

Р+РКРҮРҮР РЈРЛР|Р+РІ

РrP»CII PjPµPrPëC#PSPsPiPs P·P°CГC,PsCГCfPIP°PSPSCII P»C-РeP°CГCГCЬPePsPiPs P·P°CГCfPsP±Cf

РҮР•Р'Р+РЛРРђР

(SEVIKAR)

РҮРeP»P°Pr:

PrC-CГC#P° CГPµC#PsPIPëPSP°: PsP»PjPµCГP°CГC,P°PSCf PjPµPrPsPeCГPsPjC-P»; P°PjP»PsPrPëPïC-PSCf P±PµCГPëP»P°C,;

1 C,P°P±P»PµC,PeP°, PIPeCГPëC,P° PïP»C-PIPePsPIPsCГ PsP±PsP»PsPSPePsCГ, PjC-CГC,PëC,CГ PsP»PjPµCГP°CГC,P°PSCf PjPµPrPsPeCГPsPjC-P»Cf 20 PjPi, P°PjP»PsPrPëPïC-PSCf P±PµCГPëP»P°C,Cf 6,944 PjPi, C%oPs PµPePIC-PIP°P»PµPSC,PSPs P°PjP»PsPrPëPïC-PSCf 5 PjPi;

P°P±Ps PsP»PjPµCГP°CГC,P°PSCf PjPµPrPsPeCГPsPjC-P»Cf 40 PjPi, P°PjP»PsPrPëPïC-PSCf P±PµCГPëP»P°C,Cf 6,944 PjPi, C%oPs PµPePIC-PIP°P»PµPSC,PSPs P°PjP»PsPrPëPïC-PSCf 5 PjPi;

P°P±Ps PsP»PjPµCГP°CГC,P°PSCf PjPµPrPsPeCГPsPjC-P»Cf 40 PjPi, P°PjP»PsPrPëPïC-PSCf P±PµCГPëP»P°C,Cf 13,888 PjPi, C%oPs PµPePIC-PIP°P»PµPSC,PSPs P°PjP»PsPrPëPïC-PSCf 10 PjPi;

PrPsPïPsPjC-P¶PSC- CГPµC#PsPIPëPSPë: PeCГPsC...PjP°P»CГ PïCГPµP¶PµP»P°C,PëPSC-P·PsPIP°PSPëPNë PeCfPeCfCГCfPrP·C¶PSPëPNë, C†PµP»CГP»PsP·P° PjC-PeCГPsPeCГPëCГC,P°P»C-C†PSP° CГPëP»C-PePsPIP°PSP° (PjC-CГC,PëC,CГ 98 % PjC-PeCГPsPeCГPëCГC,P°P»C-C†PSPsC— C†PµP»CГP»PsP·Pë (Ph. Eur.) C,P° 2 % PeCГPµPjPSC-CГ PrC-PsPeCГPëPrCf PePsP»PsC—PrPSPsPiPs P±PµP·PIPsPrPSPsPiPs (Ph. Eur.)), PSP°C,CГC-CГ PeCГPsCГPeP°CГPjPµP»PsP·P°, PjP°PiPSC-CГ CГC,PµP°CГP°C, (CГPsCГP»PëPSPSPsPiPs PïPsC...PsPrP¶PµPSPSC¶);

PïP»C-PIPePsPIP° PsP±PsP»PsPSPeP° PjC-CГC,PëC,CГ: PһPïP°PrCГP°PNë II 85F18422 P±C-P»PëPNë P°P±Ps PһPïP°PrCГP°PNë II 85F22093 P¶PsPIC,PëPNë, P°P±Ps PһPïP°PrCГP°PNë II 85F25467 C†PµCГPIPSPSPëPNë (CГPïPëCГC, PïPsP»C-PIC-PSC-P»PsPIPëPNë, C,PëC,P°PSCf PrC-PsPeCГPëPr (P· 171), PïPsP»C-PµC,PëP»PµPSPiP»C-PePsP»CГ, C,P°P»CГPe, P·P°P»C-P·P° PsPeCГPëPr P¶PsPIC,PëPNë (P· 172), P·P°P»C-P·P° PsPeCГPëPr C†PµCГPIPSPSPëPNë (P· 172)).

Р·С-РeP°CГCГCЬPeP° C,,PsCГPjP°. РҮР•Р±Р»РµС,РeРë, PIPeCГPëC,C- PïP»C-PIPePsPIPsCГ PsP±PsP»PsPSPePsCГ.

PһCГPSPsPIpsc- C,,C-P·PëPePs-C...C-PjC-C†PSC- PIP»P°CГC,PëPIPсCГC,C-:

PrP»CЦ PrPsP·CзPIP°PSPSCЦ 20 PjPi/5 PjPi: P±C-P»C- PeCЦCрPiP»C- C,P°P±P»PμC,PePë, PIPeCЦPëC,C- PïP»C-PIPePsPIPsCЦ PsP±PsP»PsPSPePsCЦ, CЦPsP·PjC-CЦPsPj 6 PjPj, P· C,PëCГPSPμPSPSCЦPj B«C73B» P· PsPrPSPsPiPs P±PsPeCр;

PrP»CЦ PrPsP·CзPIP°PSPSCЦ 40 PjPi/5 PjPi: PeCЦPμPjPsPIC- PeCЦCрPiP»C- C,P°P±P»PμC,PePë, PIPeCЦPëC,C- PïP»C-PIPePsPIPsCЦ PsP±PsP»PsPSPePsCЦ, CЦPsP·PjC-CЦPsPj 8 PjPj, P· C,PëCГPSPμPSPSCЦPj B«C75B» P· PsPrPSPsPiPs P±PsPeCр;

PrP»CЦ PrPsP·CзPIP°PSPSCЦ 40 PjPi/10 PjPi: PePsCЦPëC±PSCЦPIP°C,Ps-C±PμCЦPIPsPSC- PeCЦCрPiP»C- C,P°P±P»PμC,PePë, PIPeCЦPëC,C- PïP»C-PIPePsPIPsCЦ PsP±PsP»PsPSPePsCЦ, CЦPsP·PjC-CЦPsPj 8 PjPj, P· C,PëCГPSPμPSPSCЦPj B«C77B» P· PsPrPSPsPiPs P±PsPeCр.

PrP°CЦPjP°PePsC,PμCЦP°PïPμPIC,PëC±PSP° PiCЦCрPïP°.

PŸPμCЦC†PμPIPs-CГCрPrPëPSPSP° CГPëCГC,PμPjP°. P»C-PeP°CЦCГCЦPëC- P·P°CГPsP±Pë, C%Ps PIPïP»PëPIP°CЦC,CH PSP° CЦPμPSC-PS-P°PSPiC-PsC,PμPSP·PëPSPsPICр CГPëCГC,PμPjCр. PьPsPjP±C-PSP°C†C-C— P±P»PsPeP°C,PsCЦC-PI CЦPμC†PμPïC,PsCЦP° P°PSPiC-PsC,PμPSP·PëPSCр II. PђPSC,P°PiPsPSC-CГC,Pë P°PSPiC-PsC,PμPSP·PëPSCр II C,P° P±P»PsPeP°C,PsCЦPë PeP°P»CHC†C-C”PIPëC... PeP°PSP°P»C-PI. PђP»PjPμCГP°CЦC,P°PSCр PjPμPrPsPeCГPsPjC-P» C- P°PjP»PsPrPëPïC-PS. PьPsPr PђPŸPГ PŸ09D P’02.

PrP°CЦPjP°PePsP»PsPiC-C±PSC- PIP»P°CГC,PëPIPsCГC,C-.

PrP°CЦPjP°PePsPrPëPSP°PjC-PeP°.

