

SEQUENCE LISTING

<110> Ascendis Pharma A/S

<120> Controlled-release CNP agonist combination therapy

<130> CPX70585PC

<160> 99

<170> PatentIn version 3.5

<210> 1

<211> 22

<212> PRT

<213> Homo sapiens

<220>

<221> DISULFID

<222> (6)..(22)

<400> 1

Gly	Leu	Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile	Gly	Ser
1				5					10					15	

Met	Ser	Gly	Leu	Gly	Cys
			20		

<210> 2

<211> 53

<212> PRT

<213> Homo sapiens

<220>

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<222> (37)..(53)

<400> 2

Asp	Leu	Arg	Val	Asp	Thr	Lys	Ser	Arg	Ala	Ala	Trp	Ala	Arg	Leu	Leu
1				5					10					15	

Gln	Glu	His	Pro	Asn	Ala	Arg	Lys	Tyr	Lys	Gly	Ala	Asn	Lys	Lys	Gly
			20					25					30		

Leu	Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile	Gly	Ser	Met
-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----

35

40

45

Ser Gly Leu Gly Cys
50

<210> 3
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<400> 3

Gly Asp Leu Arg Val Asp Thr Lys Ser Arg Ala Ala Trp Ala Arg Leu
1 5 10 15

Leu Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys
20 25 30

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser
35 40 45

Met Ser Gly Leu Gly Cys
50

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<400> 4

Met Asp Leu Arg Val Asp Thr Lys Ser Arg Ala Ala Trp Ala Arg Leu
1 5 10 15

Leu Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys
20 25 30

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser
35 40 45

Met Ser Gly Leu Gly Cys
50

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<400> 5

Pro Asp Leu Arg Val Asp Thr Lys Ser Arg Ala Ala Trp Ala Arg Leu
1 5 10 15

Leu Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys
20 25 30

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser
35 40 45

Met Ser Gly Leu Gly Cys
50

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<222> (37)..(53)

<400> 6

Asp	Leu	Arg	Val	Asp	Thr	Lys	Ser	Arg	Ala	Ala	Trp	Ala	Arg	Leu	Leu
1				5					10					15	

Gln	Glu	His	Pro	Asn	Ala	Arg	Lys	Tyr	Lys	Gly	Ala	Asn	Lys	Lys	Gly
			20					25					30		

Leu	Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile	Gly	Ser	Asn
		35					40					45			

Ser	Gly	Leu	Gly	Cys
	50			

<210> 7

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<223> CNP-53 with deletion of amino acids 15-31

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<222> (20)..(36)

<400> 7

Asp	Leu	Arg	Val	Asp	Thr	Lys	Ser	Arg	Ala	Ala	Trp	Ala	Arg	Gly	Leu
1				5					10					15	

Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile	Gly	Ser	Met	Ser
			20					25					30		

Gly	Leu	Gly	Cys
		35	

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<400> 8

Leu Arg Val Asp Thr Lys Ser Arg Ala Ala Trp Ala Arg Leu Leu Gln
1 5 10 15

Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu
20 25 30

Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser
35 40 45

Gly Leu Gly Cys
50

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<400> 9

Arg Val Asp Thr Lys Ser Arg Ala Ala Trp Ala Arg Leu Leu Gln Glu
1 5 10 15

His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser
20 25 30

Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly
35 40 45

Leu Gly Cys
50

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<400> 10

Val Asp Thr Lys Ser Arg Ala Ala Trp Ala Arg Leu Leu Gln Glu His
1 5 10 15

Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys
20 25 30

Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu
35 40 45

Gly Cys
50

<210> 11
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<400> 11

Asp Thr Lys Ser Arg Ala Ala Trp Ala Arg Leu Leu Gln Glu His Pro
 1 5 10 15

Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly
 20 25 30

Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly
 35 40 45

Cys

<210> 12
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 <222> (32)..(48)

<400> 12

Thr Lys Ser Arg Ala Ala Trp Ala Arg Leu Leu Gln Glu His Pro Asn
 1 5 10 15

Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly Cys
 20 25 30

Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
 35 40 45

<210> 13
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<222> (31)..(47)

<400> 13

Lys Ser Arg Ala Ala Trp Ala Arg Leu Leu Gln Glu His Pro Asn Ala
1 5 10 15

Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly Cys Phe
20 25 30

Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
35 40 45

<210> 14

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<222> (30)..(46)

