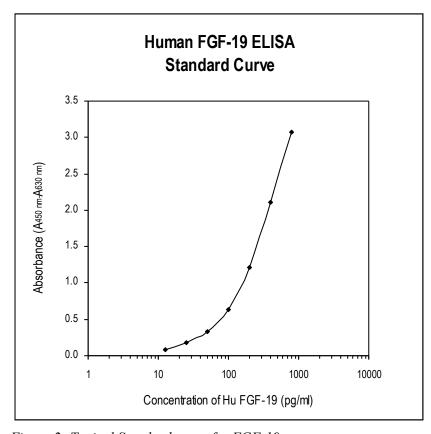


Figure 2: Typical Standard curve for FGF-19.







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13 PERFORMANCE CHARACTERISTICS

Typical analytical data of Human FGF-19 ELISA are presented in this chapter.

13.1 Sensitivity

<u>Limit of detection (LOD)</u> (defined as concentration of analyte giving absorbance higher than mean absorbance of blank* plus three standard deviations of the absorbance of blank: $A_{blank} + 3xSD_{blank}$) is calculated from the real human FGF-19 values in wells and is: 4.8 pg/ml.

13.2 Limit of assay

Results exceeding human FGF-19 level of 800 pg/ml should be repeated with more diluted samples. Dilution factor needs to be taken into consideration in calculating the FGF-19 concentration.

13.3 Specificity

The antibodies used in this ELISA are specific for human FGF-19. No cross-reactivity with recombinant human FGF-21 and recombinant human FGF-23 has been observed.

Sera of several mammalian species were measured in the assay. See results below.

For details please contact us at corp@drg-international.com

Mammalian serum	Observed
sample	cross reactivity
Bovine	no
Goat	no
Hamster	no
Horse	yes
Monkey	yes
Mouse	no
Pig	no
Rabbit	no
Rat	no
Sheep	no

13.4 Precision

Intra-assay (Within-Run, n=8)

^{*} Dilution Buffer is pipetted into Blank wells.



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Sample	Mean (pg/ml)	Standard Deviation (pg/ml)	CV (%)
1	196.0	13.7	7.0
2	384.0	19.2	5.0

Inter-assay (Run-to-Run, n=6)

Sample	Mean (pg/ml)	Standard Deviation (pg/ml)	CV (%)
1	194.0	16.5	8.5
2	462.0	30.0	6.5

13.5 Spiking Recovery

Serum samples were spiked with different amounts of human FGF-19 and assayed.

O bserved	Expected	Recovery O/E
(pg/ml)	(pg/ml)	(%)
121.1	-	-
859.2	962.1	89.3
493.5	587.1	84.1
364.8	399.6	91.3
415.2	-	-
1128.0	1165.2	96.8
724.5	790.2	91.7
545.5	602.0	90.5
	(pg/ml) 121.1 859.2 493.5 364.8 415.2 1128.0 724.5	(pg/ml) (pg/ml) 121.1 - 859.2 962.1 493.5 587.1 364.8 399.6 415.2 - 1128.0 1165.2 724.5 790.2

13.6 Linearity

Serum samples were serially diluted with Dilution Buffer and assayed.

Sample	Dilution	Observed (pg/ml)	Expected (pg/ml)	Recovery O/E (%)
	-	1023.0	-	-
1	2x	474.6	511.5	92.8
	4x	240.0	255.7	93.8
	8x	133.8	127.9	104.6
	-	1410.0	-	-
2	2x	651.0	705.0	92.3
	4x	357.0	352.5	101.3
	8x	164.1	176.2	93.1





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13.7 Effect of sample matrix

Citrate, heparin and EDTA plasmas were compared to respective serum samples from the same 10 individuals. Results are shown below:

la dividuala	Concentra	ation of FGF-	19 (pg/ml)	
Individuals No.	Serum	Plasma		
	Serum	Heparin	Citrate	EDTA
1	156.9	157.5	158.4	158.4
2	155.1	140.4	161.1	173.1
3	264.9	241.5	236.7	214.8
4	413.7	423.3	402.7	402.9
5	447.6	424.2	387.6	485.9
6	117.6	119.4	95.1	109.5
7	227.4	210.6	201.0	184.2
8	256.2	237.3	221.4	211.8
9	191.9	186.9	158.1	186.9
10	373.8	343.5	315.0	331.2
Mean	260.5	248.5	233.7	245.9
Mean Plasma/Serum	-	95%	90%	94%
Correlation coeff. R ²	-	0.99	0.97	0.94