PŸPμPIC-PeP°CЦ вђ” C†Pμ PePsPjP±C-PSPsPIP°PSPëPNë P»C-PeP°CЦCГCЦPëPëPNë P·P°CГC-P±, PrPs CГPeP»P°PrCр CЦPePsPiPs PIC...PsPrPëC,CH PsP»PjPμCГP°CЦC,P°PSCр PjPμPrPsPeCГPsPjC-P» вђ” P°PSC,P°PiPsPSC-CГC, CЦPμC†PμPïC,PsCЦC-PI P°PSPiC-PsC,PμPSP·PëPSCр P†P† вђ” C,P° P°PjP»PsPrPëPïC-PSCр P±PμCГPëP»P°C, вђ” P±P»PsPeP°C,PsCЦ PeP°P»CHC†C-C”PIPëC... PeP°PSP°P»C-PI. PьPsPjP±C-PSP°C†C-CЦ C†PëC... PrPIPsC... PrC-CЦC±PëC... CЦPμC±PsPIPëPS PIPëCЦPIP»CЦC” CГPëPSPμCЦPiC-C±PSPëPNë PμC,,PμPeC, C,P° CГPïCЦPëCЦC” P·PSPëP¶PμPSPSCЦ P°CЦC,PμCЦC-P°P»CHPSPsPiPs C,PëCГPeCр P±C-P»CHC€PsCЦ PjC-CЦPsCЦ, PSC-P¶ PePsP¶PSP° PrC-CЦC±P° CЦPμC±PsPIPëPSP° PsPeCЦPμPjPs.

Pj 8-C,PëP¶PSPμPIPsPjCр PïPsPrPIC-PNëPSPs CГP»C-PïPsPjCр, CЦP°PSPPrPsPjC-P·PsPIP°PSPsPjCр, PïP»P°C†PμP±Ps-PePsPSC,CЦPsP»CHPSPiP°PSPsPjCр C,,P°PeC,PsCЦPSPsPjCр PrPsCГP»C-PrP¶PμPSPSC- P·P° CрC±P°CГC,CЦ 1940 PïP°C†C-C”PSC,C-PI (71 % PïP°C†C-C”PSC,C-PI PSP°P»PμP¶P°P»Pë PrPs C”PICЦPsPïPμPsC—PrPSPsC— CЦP°CГPë C- 29 % – PrPs C-PSC€PëC... CЦP°CГ) P±CрP»Ps PïPsPeP°P·P°PSPs, C%Ps P»C-PeCрPIP°PSPSCЦ PïCЦPμPïP°CЦP°C,PsPj PŸPμPIC-PeP°CЦ PïCЦPëP·PIPsPrPëP»Ps PrPs C-CГC,PsC,PSPs P±C-P»CHC€ PIPëCЦP°P¶PμPSPsPiPs P·PSPëP¶PμPSPSCЦ PrC-P°CГC,PsP»C-C±PSPsPiPs C,P° CГPëCГC,PsP»C-C±PSPsPiPs C,PëCГPeCр PI PïPsCЦC-PIPSCЦPSPSC- P· PjPsPSPsC,PμCЦP°PïC-C”CЦ PIC-PrPïPsPIC-PrPSPëPjPë PePsPjPïPsPSPμPSC,P°PjPë.

PŸPµCŤPµPrPSC" P·PSPëP¶PµPSPSCŪ

CÍPëCÍC,PsP»C-C‡PSPsPiPs/PrC-P°CÍC,PsP»C-C‡PSPsPiPs C,PëCÍPeCÍ
C...P°CŤP°PeC,PµCŤPëP·CÍPIP°P»PsCÍCŪ P·P°P»PµP¶PSC-CÍC,CŤ PIC-Pr PrPsP·Pë:
-24/-14 PjPj CŤC,. CÍC,. (PrP»CŪ C,,C-PeCÍPsPIP°PSPsC— PePsPjP±C-PSP°C†C-C—
20 PjPi/5 PjPi), -25/-16 PjPj CŤC,. CÍC,. (PrP»CŪ C,,C-PeCÍPsPIP°PSPsC—
PePsPjP±C-PSP°C†C-C— 40 PjPi/5 PjPi) C- -30/-19 PjPj CŤC,. CÍC,. (PrP»CŪ
C,,C-PeCÍPsPIP°PSPsC— PePsPjP±C-PSP°C†C-C— 40 PjPi/10 PjPi).

PŸPµPIC-PeP°CŤ, 40 PjPi/5 PjPi, P·PSPëP¶CÍPIP°PI
CÍPëCÍC,PsP»C-C‡PSPëPNë/PrC-P°CÍC,PsP»C-C‡PSPëPNë C,PëCÍPe CÍ
PİPsP»PsP¶PµPSPSC- PİP°C†C-C" PSC,P° CÍPëPrCŪC‡Pë PrPsPrP°C,PePsPIPs PSP°
2,5/1,7 PjPj CŤC,. CÍC,. Pİ PİPsCŤC-PIPSCŪPSPSC- P· PŸPµPIC-PeP°CŤPsPj,
20 PjPi/5 PjPi. PŤPSP°P»PsPiC-C‡PSPs PŸPµPIC-PeP°CŤ, 40 PjPi/10 PjPi,
P·PSPëP¶CÍPIP°PI CÍPëCÍC,PsP»C-C‡PSPëPNë/PrC-P°CÍC,PsP»C-C‡PSPëPNë C,PëCÍPe CÍ
PİPsP»PsP¶PµPSPSC- PİP°C†C-C" PSC,P° CÍPëPrCŪC‡Pë PrPsPrP°C,PePsPIPs PSP°
4,7/3,5 PjPj CŤC,. CÍC,. CÍ PİPsCŤC-PIPSCŪPSPSC- P· PŸPµPIC-PeP°CŤPsPj,
40 PjPi/5 PjPi.

PŞP°CÍC,PeP° PİP°C†C-C" PSC,C-PI, CÍ CŪPePëC... PIPrP°P»PsCÍCŪ PrPsCÍCŪPiC,Pë
C†C-P»CŤPsPIPëC... P·PSP°C‡PµPSCŤ P°CŤC,PµCŤC-P°P»CŤPSPsPiPs C,PëCÍPeCÍ
(< 140/90 PjPj CŤC,. CÍC,. CÍ PİP°C†C-C" PSC,C-PI P±PµP· C†CÍPeCŤPsPIPsPiPs
PrC-P°P±PµC,CÍ C- < 130/80 PjPj CŤC,. CÍC,. CÍ PrC-P°P±PµC,PëPeC-PI), CÍPeP»P°P»P°
42,5 %, 51,0 % C- 49,1 % PİCŤPë P·P°CÍC,PsCÍCÍPIP°PSPSC- PİCŤPµPİP°CŤP°C,CÍ
PŸPµPIC-PeP°CŤ CÍ PrPsP·CÍPIP°PSPSC- 20 PjPi/5 PjPi, 40 PjPi/5 PjPi C- 40 PjPi/10 PjPi
PIC-PrPİPsPIC-PrPSPs.

PŤCÍPSPsPIPSPëPNë PiC-PİPsC,PµPSP·PëPIPSPëPNë PµC,,PµPeC, PİCŤPµPİP°CŤP°C,CÍ
PŸPµPIC-PeP°CŤ, CŪPe PİCŤP°PIPëP»Ps, PrPsCÍCŪPiP°PICÍCŪ PİCŤPsC,CŪPiPsPj
PİPµCŤC€PëC... 2 C,PëP¶PSC-PI C,PµCŤP°PİC-C—.

PJ PrCŤCÍPiPsPjCÍ PİPsPrPIC-PNëPSPs CÍP»C-PİPsPjCÍ,
CŤP°PSPrPsPjC-P·PsPIP°PSPsPjCÍ, PİP»P°C†PµP±Ps-PePsPSC,CŤPsP»CŤPsPIP°PSPsPjCÍ
PrPsCÍP»C-PrP¶PµPSPSC- PİPëPIC‡P°P»P°CÍCŪ PµC,,PµPeC,PëPIPS-CÍC,CŤ
PrPsPrP°PIP°PSPSCŪ P°PjP»PsPrPëPİC-PSCÍ PrPs CÍC...PµPjPë P»C-PeCÍPIP°PSPSCŪ
PİP°C†C-C" PSC,C-PI C" PİCŤPsPİPµPsC—PrPSPsC— CŤP°CÍPë P·
PSPµPrPsCÍC,P°C,PSCŤPsCŤ CŤPµP°PeC†C-C" CŤ PsCŤPİP°PSC-P·PjCÍ PSP°
PjPsPSPsC,PµCŤP°PİC-CŤ PsP»PjPµCÍP°CŤC,P°PSCÍ PjPµPrPsPeCÍPsPjC-P»PsPj Pİ
PrPsP·C- 20 PjPi PİCŤPsC,CŪPiPsPj 8 C,PëP¶PSC-PI.