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Ser Arg Ala Ala Trp Ala Arg Leu Leu Gln Glu His Pro Asn Ala Arg
1 5 10 15

Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly Cys Phe Gly
20 25 30

Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
35 40 45

<210> 15

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<400> 15

Arg Ala Ala Trp Ala Arg Leu Leu Gln Glu His Pro Asn Ala Arg Lys
1 5 10 15

Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu
20 25 30

Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
35 40 45

<210> 16

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<222> (28)..(44)

<400> 16

Ala Ala Trp Ala Arg Leu Leu Gln Glu His Pro Asn Ala Arg Lys Tyr
1 5 10 15

Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys
20 25 30

Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
35 40

<210> 17

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<400> 17

Ala Ala Trp Ala Arg Leu Leu Gln Glu His Pro Asn Ala Gly Leu Ser
1 5 10 15

Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly
20 25 30

Leu Gly Cys
35

<210> 18

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<223> CNP-44 with a deletion of amino acids 15-22

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<400> 18

Ala Ala Trp Ala Arg Leu Leu Gln Glu His Pro Asn Ala Arg Gly Leu
1 5 10 15

Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser
20 25 30

Gly Leu Gly Cys
35

<210> 19

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<212> PRT

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<222> (27)..(43)

<400> 19

Ala	Trp	Ala	Arg	Leu	Leu	Gln	Glu	His	Pro	Asn	Ala	Arg	Lys	Tyr	Lys
1				5					10					15	

Gly	Ala	Asn	Lys	Lys	Gly	Leu	Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu
			20					25					30		

Asp	Arg	Ile	Gly	Ser	Met	Ser	Gly	Leu	Gly	Cys
			35				40			

<210> 20

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<212> PRT

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<223> CNP-42

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<222> (26)..(42)

<400> 20

Trp	Ala	Arg	Leu	Leu	Gln	Glu	His	Pro	Asn	Ala	Arg	Lys	Tyr	Lys	Gly
1				5					10					15	

Ala	Asn	Lys	Lys	Gly	Leu	Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp
			20					25					30		

Arg	Ile	Gly	Ser	Met	Ser	Gly	Leu	Gly	Cys
			35				40		

<210> 21

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<222> (25)..(41)

<400> 21

Ala	Arg	Leu	Leu	Gln	Glu	His	Pro	Asn	Ala	Arg	Lys	Tyr	Lys	Gly	Ala
1				5					10					15	

Asn	Lys	Lys	Gly	Leu	Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg
			20					25					30		

Ile	Gly	Ser	Met	Ser	Gly	Leu	Gly	Cys
		35					40	

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<222> (24)..(40)

<400> 22

Arg	Leu	Leu	Gln	Glu	His	Pro	Asn	Ala	Arg	Lys	Tyr	Lys	Gly	Ala	Asn
1				5					10					15	

Lys	Lys	Gly	Leu	Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile
			20					25					30		

Gly	Ser	Met	Ser	Gly	Leu	Gly	Cys
		35					40

<210> 23

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<222> (23)..(39)

<400> 23

Leu Leu Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys
1 5 10 15

Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly
20 25 30

Ser Met Ser Gly Leu Gly Cys
35

<210> 24

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<222> (22)..(38)

<400> 24

Leu Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys
1 5 10 15

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser
20 25 30

Met Ser Gly Leu Gly Cys
35

<210> 25

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<212> PRT

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<222> (21)..(37)

<400> 25

Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly
1 5 10 15

Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
20 25 30

Ser Gly Leu Gly Cys
35

<210> 26

<211> 37

<212> PRT

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<220>

<223> CNP-37 mit Q1pQ (pQ = pyroglutamate)

<220>

<221> MISC_FEATURE

<222> (1)..(1)

<223> X = pyroglutamate

<220>

<221> DISULFID

<222> (21)..(37)

<400> 26

Xaa Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly
1 5 10 15

Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
20 25 30

Ser Gly Leu Gly Cys

35

<210> 27
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<400> 27

Gly	Gln	Glu	His	Pro	Asn	Ala	Arg	Lys	Tyr	Lys	Gly	Ala	Asn	Lys	Lys
1				5				10					15		

Gly	Leu	Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile	Gly	Ser
			20					25					30		

Met	Ser	Gly	Leu	Gly	Cys
			35		

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<400> 28

Pro	Gln	Glu	His	Pro	Asn	Ala	Arg	Lys	Tyr	Lys	Gly	Ala	Asn	Lys	Lys
1				5				10					15		