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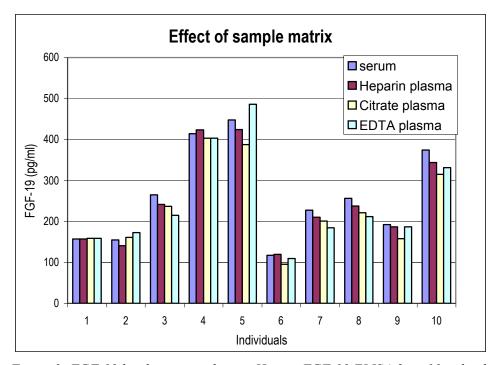


Figure 3: FGF-19 levels measured using Human FGF-19 ELISA from 10 individuals using serum, EDTA, citrate and heparin plasma, respectively.

13.8 Stability of samples stored at 2-8°C

Samples should be stored at -20°C. However, no decline in concentration of human FGF-19 was observed in serum and plasma samples after 7 days when stored at 2-8°C. To avoid microbial contamination, samples were treated with ε -aminocaproic acid and sodium azide, resulting in the final concentration of 0.03% and 0.1%, respectively.

Sample	Incubation Temp.	Serum	Plasma (pg/ml)		
Sample	Period	(pg/ml)	Heparin	Citrate	EDTA
	-20°C	256.2	237.3	221.4	211.8
1	2-8°C, 1 day	224.4	263.4	243.6	241.5
	2-8°C, 7 days	235.8	225.3	209.7	236.4
	-20°C	191.9	186.9	158.1	186.9
2	2-8°C, 1 day	186.9	170.4	169.2	162.3
	2-8°C, 7 days	183.9	183.3	151.8	176.4
	-20°C	373.8	343.5	315.0	331.2
3	2-8°C, 1 day	326.7	326.4	33.6	383.1
	2-8°C, 7 days	372.3	374.1	404.7	339.3





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13.9 Effect of Freezing/Thawing

No significant decline was observed in concentration of human FGF-19 in serum and plasma samples after repeated (5x) freeze/thaw cycles. However it is recommended to avoid unnecessary repeated freezing/thawing of the samples.

Sample	Number of f/t	Serum	Plasma (og/ml)	
Sample	cycles	(pg/ml)	Heparin	Citrate	EDTA
	1x	447.6	424.2	387.6	485.9
1	3x	463.8	459.6	387.9	408.3
	5x	411.6	364.2	386.1	439.2
	1x	117.6	119.4	95.1	109.5
2	3x	108.9	93.3	93.6	81.6
	5x	111.3	112.8	93.3	99.0
	1x	227.4	210.6	201.0	184.2
3	3x	207.6	168.9	203.4	207.3
	5x	225.6	210.6	188.7	174.6

14 DEFINITION OF THE STANDARD

The recombinant human FGF-19 is used as the Standard. The recombinant human FGF-19, produced in *E.coli*, is 23.03 kDa protein containing 192 amino acid residues of the human FGF-19 and 14 additional amino residues. The amino acid sequence of the recombinant human FGF-19 is 100% homologous to the amino acid sequence of the human FGF-19 without signal sequence.

15 PRELIMINARY POPULATION AND CLINICAL DATA

The following results were obtained when serum samples from 140 unselected donors (70 women + 70 men) 4 - 85 years old were assayed with the Human FGF-19 ELISA in our laboratory.

15.1 Reference range

The data quoted in these instructions should be used for guidance only. It is recommended that each laboratory include its own panel of control sample in the assay. Each laboratory should establish its own normal and pathological reference ranges for FGF-19 levels with the assay.