PJ PİP°C†C-C" PSC,C-PI, CŪPeC- PİCŤPsPrPsPIP¶CÍPIP°P»Pë PsC,CŤPëPjCÍPIP°C,Pë
C,C-P»CŤPePë PsP»PjPµCÍP°CŤC,P°PSCÍ PjPµPrPsPeCÍPsPjC-P» CÍ PrPsP·C- 20 PjPi,
CÍPëCÍC,PsP»C-C‡PSPëPNë/PrC-P°CÍC,PsP»C-C‡PSPëPNë C,PëCÍPe P·PSPëP·PëPICÍCŪ
PSP° 10,6/7,8 PjPj CŤC,. CÍC,. PİCŤPsC,CŪPiPsPj PSP°CÍC,CÍPİPSPëC... 8 C,PëP¶PSC-PI.
PµCŤPë PrPsPrP°PIP°PSPSC- 5 PjPi P°PjP»PsPrPëPİC-PSCÍ P·P° 8 C,PëP¶PSC-PI
PIPrP°P»PsCÍCŪ PrPsCÍCŪPiC,Pë P·PSPëP¶PµPSPSCŪ
CÍPëCÍC,PsP»C-C‡PSPsPiPs/PrC-P°CÍC,PsP»C-C‡PSPsPiPs C,PëCÍPeCÍ PSP°
-16,2/-10,6 PjPj CŤC,. CÍC,. (CŤ = 0,0006). PŞP°CÍC,PeP° PİP°C†C-C" PSC,C-PI, CÍ
CŪPePëC... PIPrP°P»PsCÍCŪ PrPsCÍCŪPiC,Pë C†C-P»CŤPsPIPëC... P·PSP°C‡PµPSCŤ

P°CĤC,PμCĤC-P°P»CHΨPSPsPiPs C,PëCÍPeCÍ (< 140/90 PjPj CĤC,.. CÍC,.. CÍ PĪP°C†C-C"ΨC,C-PI P±PμP· C†CÍPeCĤCΨsPiPs PrC-P°P±PμC,CÍ C- < 130/80 PjPj CĤC,.. CÍC,.. CÍ PrC-P°P±PμC,PëPeC-PI), CÍPeP»P°P»P° 44,5 % PrP»CÍ PePsPjP±C-PSP°C†C-C— 20 PjPi/5 PjPi PI PĪPsCĤC-PIΨCΨPSPSC- P· 28,5 % PrP»CÍ 20 PjPi PsP»PjPμCÍP°CĤC,P°PSCÍ PjPμPrPsPeCÍPsPjC-P»CÍ.

P' C...PsPrC- PĪPsPrP°P»CHΨC€PëC... PrPsCÍP»C-PrP¶PμPSCĤ PsC†C-PSCĤPIP°P»Pë PμC,,PμPeC,PëPIΨC-CÍC,CH PrPsPrP°PIP°PSPSCΨ CĤC-P·PSPëC... PrPsP· PsP»PjPμCÍP°CĤC,P°PSCÍ PjPμPrPsPeCÍPsPjC-P»CÍ PrPs CÍC...PμPjPë P»C-PeCÍPIP°PSPSCΨ PĪP°C†C-C"ΨC,C-PI C"PICĤCΨsPiPμPsC—PrPSPsC— CĤP°CÍPë P· PSPμPrPsCÍC,P°C,PSCĤPsCĤ CĤPμP°PeC†C-C"Ĥ PsCĤPiP°PSC-P·PjCÍ PSP° PjPsPSPsC,PμCĤP°PiC-CĤ P°PjP»PsPrPëPiC-PSPsPj PI PrPsP·C- 5 PjPi PiCĤPsC,CΨPiPsPj 8 C,PëP¶PSC-PI.

Pj PĪP°C†C-C"ΨC,C-PI, CΨPeC- PiCĤPsPrPsPIP¶CÍPIP°P»Pë PsC,CĤPëPjCÍPIP°C,Pë C,C-P»CHΨPePë P°PjP»PsPrPëPiC-PSCÍ 5 PjPi, CÍPëCÍC,PsP»C-C†PSPëPNë/PrC-P°CÍC,PsP»C-C†PSPëPNë C,PëCÍPe P·PSPëP·PëPICÍCΨ PSP° 9,9/5,7 PjPj CĤC,.. CÍC,.. PiCĤPsC,CΨPiPsPj PSP°CÍC,CÍPiPSPëC... 8 C,PëP¶PSC-PI. P"PsPrP°PIP°PSPSCΨ 20 PjPi PsP»PjPμCÍP°CĤC,P°PSCÍ PjPμPrPsPeCÍPsPjC-P»CÍ PiCĤPëP·PIPμP»Ps PrPs P·PSPëP¶PμPSPSCΨ CÍPëCÍC,PsP»C-C†PSPsPiPs/PrC-P°CÍC,PsP»C-C†PSPsPiPs P°CĤC,PμCĤC-P°P»CHΨPSPsPiPs C,PëCÍPeCÍ PSP° B B B B B B B B B B B B B B B B -15,3/-9,3 PjPj CĤC,.. CÍC,.. PrPsPrP°PIP°PSPSCΨ 40 PjPi PsP»PjPμCÍP°CĤC,P°PSCÍ PjPμPrPsPeCÍPsPjC-P»CÍ - PSP° -16,7/-9,5 PjPj CĤC,.. CÍC,.. (CĤ < 0,0001).

P§P°CÍC,PeP° PĪP°C†C-C"ΨC,C-PI, CÍ CΨPePëC... PIPrP°P»PsCÍCΨ PrPsCÍCΨPiC,Pë C†C-P»CHΨsPiPëC... P·PSP°C†PμPSCĤ P°CĤC,PμCĤC-P°P»CHΨPSPsPiPs C,PëCÍPeCÍ (< 140/90 PjPj CĤC,.. CÍC,.. CÍ PĪP°C†C-C"ΨC,C-PI P±PμP· C†CÍPeCĤCΨsPiPs PiPs PrC-P°P±PμC,CÍ C- < 130/80 PjPj CĤC,.. CÍC,.. CÍ PrC-P°P±PμC,PëPeC-PI), CÍPeP»P°P»P° 29,9 % PI PiCĤCÍPiC- PjPsPSPsC,PμCĤP°PiC-C— P°PjP»PsPrPëPiC-PSPsPj CÍ PrPsP·CÍPIP°PSPSC- 5 PjPi, 53,5 % CÍ PiCĤCÍPiC- P·P°CÍC,PsCÍCÍPIP°PSPSCΨ PŸPμPIC-PeP°CĤCÍ, 20 PjPi/5 PjPi, C- 50,5 % CÍ PiCĤCÍPiC- PŸPμPIC-PeP°CĤCÍ, 40 PjPi/5 PjPi.

P P°PSPrPsPjC-P·PsPIP°PSC- PrP°PSC- C%PsPrPs PĪP°C†C-C"ΨC,C-PI P· PSPμPePsPSC,CĤPsP»CHΨsPiP°PSPsCĤ P°CĤC,PμCĤC-P°P»CHΨPSPsCĤ PiC-PĪPμCĤC,PμPSP·C-C"Ĥ, C%Ps PrP°CĤC,CH P·PjPsPiCÍ PĪPsCĤC-PIΨCΨC,Pë CĤPμP·CÍP»CHΨC,P°C,Pë PePsPjP±C-PSPsPIP°PSPsC— C,PμCĤP°PiC-C— PiCĤPμPiP°CĤP°C,PsPj PŸPμPIC-PeP°CĤ PI CÍPμCĤPμPrPSC-C... PrPsP·P°C... P· PμC,,PμPeC,PsPj PIC-Pr PiC-PrPIPëC%PμPSPSCΨ PrPsP·Pë P°PjP»PsPrPëPiC-PSCÍ C,P° PsP»PjPμCÍP°CĤC,P°PSCÍ PiCĤPë PjPsPSPsC,PμCĤP°PiC-C—, PIC-PrCÍCÍC,PSC-.