Gly	Leu	Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile	Gly	Ser
			20					25					30		

Met Ser Gly Leu Gly Cys
35

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<400> 29

Met Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys
1 5 10 15

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser
20 25 30

Met Ser Gly Leu Gly Cys
35

<210> 30
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<400> 30

Pro Gly Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys
1 5 10 15

Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly
20 25 30

Ser Met Ser Gly Leu Gly Cys
35

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<400> 31

Met Gly Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys
1 5 10 15

Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly
20 25 30

Ser Met Ser Gly Leu Gly Cys
35

<210> 32
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<400> 32

Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly
1 5 10 15

Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Asn
20 25 30

Ser Gly Leu Gly Cys
35

<210> 33
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<400> 33

Gly Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys
1 5 10 15

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser
20 25 30

Asn Ser Gly Leu Gly Cys
35

<210> 34
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<400> 34

Gly Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Gln Lys
1 5 10 15

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser

20

25

30

Met Ser Gly Leu Gly Cys
35

<210> 35

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<400> 35

Gly	Gln	Glu	His	Pro	Asn	Ala	Arg	Lys	Tyr	Lys	Gly	Ala	Asn	Pro	Lys
1				5				10					15		

Gly	Leu	Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile	Gly	Ser
			20					25					30		

Met Ser Gly Leu Gly Cys
35

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<223> G-CNP-37 K14Q, deletion of amino acid 15

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<222> (21)..(37)

<400> 36

Gly	Gln	Glu	His	Pro	Asn	Ala	Arg	Lys	Tyr	Lys	Gly	Ala	Asn	Gln	Gly
1				5				10					15		

Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
 20 25 30

Ser Gly Leu Gly Cys
 35

<210> 37
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<400> 37

Gly Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Gln Gln
 1 5 10 15

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser
 20 25 30

Met Ser Gly Leu Gly Cys
 35

<210> 38
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<400> 38

Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu
 1 5 10 15

Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser
20 25 30

Gly Leu Gly Cys
35

<210> 39
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<400> 39

His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser
1 5 10 15

Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly
20 25 30

Leu Gly Cys
35

<210> 40
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<222> (18)..(34)

<400> 40

Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys
1 5 10 15

Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu
 20 25 30

Gly Cys

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<400> 41

Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly
 1 5 10 15

Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly
 20 25 30

Cys

<210> 42
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<400> 42

Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly Cys

1	5	10	15
Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys			
	20	25	30

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<400> 43

Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly Cys Phe
1 5 10 15

Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25 30

<210> 44
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 <222> (14)..(30)

<400> 44

Lys Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly Cys Phe Gly
1 5 10 15

Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25 30

<210> 45
<211> 29
<212> PRT
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<400> 45

Tyr Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu
1 5 10 15

Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25

<210> 46
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<400> 46

Lys Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys
1 5 10 15

Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25

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<222> (22)..(38)

<400> 47

Gly His Lys Ser Glu Val Ala His Arg Phe Lys Gly Ala Asn Lys Lys
1 5 10 15

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser
20 25 30

Met Ser Gly Leu Gly Cys
35

<210> 48

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<222> (11)..(27)

<400> 48

Gly Ala Asn Lys Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu
1 5 10 15

Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25

<210> 49

<211> 27

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<220>

<223> CNP-27 K4Q, K5Q

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<222> (11)..(27)

<400> 49

Gly Ala Asn Gln Gln Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu
1 5 10 15

Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25

<210> 50
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<400> 50

Gly Ala Asn Arg Arg Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu
1 5 10 15

Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25

<210> 51
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<220>
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<400> 51

Gly Ala Asn Pro Arg Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu
1 5 10 15

Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25

<210> 52
<211> 27
<212> PRT
<213> Artificial Sequence

<220>
<223> CNP-27 K4S, K5S

<220>
<221> DISULFID
<222> (11)..(27)

<400> 52

Gly Ala Asn Ser Ser Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu
1 5 10 15

Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25

<210> 53
<211> 30
<212> PRT
<213> Artificial Sequence

<220>
<223> GAN-CNP-27 K4P, K5R

<220>
<221> DISULFID
<222> (14)..(30)

<400> 53

Gly Ala Asn Gly Ala Asn Pro Arg Gly Leu Ser Arg Gly Cys Phe Gly
1 5 10 15

Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25 30

<210> 54

<211> 27
<212> PRT
<213> Artificial Sequence

<220>
<223> CNP-27 K4R, K5R, K9R

<220>
<221> DISULFID
<222> (11)..(27)