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15.2 Age and sex dependent distribution of FGF-19

Sex	Age	Age	Mean	SD	Min	Max	
Sex	(years)	"	FGF-19 (pg/ml)				
	4-19	7	155.9	72.8	47.7	266.7	
	20-39	16	282.9	189.7	45.9	615.3	
Men	40-59	20	235.3	160.0	55.5	511.8	
	60-79	20	304.3	263.0	64.8	1218.0	
	80-85	7	235.5	124.8	66.9	490.5	
	4-19	8	272.8	151.3	72.0	482.7	
	20-39	18	181.9	126.0	56.4	480.6	
Women	40-59	16	208.4	173.2	27.3	733.6	
	60-79	20	167.3	83.5	46.5	375.6	
	80-82	8	297.7	143.2	79.8	470.4	

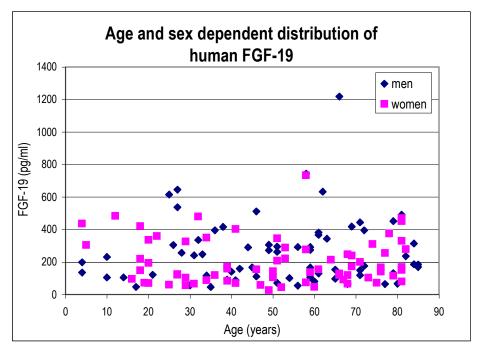


Figure 4: Human FGF-19 concentration plotted against donor age and sex.

16 METHOD COMPARISON

The Human FGF-19 ELISA was compared to the other commercial ELISA immunoassay, by measuring 39 serum samples. The following correlation graph was obtained.







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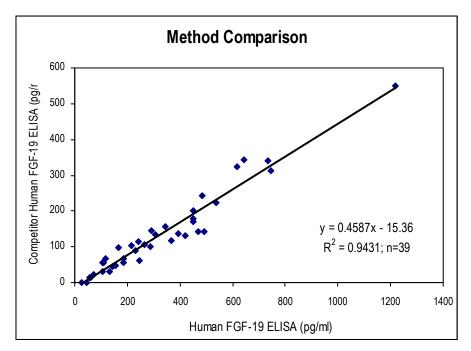


Figure 5: Method comparison

17 TROUBLESHOOTING AND FAQS

Weak signal in all wells

Possible explanations:

- Omission of a reagent or a step
- Improper preparation or storage of a reagent
- Assay performed before reagents were allowed to come to room temperature
- Improper wavelength when reading absorbance

High signal and background in all wells

Possible explanations:

- Improper or inadequate washing
- Overdeveloping; incubation time with Substrate Solution should be decreased before addition of Stop Solution
- Incubation temperature over 30°C





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High coefficient of variation (CV)

Possible explanation:

- Improper or inadequate washing
- Improper mixing Standards, Quality Controls or samples

18 REFERENCES / LITERATURE

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- 8. Ornitz DM and Itoh N: Fibroblast growth factors. Genome Biology 2(3), 3005.1-3005.12 (2001), Review