P PμP·CÍP»CHΨC,P°C,Pë C,CĤCHΨPsC... PrPsCÍP»C-PrP¶PμPSCĤ PiC-PrC,PIPμCĤPrP¶CÍCĤC,CH, C%Ps PiC-PĪPsC,PμPSP·PëPIΨPëPNë PμC,,PμPeC, PiCĤPμPiP°CĤP°C,CÍ PŸPμPIC-PeP°CĤ PiCĤPë C†P°CÍC,PsC,C- PiCĤPëPNëPsPjCÍ PsPrPëPS CĤP°P· PSP° PrPsP±CÍ P·P±PμCĤC-PiP°PICÍCΨ PiCĤPë C-PSC,PμCĤPIP°P»C- PjC-P¶ PiCĤPëPNëPsPjP°PjPë 24 PiPsPrPëPSPë, PiCĤPë C†CHΨPsPjCÍ

СГРІС-PIPIC-PrPSPsC€PµPSPSCЦ PjC-P¶ PjC-PSC-PjP°P»CHΨPSPëPj C-
PjP°PeCГPëPjP°P»CHΨPSPëPj P·PSP°C‡PµPSPSCЦPjPë CГPëCГC,PsP»C-C‡PSPsPiPs C-
PrC-P°CГC,PsP»C-C‡PSPsPiPs C,PëCГPeC PIP°CГC-CГPIP°P»Ps PIC-Pr 71 PrPs 82 %.
P·C,,PµPeC,PëPIPSC-CГC,CH PïCГPµPïP°CГP°C,Cr PïCГPsC,CЦPiPsPj PrPsP±Pë
P±CfP»P° PïC-PrC,PIPµCГPrP¶PµPSP° PïCГPë P°PjP±CfP»P°C,PsCГPSPsPjCf
PjPsPSC-C,PsCГPëPSPiCf P°CГC,PµCГC-P°P»CHΨPSPsPiPs C,PëCГPeCf.

P°C-PïPsC,PµPSP·PëPIPSPëPNë PµC,,PµPeC, PïCГPµPïP°CГP°C,Cr PŸPµPIC-PeP°CГ PSPµ
P·P°P»PµP¶P°PI PIC-Pr PIC-PeCf C- CГC,P°C,C- C...PIPsCГPsPiPs, P° C,P°PePsP¶ PIC-Pr
PSP°CЦPIPSPsCГC,C- Cr PïP°C†C-C"PSC,P° C†CfPeCГPsPiPsPiPs PrC-P°P±PµC,Cf.

PJ PrPIPsC... PIC-PrPeCГPëC,PëC... PSPµCГP°PSPPrPsPjC-P·PsPIP°PSPëC...
CГPsP·C€PëCГPµPSPëC... PrPsCГP»C-PrP¶PµPSPSCЦC... CГC,C-PNëPeP°
PµC,,PµPeC,PëPIPSC-CГC,CH PïCГPµPïP°CГP°C,Cr PŸPµPIC-PeP°CГ, 40 PjPi/5 PjPi,
P±CfP»P° PïPsPeP°P·P°PSP° Cr 49-67 % PïP°C†C-C"PSC,C-PI C‡PµCГPµP· PsPrPëPS
CГC-Pe P·P°CГC,PsCГCfPIP°PSPSCЦ.

*PħP»PjPµCГP°CГC,P°PSCz PjPµPrPsPeCГPsPjC-P» (PrC-CГC‡P° CГPµC‡PsPIPëPSP°
PŸPµPIC-PeP°CГCz)*

PħP»PjPµCГP°CГC,P°PSCr PjPµPrPsPeCГPsPjC-P», C%Ps PIC...PsPrPëC,CH PrPs
CГPeP»P°PrCf PïCГPµPïP°CГP°C,Cr PŸPµPIC-PeP°CГ, C" CГPµP»PµPeC,PëPIPSPëPj
P°PSC,P°PiPsPSC-CГC,PsPj CГPµC†PµPïC,PsCГC-PI P°PSPiC-PsC,PµPSP·PëPSCr II
PïPµCГC€PsPiPs C,PëPïCf (AT₁). P' PsCГPïP°PSC-P·PjC- PsP»PjPµCГP°CГC,P°PSCr
PjPµPrPsPeCГPsPjC-P» C€PIPëPrPePs PïPµCГPµC,PIPsCГCГC"C,CHCГCЦ Cr
C,,P°CГPjP°PePsP»PsPiC-C‡PSPs P°PeC,PëPIPSPëPNë PjPµC,P°P±PsP»C-C,
PsP»PjPµCГP°CГC,P°PS. PħPSPiC-PsC,PµPSP·PëPS P†P† BГ " C†Pµ PïPµCГPIPëPSPSPëPNë
PIP°P·PsP°PeC,PëPIPSPëPNë PiPsCГPjPsPS CГPµPSC-PS-P°PSPiC-PsC,PµPSP·PëPS-
P°P»CHPrPsCГC,PµCГPsPSPsPIPsC— CГPëCГC,PµPjPë, C%Ps PIC-PrC-PiCГP°C"
PIP°P¶P»PëPICr CГPsP»CH Cr PïP°C,PsC,,C-P·C-PsP»PsPiC-C—
P°CГC,PµCГC-P°P»CHΨPSPsC— PiC-PïPµCГC,PµPSP·C-C—. PħPSPiC-PsC,PµPSP·PëPS
P†P† PïCГPëP·PIPsPrPëC,CH PrPs P·PICrP¶PµPSPSCЦ CГCfPrPëPS,
CГC,PëPjCfP»CЦC†C-C— CГPëPSC,PµP·Cr C,P° PIPëPIC-P»CHΨPSPµPSPSCЦ
P°P»CHPrPsCГC,PµCГPsPSCr, PeP°CГPrC-PsCГC,PëPjCfP»CЦC†C-C— C,P°
PSPëCГPePsPIPsC— CГPµP°P±CfPsCГP±C†C-C— PSP°C,CГC-CГ.
PħP»PjPµCГP°CГC,P°PS PïCГPëPiPSC-C‡CfC" CГCfPrPëPSPsP·PICrP¶CfPIP°P»CHΨPSCr
C,P° P°P»CHPrPsCГC,PµCГPsPSPsCГPµPeCГPµC,CfCГC‡Cf PrC-CГ
P°PSPiC-PsC,PµPSP·PëPSCr P†P† C€P»CЦC...PsPj P±P»PsPeCfPIP°PSPSCЦ
CГPµC†PµPïC,PsCГC-PI AT₁ Cr C,PeP°PSPëPSP°C..., Cr C,PsPjCf C‡PëCГP»C- PI
PiP»P°PrPePëC... PjBГ™ CЦP·P°C... CГCfPrPëPS C,P° PSP°PrPSPëCГPePsPIPëC...
P·P°P»PsP·. P°C-CЦ PsP»PjPµCГP°CГC,P°PSCr PSPµ P·P°P»PµP¶PëC,CH PIC-Pr
PrP¶PµCГPµP»P° C- C€P»CЦC...Cr CГPëPSC,PµP·Cr P°PSPiC-PsC,PµPSP·PëPSCr P†P†.
P'PëP±C-CГPePsPIPëPNë P°PSC,P°PiPsPSC-P·Pj CГC,PsCГPsPIPs
CГPµC†PµPïC,PsCГC-PI AT₁ P°PSPiC-PsC,PµPSP·PëPSCr P†P† PïCГPëP·PIPsPrPëC,CH
PrPs P·P±C-P»CHC€PµPSPSCЦ CГC-PIPsCЦ CГPµPSC-PSCr Cr PïP»P°P·PjC-
PeCГPsPIC- C,P° PePsPSC†PµPSC,CГP°C†C-C— P°PSPiC-PsC,PµPSP·PëPSCr P† C-
P°PSPiC-PsC,PµPSP·PëPSCr P†P†, P° C,P°PePsP¶ PrPs PrPµCЦPePsPiPs
P·PjPµPSC€PµPSPSCЦ CГC-PIPsCЦ P°P»CHPrPsCГC,PµCГPsPSCr Cr PïP»P°P·PjC-

PeC7PsPIC-. P4C7Pë P°C7C,P4C7C-P°P»CH7PSC-PNë PiC-PiP4C7C,P4PSP·C-C—
PsP»PjP4C7P°C7C,P°PSCr PjP4PrPsPeC7PsPjC-P» C7PiC7PëC7PëPSC7C”
C,C7PëPIP°P»P4 P·PSPëP4P4PSPSC4 P°C7C,P4C7C-P°P»CH7PSPsPiPs C,PëC7PeC7,
C%0Ps P·P°P»P4P4PëC,CH PIC-Pr PrPsP·Pë.