<400> 54

Gly Ala Asn Arg Arg Gly Leu Ser Arg Gly Cys Phe Gly Leu Lys Leu
1 5 10 15

Asp Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25

<210> 55
<211> 27
<212> PRT
<213> Artificial Sequence

<220>
<223> CNP-27 K4R, K5R, K9R, M22N

<220>
<221> DISULFID
<222> (11)..(27)

<400> 55

Gly Ala Asn Arg Arg Gly Leu Ser Arg Gly Cys Phe Gly Leu Lys Leu
1 5 10 15

Asp Arg Ile Gly Ser Asn Ser Gly Leu Gly Cys
20 25

<210> 56
<211> 28
<212> PRT
<213> Artificial Sequence

<220>
<223> P-CNP-27 K4R, K5R, K9R

<220>

<221> DISULFID

<222> (12)..(28)

<400> 56

Pro	Gly	Ala	Asn	Arg	Arg	Gly	Leu	Ser	Arg	Gly	Cys	Phe	Gly	Leu	Lys
1				5					10					15	

Leu	Asp	Arg	Ile	Gly	Ser	Met	Ser	Gly	Leu	Gly	Cys
			20					25			

<210> 57

<211> 28

<212> PRT

<213> Artificial Sequence

<220>

<223> M-CNP-27 K4R, K5R, K9R

<220>

<221> DISULFID

<222> (12)..(28)

<400> 57

Met	Gly	Ala	Asn	Arg	Arg	Gly	Leu	Ser	Arg	Gly	Cys	Phe	Gly	Leu	Lys
1				5					10					15	

Leu	Asp	Arg	Ile	Gly	Ser	Met	Ser	Gly	Leu	Gly	Cys
			20					25			

<210> 58

<211> 38

<212> PRT

<213> Artificial Sequence

<220>

<223> Human Serum Albumine Fragment - CNP-27

<220>

<221> DISULFID

<222> (22)..(38)

<400> 58

Gly His Lys Ser Glu Val Ala His Arg Phe Lys Gly Ala Asn Lys Lys
1 5 10 15

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser
20 25 30

Met Ser Gly Leu Gly Cys
35

<210> 59
<211> 38
<212> PRT
<213> Artificial Sequence

<220>
<223> Human Serum Albumine Fragment - CNP-27 M22N

<220>
<221> DISULFID
<222> (22)..(38)

<400> 59

Gly His Lys Ser Glu Val Ala His Arg Phe Lys Gly Ala Asn Lys Lys
1 5 10 15

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser
20 25 30

Asn Ser Gly Leu Gly Cys
35

<210> 60
<211> 39
<212> PRT
<213> Artificial Sequence

<220>
<223> methionine - Human Serum Albumine Fragment - CNP-27

<220>
<221> DISULFID
<222> (23)..(39)

<400> 60

Met Gly His Lys Ser Glu Val Ala His Arg Phe Lys Gly Ala Asn Lys
1 5 10 15

Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly
20 25 30

Ser Met Ser Gly Leu Gly Cys
35

<210> 61
<211> 39
<212> PRT
<213> Artificial Sequence

<220>
<223> proline - Human Serum Albumine Fragment - CNP-27

<220>
<221> DISULFID
<222> (23)..(39)

<400> 61

Pro Gly His Lys Ser Glu Val Ala His Arg Phe Lys Gly Ala Asn Lys
1 5 10 15

Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly
20 25 30

Ser Met Ser Gly Leu Gly Cys
35

<210> 62
<211> 26
<212> PRT
<213> Artificial Sequence

<220>
<223> CNP-26

<220>
<221> DISULFID
<222> (10)..(26)

<400> 62

Ala Asn Lys Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp
1 5 10 15

Arg Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25

<210> 63

<211> 25

<212> PRT

<213> Artificial Sequence

<220>

<223> CNP-25

<220>

<221> DISULFID

<222> (9)..(25)

<400> 63

Asn Lys Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg
1 5 10 15

Ile Gly Ser Met Ser Gly Leu Gly Cys
20 25

<210> 64

<211> 24

<212> PRT

<213> Artificial Sequence

<220>

<223> CNP-24

<220>

<221> DISULFID

<222> (8)..(24)

<400> 64

Lys Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile
1 5 10 15

Gly Ser Met Ser Gly Leu Gly Cys

20

<210> 65
<211> 23
<212> PRT
<213> Artificial Sequence

<220>
<223> CNP-23

<220>
<221> DISULFID
<222> (7)..(23)

