SEQUENCE LISTING

<110> Ascendis Pharma A/S

<120> Controlled-release CNP agonists with increased NEP stability

<130> CPX70164PC

<160> 98

<170> PatentIn version 3.5

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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

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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

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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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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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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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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

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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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<210> 8
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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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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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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
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<210>  11
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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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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

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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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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

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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

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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

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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

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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

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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

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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

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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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<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

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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

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<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

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<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
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<220>
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<223> X = pyroglutamate

<220>
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<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

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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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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

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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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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

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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

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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

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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

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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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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

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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

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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

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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

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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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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

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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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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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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
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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

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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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<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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<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

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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

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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

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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

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<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

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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
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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

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<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
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<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