P4C7Pë PNëPsPiPs P·P°C7C,PsC7C7PIP°PSPSC- PSP4 PIC-PrPjC-C7P°P»PsC7C4
C7PsP·PIPëC,PëC7 P°C7C,P4C7C-P°P»CH7PSPsC— PiC-PiPsC,P4PSP·C-C— PiC-C7P»C4
PiC7PëPNëPsPjC7 PiP4C7C€PsC— PrPsP·Pë, PsP·PSP°Pe C,P°C...C-C,,C-P»P°PeC7C-C—
PiC7Pë C,C7PëPIP°P»PsPjC7 P·P°C7C,PsC7C7PIP°PSPSC- C,P° C7P4C7PëPrPëPIC7
P°C7C,P4C7C-P°P»CH7PSPsC— PiC-PiP4C7C,P4PSP·C-C— PiC-C7P»C4
PiC7PëPiPëPSP4PSPSC4 P·P°C7C,PsC7C7PIP°PSPSC4.

P4C7Pë P·P°C7C,PsC7C7PIP°PSPSC- PsP»PjP4C7P°C7C,P°PSCr PjP4PrPsPeC7PsPjC-P»C7
C...PIPsc7PëPj PSP° P°C7C,P4C7C-P°P»CH7PSCr PiC-PiP4C7C,P4PSP·C-C7 1 C7P°P·
PSP° PrPsP±C7 PIC-PrP±C7PIP°C”C,CH7C7C4 P4C,,P4PeC,PëPIPSP4 C,P° PiP»P°PIPSP4
P·PSPëP4P4PSPSC4 P°C7C,P4C7C-P°P»CH7PSPsPiPs C,PëC7PeC7 PiC7PsC,C4PiPsPj 24-
PiPsPrPëPSPSPsPiPs C-PSC,P4C7PIP°P»C7 PjC-P4 P·P°C7C,PsC7C7PIP°PSPSC4PjPë.

P C-PIP4PSC7 P°PSC,PëPiC-PiP4C7C,P4PSP·PëPIPSPsC— PrC-C— PiC7Pë
P·P°C7C,PsC7C7PIP°PSPSC- PiC7P4PiP°C7P°C,C7 1 C7Pë 2 C7P°P·Pë PSP° PrPsP±C7 PI
PsPrPSC-PNë C- C,C-PNë C7P°PjC-PNë PrPsP±PsPIC-PNë PrPsP·C- P±C7PI
PsPrPSP°PePsPIPëPNë. P7P°PeC7PëPjP°P»CH7PSP4 P·PSPëP4P4PSPSC4
P°C7C,P4C7C-P°P»CH7PSPsPiPs C,PëC7PeC7 PrPsC7C4PiP°C”C,CH7C7C4 C7P4C7P4P·
8 C,PëP4PSC-PI PiC-C7P»C4 PiPsC7P°C,PëC7 P»C-PëC7PIP°PSPSC4, C...PsC7P°
C7C7C,C,C”PIPëPNë PiC-PiPsC,P4PSP·PëPIPSPëPNë P4C,,P4PeC,
C7PiPsC7C,P4C7C-PiP°C”C,CH7C7C4 PIP4P4 C7P4C7P4P· 2 C,PëP4PSC-
P»C-PëC7PIP°PSPSC4.

P’PiP»PëPI PsP»PjP4C7P°C7C,P°PSCr PjP4PrPsPeC7PsPjC-P»C7 PSP°
P·P°C...PIPsC7C7PIP°PSC-C7C,CH C,P° C7PjP4C7C,PSC-C7C,CH PSP4
PIC7C,P°PSPsPIP»P4PSPëPNë.

P P°PSPPrPsPjC-P·PsPIP°PSP4 PrPsC7P»C-PrP4P4PSPSC4 P·P°C7C,PsC7C7PIP°PSPSC4
PsP»PjP4C7P°C7C,P°PSCr PrP»C4 PiC7PsC,,C-P»P°PeC,PëPePë
PrC-P°P±P4C,PëC7PSPsC— PjC-PëC7PsP°P»CH7P±C7PjC-PSC7C7C— (ROADMAP) C-P·
P·P°P»C7C7P4PSPSC4Pj 4447 PiP°C7C-C”PSC,C-PI C-P· C7C7PeC7PsPIPëPj
PrC-P°P±P4C,PsPj 2 C,PëPiC7 P· PSPsC7PjP°P»CH7PSPëPj C7C-PIPSP4Pj
P°P»CH7P±C7PjC-PSC7C7C— C- C4Pe PjC-PSC-PjC7Pj PsPrPSPëPj
PrPsPrP°C,PePsPIPëPj C,,P°PeC,PsC7PsPj C7PëP·PëPeC7 PIPëPSPëPePSP4PSPSC4
C7P4C7C7P4PIPsc-C7C7PrPëPSPSPëC... P·P°C...PIPsC7C7PIP°PSC7,
PiC7PsPiPsPrPëP»PsC7C4 P· PjP4C,PsC7 P·B7™ C4C7C7PIP°C,Pë, C7Pë PjPsP4P4
C,P4C7P°PiC-C4 PsP»PjP4C7P°C7C,P°PSPsPj P·P°C,C7PëPjP°C,Pë C7P°C7 PiPsC4PIPë
PjC-PëC7PsP°P»CH7P±C7PjC-PSC7C7C— P4C-Pr C7P°C7 C7P4C7P4PrPSC7PsPiPs
PiP4C7C-PsPrC7 PIC-PrC7C,P4P4P4PSPSC4 C,C7PëPIP°P»C-C7C,C7 3,2 C7PsPeC7
PiP°C7C-C”PSC,Pë PsC,C7PëPjC7PIP°P»Pë PsP»PjP4C7P°C7C,P°PS P°P±Ps
PiP»P°C7P4P±Ps PSP° PrPsPrP°C,PsPe PrPs C-PSC€PëC...
P°PSC,PëPiC-PiP4C7C,P4PSP·PëPIPSPëC... P·P°C7PsP±C-PI, P·P° PIPëPSC4C,PePsPj
C-PSPiC-P±C-C,PsC7C-PI P7P4P7 P°P±Ps P’P P7.

PJ PīPμCḡPIPēPSPSC-PNē PeC-PSC†PμPIC-PNē C,PsC‡C†C- PrPsCĠP»C-PrP¶PμPSPSCĬ PīCḡPsPrPμPjPsPSCĠC,CḡCḡPIP°P»Ps P·PSP°C‡PSPμ P·PSPēP¶PμPSPSCĬ CḡPēP·PēPeCḡ C%oPsPrPs C‡P°CĠCḡ PīPsCĬPIPē PjC-PeCḡPsP°P»CḡP±CḡPjC-PSCḡCḡC-C— PSP° PePsCḡPēCĠC,Cḡ PsP»PjPμCĠP°CḡC,P°PSCḡ. PμC-CĠP»CĬ PePsCḡPēPiCḡPIP°PSPSCĬ PIC-PrPjC-PSPSPsCĠC,PμPNē Cḡ PīPsPeP°P·PSPēPeP°C... P°CḡC,PμCḡC-P°P»CḡPSPsPiPs C,PēCĠPeCḡ (PḡPḡ) C†Pμ P·PSPēP¶PμPSPSCĬ CḡPēP·PēPeCḡ PīPμCḡPμCĠC,P°P»Ps P±CḡC,Pē CḡC,P°C,PēCḡC,PēC‡PSPs P·PSP°C‡CḡC%oPēPj. PJ 8,2 % (Cḡ 178 P· 2160) PīP°C†C-C”PSC,C-PI Cḡ PiCḡCḡPiC- PsP»PjPμCĠP°CḡC,P°PSCḡ C- Cḡ 9,8 % (Cḡ 210 P· 2139) Cḡ PiCḡCḡPiC- PīP»P°C†PμP±Ps CḡPsP·PIPēPSCḡP»P°CĠCĬ PjC-PeCḡPsP°P»CḡP±CḡPjC-PSCḡCḡC-CĬ.