<400> 65

Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly
1 5 10 15

Ser Met Ser Gly Leu Gly Cys
20

<210> 66
<211> 23
<212> PRT
<213> Artificial Sequence

<220>
<223> R-CNP-22

<220>
<221> DISULFID
<222> (7)..(23)

<400> 66

Arg Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly
1 5 10 15

Ser Met Ser Gly Leu Gly Cys
20

<210> 67
<211> 24
<212> PRT
<213> Artificial Sequence

<220>
<223> ER-CNP-22

<220>
<221> DISULFID
<222> (8)..(24)

<400> 67

Glu	Arg	Gly	Leu	Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile
1				5					10					15	

Gly	Ser	Met	Ser	Gly	Leu	Gly	Cys
				20			

<210> 68
<211> 23
<212> PRT
<213> Artificial Sequence

<220>
<223> R-CNP-22 K4R

<220>
<221> DISULFID
<222> (7)..(23)

<400> 68

Arg	Gly	Leu	Ser	Arg	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile	Gly
1				5					10					15	

Ser	Met	Ser	Gly	Leu	Gly	Cys
				20		

<210> 69
<211> 24
<212> PRT
<213> Artificial Sequence

<220>
<223> ER-CNP-22 4KR

<220>
<221> DISULFID

<222> (8)..(24)

<400> 69

Glu Arg Gly Leu Ser Arg Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile
1 5 10 15

Gly Ser Met Ser Gly Leu Gly Cys
20

<210> 70

<211> 24

<212> PRT

<213> Artificial Sequence

<220>

<223> RR-CNP-22

<220>

<221> DISULFID

<222> (8)..(24)

<400> 70

Arg Arg Gly Leu Ser Arg Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile
1 5 10 15

Gly Ser Met Ser Gly Leu Gly Cys
20

<210> 71

<211> 37

<212> PRT

<213> Artificial Sequence

<220>

<223> HRGP fragment - CNP-22

<220>

<221> DISULFID

<222> (21)..(37)

<400> 71

Gly His His Ser His Glu Gln His Pro His Gly Ala Asn Gln Gln Gly
1 5 10 15

Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
 20 25 30

Ser Gly Leu Gly Cys
 35

<210> 72
 <211> 38
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> HRGP fragment - CNP-22

<220>
 <221> DISULFID
 <222> (22)..(38)

<400> 72

Gly Ala His His Pro His Glu His Asp Thr His Gly Ala Asn Gln Gln
 1 5 10 15

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser
 20 25 30

Met Ser Gly Leu Gly Cys
 35

<210> 73
 <211> 37
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> HRGP fragment - CNP-22

<220>
 <221> DISULFID
 <222> (21)..(37)

<400> 73

Gly His His Ser His Glu Gln His Pro His Gly Ala Asn Pro Arg Gly
 1 5 10 15

Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
20 25 30

Ser Gly Leu Gly Cys
35

<210> 74
<211> 36
<212> PRT
<213> Artificial Sequence

<220>
<223> IgG1(FC) fragment - CNP-22

<220>
<221> DISULFID
<222> (20)..(36)

<400> 74

Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Ser Gly Leu
1 5 10 15

Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser
20 25 30

Gly Leu Gly Cys
35

<210> 75
<211> 39
<212> PRT
<213> Artificial Sequence

<220>
<223> Human Serum Albumine - CNP-22

<220>
<221> DISULFID
<222> (23)..(39)

<400> 75

Gly Gln His Lys Asp Asp Asn Pro Asn Leu Pro Arg Gly Ala Asn Pro

1	5	10	15
Arg Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly			
	20	25	30

Ser Met Ser Gly Leu Gly Cys
35

<210> 76
 <211> 37
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> Human Serum Albumine - CNP-22

<220>
 <221> DISULFID
 <222> (21)..(37)

<400> 76

Gly Glu Arg Ala Phe Lys Ala Trp Ala Val Ala Arg Leu Ser Gln Gly
1 5 10 15

Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
20 25 30

Ser Gly Leu Gly Cys
35

<210> 77
 <211> 35
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> osteocrin NPR C inhibitor fragment - CNP-22

<220>
 <221> DISULFID
 <222> (19)..(35)

<400> 77

Phe Gly Ile Pro Met Asp Arg Ile Gly Arg Asn Pro Arg Gly Leu Ser
 1 5 10 15

Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly
 20 25 30

Leu Gly Cys
 35

<210> 78
 <211> 40
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> FGF2 heparin-binding domain fragment - CNP-22