For more references on this product ask corp@drg-international.com

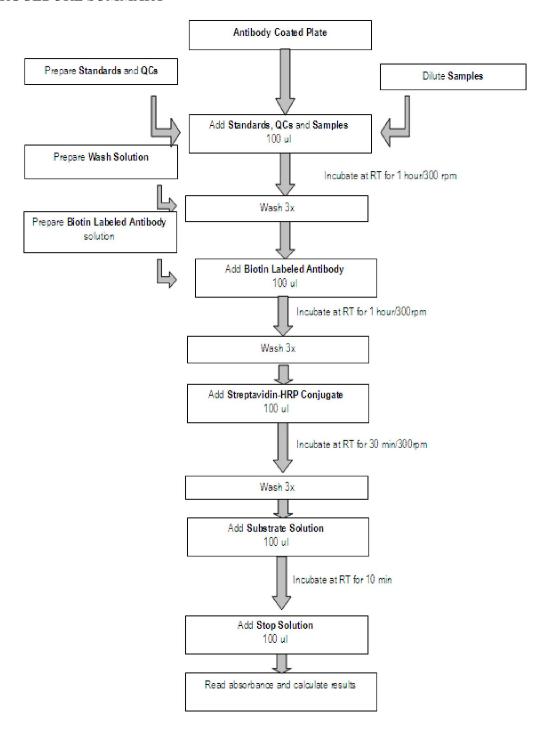




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19 ASSAY PROCEDURE SUMMARY

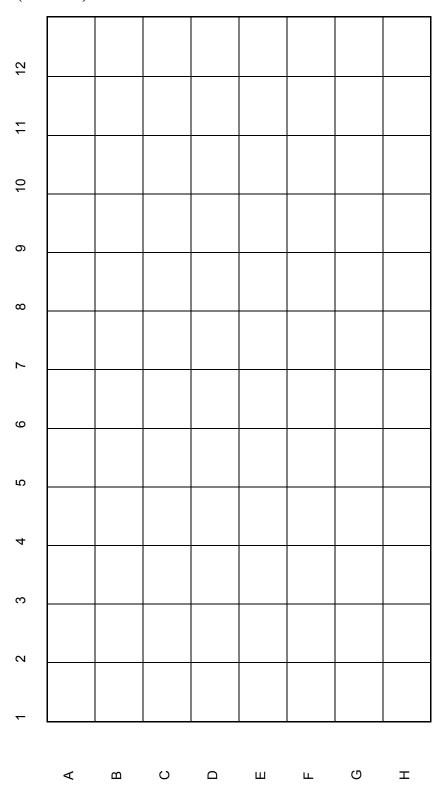






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SYMBOLS USED WITH DRG ASSAYS

Symbol	English	Deutsch	Français	Español	Italiano
[]i	Consult instructions for use	Gebrauchsanweisung beachten	Consulter les instructions d'utilisation	Consulte las instrucciones de uso	Consultare le istruzioni per l'uso
(€	European Conformity	CE-Konfirmitäts- kennzeichnung	Conformité aux normes européennes	Conformidad europea	Conformità europea
IVD	In vitro diagnostic device	In-vitro-Diagnostikum	Usage Diagnostic in vitro	Para uso Diagnóstico in vitro	Per uso Diagnostica in vitro
RUO	For research use only	Nur für Forschungszwecke	Seulement dans le cadre de recherches	Sólo para uso en investigación	Solo a scopo di ricerca
REF	Catalogue number	Katalog-Nr.	Numéro de catalogue	Número de catálogo	Numero di Catalogo
LOT	Lot. No. / Batch code	Chargen-Nr.	Numéro de lot	Número de lote	Numero di lotto
\sum	Contains sufficient for <n> tests/</n>	Ausreichend für "n" Ansätze	Contenu suffisant pour "n" tests	Contenido suficiente para <n> ensayos</n>	Contenuto sufficiente per "n" saggi
1	Storage Temperature	Lagerungstemperatur	Température de conservation	Temperatura de conservación	Temperatura di conservazione
	Expiration Date	Mindesthaltbarkeits-datum	Date limite d'utilisation	Fecha de caducidad	Data di scadenza
***	Legal Manufacturer	Hersteller	Fabricant	Fabricante	Fabbricante
Distributed by	Distributor	Vertreiber	Distributeur	Distribuidor	Distributore
Content	Content	Inhalt	Conditionnement	Contenido	Contenuto
Volume/No.	Volume / No.	Volumen/Anzahl	Volume/Quantité	Volumen/Número	Volume/Quantità

Symbol	Portugues	Dansk	Svenska	Ελληνικά
(li)	Consulte as instruções de utilização	Se brugsanvisning	Se bruksanvisningen	Εγχειρίδιο χρήστη
((Conformidade com as normas europeias	Europaeisk overensstemmelse	Europeisk överensstämmelse	Ευρωπαϊκή Συμμόρφωση
IVD	Diagnóstico in vitro	In vitro diagnostik	Diagnostik in vitro	in vitro διαγνωστικό
RUO				
REF	Catálogo n.º	Katalognummer	Katalog nummer	Αριθμός καταλόγου
LOT	No do lote	Lot nummer	Batch-nummer	Αριθμός Παρτίδος
\sum		Indeholder tilsttrækkeligt til "n" test	Innehåller tillräckligt till "n" tester	Περιεχόμενο επαρκές για «n» εξετάσεις
1	Temperatura de conservação	Opbevarings-temperatur	Förvaringstempratur	Θερμοκρασία αποθήκευσης
	Prazo de validade	Udløbsdato	Bäst före datum	Ημερομηνία λήξης
	Fabricante	Producent	Tillverkare	Κατασκευαστής
Distributed by				
Content	Conteúdo	Indhold	Innehåll	Περιεχόμενο
Volume/No.	Volume/Número	Volumen/antal	Volym/antal	Όγκος/αριθ