PJ PIC,PsCḡPēPSPSC-PNē PeC-PSC†PμPIC-PNē C,PsC‡C†C- CĠPμCḡC†PμPIPs- CĠCḡPrPēPSPSC- CĬPIPēC%oP° PIC-PrP·PSP°C‡P°P»PēCĠCĬ Cḡ 96 PīP°C†C-C”PSC,C-PI (4,3 %), CĬPeC- PsC,CḡPēPjCḡPIP°P»Pē PsP»PjPμCĠP°CḡC,P°PS, C- Cḡ 94 PīP°C†C-C”PSC,C-PI (4,2 %), CĬPeC- PsC,CḡPēPjCḡPIP°P»Pē PīP»P°C†PμP±Ps. PḡPjPμCḡC,PSC-CĠC,Cḡ PIC-Pr CĠPμCḡC†PμPIPs-CĠCḡPrPēPSPSPēC... P·P°C...PIPsCḡCḡPIP°PSCḡ P±CḡP»P° PIPēC%oPsCḡ PI PiCḡCḡPiC- PsP»PjPμCĠP°CḡC,P°PSCḡ PI PīPsCḡC-PIPsCĬPSPSC- P· PiCḡCḡPīPsCḡ PīP»P°C†PμP±Ps (15 PīP°C†C-C”PSC,C-PI (0,7 %) C- 3 PīP°C†C-C”PSC,Pē (0,1 %)), PSPμP·PIP°P¶P°CḡC‡Pē PSP° PīPsPrC-P±PSCḡ C‡P°CĠC,PsC,Cḡ PIPēPSPēPePSPμPSPSCĬ C-PSCḡCḡP»CḡC,Cḡ P±PμP· P»PμC,P°P»CḡPSPsPiPs PSP°CĠP»C-PrPeCḡ (14 PīP°C†C-C”PSC,C-PI (0,6 %) C- 8 PīP°C†C-C”PSC,C-PI (0,4 %)), C-PSC,,P°CḡPeC,Cḡ PjC-PsPeP°CḡPrP° P±PμP· P»PμC,P°P»CḡPSPsPiPs PSP°CĠP»C-PrPeCḡ (17 PīP°C†C-C”PSC,C-PI (0,8 %) C- 26 PīP°C†C-C”PSC,C-PI (1,2 %)) C,P° PīPsPeP°P·PSPēPeP° CĠPjPμCḡC,PSPsCĠC,C-, PSPμ PīPsPiBḡ™ CĬP·P°PSPsC— P· CĠPμCḡC†PμPIPs-CĠCḡPrPēPSPSPēPjPē PīCḡPēC‡PēPSP°PjPē (11 PīP°C†C-C”PSC,C-PI (0,5 %) C- 12 PīP°C†C-C”PSC,C-PI (0,5 %)). P—P°PīP°P»CḡPSP° CĠPjPμCḡC,PSC-CĠC,Cḡ Cḡ PiCḡCḡPiC- PsP»PjPμCĠP°CḡC,P°PSCḡ P±CḡP»P° PIPēC%oPsCḡ (26 PīP°C†C-C”PSC,C-PI (1,2 %) C- 15 PīP°C†C-C”PSC,C-PI (0,7 %)), PiPsP»PsPIPsPēPj C‡PēPSPsPj, P·P° CḡP°C...CḡPSPsPe PIPēC%oPsC— CĠPjPμCḡC,PSPsCĠC,C- P· CĠPμCḡC†PμPIPs-CĠCḡPrPēPSPSPēC... PīCḡPēC‡PēPS.

PJ PrPsCĠP»C-PrP¶PμPSPSC- ORIENT (The Olmesartan Reducing Incidence of End-stage Renal Disease in Diabetic Nephropathy Trial) PIPēPIC‡P°P»Pē PIPiP»PēPI PsP»PjPμCĠP°CḡC,P°PSCḡ PSP° CḡPμP·CḡP»CḡC,P°C, PSPēCḡPePsPIPēC... C- CĠPμCḡC†PμPIPs-CĠCḡPrPēPSPSPēC... P·P°C...PIPsCḡCḡPIP°PSCḡ Cḡ 577 CḡP°PSPPrPsPjC-P·PsPIP°PSPēC... PīP°C†C-C”PSC,C-PI PI PīPīPsPSC-C— C- PI PḡPēC,P°C— P· C†CḡPeCḡPsPIPēPj PrC-P°P±PμC,PsPj 2 C,PēPiCḡ C- PIPēCḡP°P¶PμPSPsCḡ PSPμC,,CḡPsPīP°C,C-C”Cḡ. PμC-Pr C‡P°CĠ CĠPμCḡPμPrPSCḡPsPiPs PīPμCḡC-PsPrCḡ PIC-PrCĠC,PμP¶PμPSPSCĬ C,CḡPēPIP°P»C-CĠC,Cḡ 3,1 CḡPsPeCḡ PīP°C†C-C”PSC,Pē PsC,CḡPēPjCḡPIP°P»Pē PsP»PjPμCĠP°CḡC,P°PS P°P±Ps PīP»P°C†PμP±Ps PSP° PrPsPrP°C,PsPe PrPs C-PSC€PēC... P°PSC,PēPiC-PīPμCḡC,PμPSP·PēPIPsPēC... P·P°CĠPsP±C-PI, Cḡ C,PsPjCḡ C‡PēCĠP»C- C-PSPiC-P±C-C,PsCḡC-PI PḡPμPḡ.

PμPμCḡPIPēPSPSP° PsP±Bḡ™ C”PrPSP°PSP° PeC-PSC†PμPIP° C,PsC‡PeP° (C‡P°CĠ PīPμCḡC€PsC— PīPsCĬPIPē PīPsPrPIPsC”PSPSCĬ CĠPēCḡPsPIP°C,PePsPIPsPiPs PeCḡPμP°C,PēPSC-PSCḡ, PSPēCḡPePsPIPμ P·P°C...PIPsCḡCḡPIP°PSPSCĬ PI

C, P_μ C₇ P_j C - P S P ° P » C H P S C - P N₂ C₁ C, P ° P r C - C —, C₁ P_j P_μ C₇ C, C H P · C₁ C₁ C - C... P_i C₇ P_e C † P_e P S) P ± C₁ P » P ° P r P s C₁ C₁ P_i P S C₁ C, P ° C₁ 116 P_i P ° C † C - C " P S C, C - P I C₁ P_i C₇ C₁ P_i C - P s P » P_j P_μ C₁ P ° C₇ C, P ° P S C₁ (41,1 %) C - C₁ 129 P_i P ° C † C - C " P S C, C - P I, C₁ P_e C - P s C, C₇ P_e P_j C₁ P_i P ° P » P_e P_i P » P ° C † P_μ P ± P s (45,4 %) (H R 0,97 (95 % P " P † P I C - P r 0,75 P r P s 1,24); p = 0,791). P ° C, P s C₇ P_e P S P S P ° P s P ± B₇ ° C " P r P S P ° P S P ° C₁ P_μ C₇ C † P_μ P I P s - C₁ C₁ P r P_e P S P S P ° P e C - P S C † P_μ P I P ° C, P s C † P e P ° P ± C₁ P » P ° P r P s C₁ C₁ P_i P S C₁ C, P ° C₁ 40 P_i P ° C † C - C " P S C, C - P I, C₁ P_e C - P s C, C₇ P_e P_j C₁ P_i P ° P » P_e P s P » P_j P_μ C₁ P ° C₇ C, P ° P S (14,2 %), C - C₁ 53 P_i P ° C † C - C " P S C, C - P I, C₁ P_e C - P s C, C₇ P_e P_j C₁ P_i P ° P » P_e P_i P » P ° C † P_μ P ± P s (18,7 %). P_i C₁ P s P ± B₇ ° C " P r P S P ° P S P ° C₁ P_μ C₇ C † P_μ P I P s - C₁ C₁ P r P_e P S P S P ° P e C - P S C † P_μ P I P ° C, P s C † P e P ° P I P e P » C H C † P ° P » P ° C₁ P_j P_μ C₇ C, P S C - C₁ C, C H P I C - P r C₁ P_μ C₇ C † P_μ P I P s - C₁ C₁ P r P_e P S P S P e C... P · P ° C... P I P s C₇ C₇ P I P ° P S C H C₁ C₁ C₁ 10 (3,5 %) P_i P ° C † C - C " P S C, C - P I, C₁ P_e C - P s C, C₇ P_e P_j C₁ P_i P ° P » P_e P s P » P_j P_μ C₁ P ° C₇ C, P ° P S, C - C₁ 3 (1,1 %) P_i P ° C † C - C " P S C, C - P I, C₁ P_e C - P s C, C₇ P_e P_j C₁ P_i P ° P » P_e P_i P » P ° C † P_μ P ± P s; P · P ° P_i P ° P » C H P S P e P N₂ P_i P s P e P ° P · P S P e P e C₁ P_j P_μ C₇ C, P S P s C₁ C, C - P r P s C₇ C - P I P S C H P I P ° P I 19 (6,7 %) C - 20 (7,0 %), C - P S C₁ C₁ P » C H C, P ± P_μ P · P » P_μ C, P ° P » C H P S P s P_i P s P S P ° C₁ P » C - P r P e C₁ - 8 (2,8 %) C, P ° 11 (3,9 %) C, P ° C - P S C, P ° C₇ P e C, P_j C - P s P e P ° C₇ P r P ° P ± P_μ P · P » P_μ C, P ° P » C H P S P s P_i P s P S P ° C₁ P » C - P r P e C₁ B₇ ° 3 (1,1 %) C - 7 (2,5 %) P I C - P r P_i P s P I C - P r P S P s.