<220>
 <221> DISULFID
 <222> (24)..(40)

<400> 78

Gly Lys Arg Thr Gly Gln Tyr Lys Leu Gly Ser Lys Thr Gly Pro Gly
 1 5 10 15

Pro Lys Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile
 20 25 30

Gly Ser Met Ser Gly Leu Gly Cys
 35 40

<210> 79
 <211> 37
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> IgG1(FC) fragment - CNP-22 K4R

<220>
 <221> DISULFID
 <222> (21)..(37)

<400> 79

Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Gly Ala Asn Gln Gln Gly
 1 5 10 15

Leu Ser Arg Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
 20 25 30

Ser Gly Leu Gly Cys
 35

<210> 80
 <211> 36
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> Human Serum Albumine fragment - CNP-22 K4R

<220>
 <221> DISULFID
 <222> (20)..(36)

<400> 80

Gly Val Pro Gln Val Ser Thr Ser Thr Gly Ala Asn Gln Gln Gly Leu
 1 5 10 15

Ser Arg Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser
 20 25 30

Gly Leu Gly Cys
 35

<210> 81
 <211> 37
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> fibronectin fragment - CNP-22

<220>
 <221> DISULFID
 <222> (21)..(37)

<400> 81

Gly Gln Pro Ser Ser Ser Ser Gln Ser Thr Gly Ala Asn Gln Gln Gly
1 5 10 15

Leu Ser Arg Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
20 25 30

Ser Gly Leu Gly Cys
35

<210> 82

<211> 37

<212> PRT

<213> Artificial Sequence

<220>

<223> fibronectin fragment - CNP-22 K4R

<220>

<221> DISULFID

<222> (21)..(37)

<400> 82

Gly Gln Thr His Ser Ser Gly Thr Gln Ser Gly Ala Asn Gln Gln Gly
1 5 10 15

Leu Ser Arg Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
20 25 30

Ser Gly Leu Gly Cys
35

<210> 83

<211> 37

<212> PRT

<213> Artificial Sequence

<220>

<223> fibronectin fragment - CNP-22 K4R

<220>

<221> DISULFID

<222> (21)..(37)

<400> 83

Gly Ser Thr Gly Gln Trp His Ser Glu Ser Gly Ala Asn Gln Gln Gly
1 5 10 15

Leu Ser Arg Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
20 25 30

Ser Gly Leu Gly Cys
35

<210> 84

<211> 37

<212> PRT

<213> Artificial Sequence

<220>

<223> zinc finger fragment - CNP-22 K4R

<220>

<221> DISULFID

<222> (21)..(37)

<400> 84

Gly Ser Ser Ser Ser Ser Ser Ser Ser Ser Gly Ala Asn Gln Gln Gly
1 5 10 15

Leu Ser Arg Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
20 25 30

Ser Gly Leu Gly Cys
35

<210> 85

<211> 21

<212> PRT

<213> Artificial Sequence

<220>

<223> CNP-21

<220>

<221> DISULFID

<222> (5)..(21)

<400> 85

Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
1 5 10 15

Ser Gly Leu Gly Cys
20

<210> 86

<211> 20

<212> PRT

<213> Artificial Sequence

<220>

<223> CNP-20

<220>

<221> DISULFID

<222> (4)..(20)

<400> 86

Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser
1 5 10 15

Gly Leu Gly Cys
20

<210> 87

<211> 19

<212> PRT

<213> Artificial Sequence

<220>

<223> CNP-19

<220>

<221> DISULFID

<222> (3)..(19)

<400> 87

Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met Ser Gly
1 5 10 15

Leu Gly Cys

<210> 88
<211> 18
<212> PRT
<213> Artificial Sequence

<220>
<223> CNP-18

<220>
<221> DISULFID
<222> (2)..(18)

<400> 88

Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile	Gly	Ser	Met	Ser	Gly	Leu
1				5					10					15	

Gly Cys

<210> 89
<211> 17
<212> PRT
<213> Artificial Sequence

<220>
<223> CNP-17

<220>
<221> DISULFID
<222> (1)..(17)

<400> 89

Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile	Gly	Ser	Met	Ser	Gly	Leu	Gly
1				5					10					15	

Cys

<210> 90
<211> 32

<212> PRT
<213> Artificial Sequence

<220>
<223> BNP fragment - CNP-17 - BNP- fragment

<220>
<221> DISULFID
<222> (10)..(26)

<400> 90

Ser Pro Lys Met Val Gln Gly Ser Gly Cys Phe Gly Leu Lys Leu Asp
1 5 10 15

Arg Ile Gly Ser Met Ser Gly Leu Gly Cys Lys Val Leu Arg Arg His
20 25 30

<210> 91
<211> 38
<212> PRT
<213> Artificial Sequence

<220>
<223> CNP-38 L1G

<220>
<221> DISULFID
<222> (22)..(38)

<400> 91

Gly Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys
1 5 10 15

Gly Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser
20 25 30

Met Ser Gly Leu Gly Cys
35

<210> 92
<211> 37
<212> PRT
<213> Artificial Sequence

<220>
<223> Ac-CNP-37

<220>
<221> MOD_RES
<222> (1)..(1)
<223> ACETYLATION

<220>
<221> DISULFID
<222> (21)..(37)

<400> 92

Gln Glu His Pro Asn Ala Arg Lys Tyr Lys Gly Ala Asn Lys Lys Gly
1 5 10 15

Leu Ser Lys Gly Cys Phe Gly Leu Lys Leu Asp Arg Ile Gly Ser Met
20 25 30

Ser Gly Leu Gly Cys
35

<210> 93
<211> 37
<212> PRT
<213> Artificial Sequence

<220>
<223> CNP-37, Xaa = K or R, with the provision that at least one Xaa is R

<220>
<221> MISC_FEATURE
<222> (8)..(8)
<223> Xaa = Lys, Arg, Pro, Ser or Gln, with the provision that at least one of amino acids 8, 10, 14, 15, 19 or 25 is selected from the group consisting of Arg, Pro, Ser and Gln

<220>
<221> MISC_FEATURE
<222> (10)..(10)
<223> Xaa = Lys, Arg, Pro, Ser or Gln, with the provision that at least one of amino acids 8, 10, 14, 15, 19 or 25 is selected from the group consisting of Arg, Pro, Ser and Gln

<220>
<221> MISC_FEATURE

<222> (14)..(14)
 <223> Xaa = Lys, Arg, Pro, Ser or Gln, with the provision that at least one of amino acids 8, 10, 14, 15, 19 or 25 is selected from the group consisting of Arg, Pro, Ser and Gln

<220>
 <221> MISC_FEATURE
 <222> (15)..(15)
 <223> Xaa = Lys, Arg, Pro, Ser or Gln, with the provision that at least one of amino acids 8, 10, 14, 15, 19 or 25 is selected from the group consisting of Arg, Pro, Ser and Gln

<220>
 <221> MISC_FEATURE
 <222> (19)..(19)
 <223> Xaa = Lys, Arg, Pro, Ser or Gln, with the provision that at least one of amino acids 8, 10, 14, 15, 19 or 25 is selected from the group consisting of Arg, Pro, Ser and Gln

<220>
 <221> DISULFID
 <222> (21)..(37)

<220>
 <221> MISC_FEATURE
 <222> (25)..(25)
 <223> Xaa = Lys, Arg, Pro, Ser or Gln, with the provision that at least one of amino acids 8, 10, 14, 15, 19 or 25 is selected from the group consisting of Arg, Pro, Ser and Gln

<400> 93

Gln	Glu	His	Pro	Asn	Ala	Arg	Xaa	Tyr	Xaa	Gly	Ala	Asn	Xaa	Xaa	Gly
1				5					10					15	

Leu	Ser	Xaa	Gly	Cys	Phe	Gly	Leu	Xaa	Leu	Asp	Arg	Ile	Gly	Ser	Met
			20					25					30		

Ser	Gly	Leu	Gly	Cys
				35

<210> 94
 <211> 37
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> mutated CNP-37

<220>
 <221> MISC_FEATURE
 <222> (14)..(14)
 <223> X is selected from the group consising of Lys, Arg, Pro, Ser and Gln, with the provision that at least one of the amino acids at position 14, 15, 19 and 25 is selected from the group consisting of Arg, Pro, Ser and Gln

<220>
 <221> MISC_FEATURE
 <222> (15)..(15)
 <223> X is selected from the group consising of Lys, Arg, Pro, Ser and Gln, with the provision that at least one of the amino acids at position 14, 15, 19 and 25 is selected from the group consisting of Arg, Pro, Ser and Gln