P_h P_j P » P s P r P_e P_i C - P S (P r C - C H C † P ° C₇ P_μ C † P s P I P e P S P ° P_Y P_μ P I C - P e P ° C₇ C₁)

P_h P_j P » P s P r P_e P_i C - P S, C % P s P I C... P s P r P_e C, C H P r P s C₁ P e P » P ° P r C₁ P_i C₇ P_μ P_i P ° C₇ P ° C, C₁ P_Y P_μ P I C - P e P ° C₇, C " P ± P » P s P e P ° C, P s C₇ P s P_j P e P ° P » C H C † C - C " P I P e C... P e P ° P S P ° P » C - P I C - P_i P ° P » C H P_j C₁ C " C, C₇ P ° P S C₁ P_j P_μ P_j P ± C₇ P ° P S P S P e P N₂ P_i P_μ C₇ P_μ P S P s C₁ C - P s P S C - P I P e P ° P » C H C † C - C H C † P_μ C₇ P_μ P · P_i P s C, P_μ P S C † C - P ° P » P · P ° P » P_μ P P S C - P e P ° P S P ° P » P_e L - C, P_e P_i C₁ P I C₁ P_μ C₇ C † C - C - P_i P » P ° P r P e P_e C... P_j B₇ ° C₁ P · P ° C... P · P e C₁ P_i P_μ C₇ P_e P_j P_μ P S C, P ° P » C H P S C - P r P ° P S C - C₁ P I C - P r C † P ° C, C H, C % P s P ° P_j P » P s P r P_e P_i C - P S P I P · P ° C " P_j P s P r C - C " C₁ P e P · P r C - P » C₁ P S P e P ° P_j P_e P · P I B₇ ° C₁ P · C₁ P I P ° P S P S C₁ P r P_e P_i C - P r C₇ P s P_i C - C₇ P_e P r P_e P S C₁, C, P ° P e C - P · C - P S C € P_e P_j P_e P r C - P » C₁ P S P e P ° P_j P_e. P_h P_j P » P s P r P_e P_i C - P S P_j P ° C " P I C - P r P S P s C₁ P S C₁ P I P ° P · P s C₁ P_μ P » P_μ P e C, P_e P I P S C - C₁ C, C H C - P ± C - P » C H C € P_μ P I P_i P » P_e P I P ° C " P S P ° P e P » C - C, P_e P S P e P_i P » P ° P r P e P_e C... P_j B₇ ° C₁ P · C - P I C₁ C₁ P r P_e P S, P S C - P P P S P ° P e P ° C₇ P r C - P s P_j C - P s C † P_e C, P_e. P " C - P_i P s C, P_μ P S P · P_e P I P S P e P N₂ P_μ C, P_μ P e C, P ° P_j P » P s P r P_e P_i C - P S C₁ P · C₁ P_j P s P I P » P_μ P S P e P N₂ P_i C₇ C₁ P_j P s C₇ C₇ P s P · C₁ P » P ° P ± P » C H C H C † P s C₇ P r C - C " C₇ P S P ° P_i P » P ° P r P e P s P_j B₇ ° C₁ P · P s P I C - P e P » C - C, P_e P S P e P ° C₇ C, P_μ C₇ C - P N₂, C % P s C₁ P_i C₇ P_e C₁ C " P · P_j P_μ P S C € P_μ P S P S C₇ P_i P_μ C₇ P_e C, P_μ C₇ P_e C † P S P s P_i P s P s P_i P s C₇ C₁ C₁ C₁ P r P_e P S C -, P s C, P P P_μ, P · P S P e P P P_μ P S P S C₇ P ° C₇ C, P_μ C₇ C - P ° P » C H P S P s P_i P s C, P_e C₁ P e C₁.

P_μ C₇ P_e P ° C₇ C, P_μ C₇ C - P ° P » C H P S C - P N₂ P_i C - P_i P_μ C₇ C, P_μ P S P · C - C — P ° P_j P » P s P r P_e P_i C - P S P I P e P e P » P_e P e P ° C " C, C₇ P_e P I P ° P » P_μ P · P S P e P P P_μ P S P S C₁ P ° C₇ C, P_μ C₇ C - P ° P » C H P S P s P_i P s C, P_e C₁ P e C₁, P · P ° P » P_μ P P P S P_μ P I C - P r P r P s P · P_e. P P s P · P I P e C, P e C₁ P ° C₇ C, P_μ C₇ C - P ° P » C H P S P s C — P_i C - P_i P s C, P_μ P S P · C - C — P_i C - C₁ P » C₁ P_i C₇ P_e P N₂ P s P_j C₁ P_i P_μ C₇ C € P s C — P r P s P · P_e, P s P · P S P ° P e C, P ° C... C - C, C - P » P ° P e C₁ C - C — P_i C₇ P_e C, C₇ P_e P I P ° P » P s P_j C₁ P » C - P e C₁ P I P ° P S P S C - P ° P ± P s C₇ P_μ C † P_e P r P_e P I C₁ P ° C₇ C, P_μ C₇ C - P ° P » C H P S P s C — P_i C - P_i P_μ C₇ C, P_μ P S P · C - C — P_i C - C₁ P » C₁ P_i C₇ P_e P_i P_e P S P_μ P S P S C₁ P » C - P e C₁ P I P ° P S P S C₁ P S P_μ P I C - P r P · P S P ° C † P ° P » P s C₁ C₁.

PμC-CΓP»CΠ PiCηPēPNεPsPjCř PI C,PμCηP°PīPμPIC,PēC‡PSPēC... PrPsP·P°C... Cř
PīP°C†C-C"psc,c-pi P· P°CηC,PμCηC-P°P»CHbPSPsCη PiC-PīPμCηC,PμPSP·C-C"CH
P°PjP»PsPrPēPīC-PS P·P°P±PμP·PīPμC‡Cř" PμC,,PμPeC,PēPIPSPμ P·PSPēP¶PμPSPSCΠ
P°CηC,PμCηC-P°P»CHbPSPsPiPs C,PēCΓPeCř PI PīPsP»PsP¶PμPSPSCΠC...
PīP°C†C-C"psc,P° P»PμP¶P°C‡Pē, CΓPēPrCΠC‡Pē C- CΓC,PsCΠC‡Pē. PŷCηPēPIP°P»Pμ
P·P°CΓC,PsCΓCřPIP°PSPSCΠ P°PjP»PsPrPēPīC-PSCř PSPμ PīPsPIbη™ CΠP·P°PSPμ P·
C-CΓC,PsC,PSPēPjPē P·PjC-PSP°PjPē C‡P°CΓC,PsC,Pē CΓPμCηC†PμPIPēC...
CΓPePsCηPsC‡PμPSCη P°P±Ps CηC-PIPSCΠ PeP°C,PμC...PsP»P°PjC-PSC-PI Cř
PīP»P°P·PjC-. PJ PīP°C†C-C"psc,c-pi P· P°CηC,PμCηC-P°P»CHbPSPsCη
PiC-PīPμCηC,PμPSP·C-C"CH C,P° PSPsCηPjP°P»CHbPSPsCη C,,CřPSPeC†C-C"CH
PSPēCηPsPe P°PjP»PsPrPēPīC-PS PI C,PμCηP°PīPμPIC,PēC‡PSPēC... PrPsP·P°C...
P·PjPμPSC€CřPIP°PI PsPīC-Cη PSPēCηPePsPIPēC... CΓCřPrPēPS C-
PīC-PrPIPēC%oCřPIP°PI C€PIPēPrPeC-CΓC,CH PeP»CřP±PsC‡PePsPIPsC—
C,,C-P»CHbC,CηP°C†C-C— C,P° PμC,,PμPeC,PēPIPSCř C€PIPēPrPeC-CΓC,CH
PīPsC,PsPeCř PīP»P°P·PjPē PI PSPēCηPeP°C..., PSPμ P·PjC-PSCηCηC‡Pē
C,,C-P»CHbC,CηP°C†C-PMεPSPsC— C,,CηP°PeC†C-C— C- PSPμ PiCηPsPIPsPeCřCηC‡Pē
CηPsP·PIPēC,PeCř PiCηPsC,PμC—PSCřCηC-C—.