<220>
 <221> MISC_FEATURE
 <222> (19)..(19)
 <223> X is selected from the group consising of Lys, Arg, Pro, Ser and Gln, with the provision that at least one of the amino acids at position 14, 15, 19 and 25 is selected from the group consisting of Arg, Pro, Ser and Gln

<220>
 <221> DISULFID
 <222> (21)..(37)

<220>
 <221> MISC_FEATURE
 <222> (25)..(25)
 <223> X is selected from the group consising of Lys, Arg, Pro, Ser and Gln, with the provision that at least one of the amino acids at position 14, 15, 19 and 25 is selected from the group consisting of Arg, Pro, Ser and Gln

<400> 94

Gln	Glu	His	Pro	Asn	Ala	Arg	Lys	Tyr	Lys	Gly	Ala	Asn	Xaa	Xaa	Gly
1				5					10					15	

Leu	Ser	Xaa	Gly	Cys	Phe	Gly	Leu	Xaa	Leu	Asp	Arg	Ile	Gly	Ser	Met
			20					25					30		

Ser	Gly	Leu	Gly	Cys
				35

<210> 95
 <211> 37
 <212> PRT

<213> Artificial Sequence

<220>

<223> mutated CNP-37

<220>

<221> MISC_FEATURE

<222> (14)..(15)

<223> Xaa Xaa is selected from the group consisting of Lys Arg, Arg Lys, Lys Pro, Pro Lys, Ser Ser, Arg Ser, Ser Arg, Gln Lys, Gln Arg, Lys Gln, Arg Gln, Arg Arg and Gln Gln

<220>

<221> DISULFID

<222> (21)..(37)

<400> 95

Gln	Glu	His	Pro	Asn	Ala	Arg	Lys	Tyr	Lys	Gly	Ala	Asn	Xaa	Xaa	Gly
1				5					10					15	

Leu	Ser	Lys	Gly	Cys	Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile	Gly	Ser	Met
			20					25					30		

Ser	Gly	Leu	Gly	Cys
				35

<210> 96

<211> 15

<212> PRT

<213> Homo sapiens

<400> 96

Phe	Gly	Leu	Lys	Leu	Asp	Arg	Ile	Gly	Ser	Met	Ser	Gly	Leu	Gly
1				5					10					15

<210> 97

<211> 20

<212> PRT

<213> Artificial Sequence

<220>

<223> Artificial random coil

<400> 97

Gly Gly Pro Gly Gly Pro Gly Pro Gly Gly Pro Gly Gly Pro Gly Pro

1 5 10 15

Gly Gly Pro Gly
20

<210> 98
<211> 13
<212> PRT
<213> Artificial Sequence

<220>
<223> NPR-C inhibitor

<400> 98

Phe Gly Ile Pro Met Asp Arg Ile Gly Arg Asn Pro Arg
1 5 10

<210> 99
<211> 191
<212> PRT
<213> Homo sapiens

<400> 99

Phe Pro Thr Ile Pro Leu Ser Arg Leu Phe Asp Asn Ala Met Leu Arg
1 5 10 15

Ala His Arg Leu His Gln Leu Ala Phe Asp Thr Tyr Gln Glu Phe Glu
20 25 30

Glu Ala Tyr Ile Pro Lys Glu Gln Lys Tyr Ser Phe Leu Gln Asn Pro
35 40 45

Gln Thr Ser Leu Cys Phe Ser Glu Ser Ile Pro Thr Pro Ser Asn Arg
50 55 60

Glu Glu Thr Gln Gln Lys Ser Asn Leu Glu Leu Leu Arg Ile Ser Leu
65 70 75 80

Leu Leu Ile Gln Ser Trp Leu Glu Pro Val Gln Phe Leu Arg Ser Val
85 90 95

Phe Ala Asn Ser Leu Val Tyr Gly Ala Ser Asp Ser Asn Val Tyr Asp

100

105

110

Leu Leu Lys Asp Leu Glu Glu Gly Ile Gln Thr Leu Met Gly Arg Leu
 115 120 125

Glu Asp Gly Ser Pro Arg Thr Gly Gln Ile Phe Lys Gln Thr Tyr Ser
 130 135 140

Lys Phe Asp Thr Asn Ser His Asn Asp Asp Ala Leu Leu Lys Asn Tyr
 145 150 155 160

Gly Leu Leu Tyr Cys Phe Arg Lys Asp Met Asp Lys Val Glu Thr Phe
 165 170 175

Leu Arg Ile Val Gln Cys Arg Ser Val Glu Gly Ser Cys Gly Phe
 180 185 190