PJ PrPsCΓP»C-PrP¶PμPSPSCΠC... PiPμPjPsPrPēPSP°PjC-PePē Cř PīP°C†C-C"psc,c-pi
C-P· CΓPμCηC†PμPIPsCη PSPμPrPsCΓC,P°C,PSC-CΓC,Cη, P° C,P°PePsP¶ Cř
PeP»C-PSC-C‡PSPēC... PrPsCΓP»C-PrP¶PμPSPSCΠC... P·C- CΓC,CηPμCΓC,PμCΓC,PsPj
PīCηPē CΓPμCηC†PμPIC-PMε PSPμPrPsCΓC,P°C,PSPsCΓC,C- (PeP»P°CΓPē II-IV P·P°
NYHA) P°PjP»PsPrPēPīC-PS PSPμ PīPsPiC-CηC€CřPIP°PI CΓC,P°PSCř
CřC‡P°CΓPSPēPeC-PI PrPsCΓP»C-PrP¶PμPSPSCΠ, C%oPs PsC†C-PSCηPIP°PICΓCΠ P·P°
PīPμCηPμPSPsCΓPēPjC-CΓC,Cη PSP°PIP°PSC,P°P¶PμPSCη, C,,CηP°PeC†C-C"CH
PIPēPePēPrCř P»C-PIPsPiPs C€P»CřPSPsC‡PeP°, P° C,P°PePsP¶ P·P°
PeP»C-PSC-C‡PSPēPjPē PsP·PSP°PeP°PjPē C- CΓPēPjPiC,PsPjP°PjPē.

PJ PīP»P°C†PμP±Ps-PePsPSC,CηPsP»CHbPsPIP°PSPsPjCř PrPsCΓP»C-PrP¶PμPSPSC-
(PRAISE) P·P° CřC‡P°CΓC,Cη PīP°C†C-C"psc,c-pi C-P· CΓPμCηC†PμPIPsCη
PSPμPrPsCΓC,P°C,PSC-CΓC,Cη (PeP»P°CΓPē III-IV P·P° NYHA), CΠPeC-
PsC,CηPēPjCřPIP°P»Pē PrPēPiPsPeCΓPēPS, PrC-CřCηPμC,PēPePē C,P°
C-PSPiC-P±C-C,PsCηPē PηPμPα, P±CřP»Ps PīPsPeP°P·P°PSPs, C%oPs
P°PjP»PsPrPēPīC-PS PSPμ P·P±C-P»CHbC€CřC" CηPēP·PēPe Cř P»PμC,P°P»CHbPSPsPiPs
PSP°CΓP»C-PrPeCř P°P±Ps PīC-PrPIPēC%oPμPSPSCΠ CΓPjPμCηC,PSPsCΓC,C- C,P°
P·P°C...PIPsCηCηPIP°PSPsCΓC,C- Cř PīP°C†C-C"psc,c-pi C-P· CΓPμCηC†PμPIPsCη
PSPμPrPsCΓC,P°C,PSC-CΓC,Cη.

PJ PīPsPrP°P»CHbC€PsPjCř PrPsPIPiPsCΓC,CηPsPePsPIPsPjCř PīP»P°C†PμP±Ps-
PePsPSC,CηPsP»CHbPsPIP°PSPsPjCř PrPsCΓP»C-PrP¶PμPSPSC- (PRAISE-2)
P°PjP»PsPrPēPīC-PSCř P·P° CřC‡P°CΓC,Cη PīP°C†C-C"psc,c-pi C-P·
CΓPμCηC†PμPIPsCη PSPμPrPsCΓC,P°C,PSC-CΓC,Cη (NYHA, III C- IV) P±PμP·
PeP»C-PSC-C‡PSPēC... CΓPēPjPiC,PsPjC-PI P°P±Ps PsP±bη™ C"PeC,PēPIPSPēC...
PrP°PSPēC..., CΠPeC- P± CΓPIC-PrC‡PēP»Pē PiCηPs C-C€PμPjC-C‡PSCř
C...PIPsCηPsP±Cř CΓPμCηC†CΠ, PiCηPē P»C-PeCřPIP°PSPSC-
C-PSPiC-P±C-C,PsCηP°PjPē PηPμPα, PiCηPμPīP°CηP°C,P°PjPē
PSP°PīPμCηCΓC,CΠPSPePē C,P° PrC-CřCηPμC,PēPeP°PjPē PI PīPsCΓC,C-PMεPSPēC...

PrPsP·P°C... P°PjP»PsPrPëPïC-PS PSPµ PIPïP»PëPIP°PI PSP° CÍPjPµCĤC,PSC-CÍC,CH
P·P°PiP°P»PsPj C- CÍPjPµCĤC,PSC-CÍC,CH PI CĤPµP·CfP»CHC,P°C,C-
CÍPµCĤC†PµPIPs-CÍCfPrPëPSPSPëC... P·P°C...PIPsCĤCĤPIP°PSCĤ P·PsPeCĤPµPjP°. PJ
C†C-PNë PiCĤCfPïC- PïP°C†C-C" PSC,C-PI PIC-PrP·PSP°C†P°P»PsCÍCĬ
P·P±C-P»CHC€PµPSPSCĬ PIPëPïP°PrPeC-PI CĤPsP·PIPëC,PeCf PSP°P±CĤCĬPeCf
P»PµPiPµPSCĤ, PïPsPIBĤ™ CĬP·P°PSPsPiPs P· PïCĤPëPNëPsPjPsPj P°PjP»PsPrPëPïC-PSCf,
PïCĤPsC,Pµ CÍC,P°C,PëCÍC,PëC†PSPs P·PSP°C†CfC%PëC... PIC-PrPjC-PSPSPsCÍC,PµPNë
C†P°CÍC,PsC,Pë C,,P°PeC,C-PI PïPsPiC-CĤC€PµPSPSCĬ CÍPµCĤC†PµPIPsC—
PSPµPrPsCÍC,P°C,PSPsCÍC,C- PI PïPsCĤC-PIPSCĬPSPSC- P· C,P°PePsCĤ PïCĤPë
P·P°CÍC,PsCÍCfPIP°PSPSC- PïP»P°C†PµP±Ps PSPµ CÍPïPsCÍC,PµCĤC-PiP°P»PsCÍCĬ.

*PµCĤPsC,,C-P»P°PeC,PëC†PSP° C,PµCĤP°PiC-CĬ C-PSC,,P°CĤPeC,Cz PjC-PsPeP°CĤPrP°
(ALLHAT)*

P°P»CĬ PïPsCĤC-PIPSCĬPSPSCĬ PSPsPIC-C,PSC-C... C,PµCĤP°PiPµPIC,PëC†PSPëC...
PïC-PrC...PsPrC-PI P±CfP»Ps PïCĤPsPIPµPrPµPSPs PïPsPrPIC-PNëPSPs CÍP»C-PiPµ
CĤP°PSPPrPsPjC-P·PsPIP°PSPµ PrPsCÍP»C-PrP¶PµPSPSCĬ PIPïP»PëPICf PSP°
P·P°C...PIPsCĤCĤPIP°PSC-CÍC,CH C,P° CÍPjPµCĤC,PSC-CÍC,CH PïC-Pr PSP°P·PIPsCĤ
B«P°PëPïCĤPsP±CfPIP°PSPSCĬ C%PsPrPs P·P°CÍC,PsCÍCfPIP°PSPSCĬ
P°PSC,PëPiC-PiPµCĤC,PsPSC-C†PSPsC— C- P»C-PiC-PrPsP·PSPëP¶CfPIP°P»CHPSPsC—
C,PµCĤP°PiC-C— PrP»CĬ PïCĤPsC,,C-P»P°PeC,PëPePë C-PSC,,P°CĤPeC,Cf
PjC-PsPeP°CĤPrP°B» (ALLHAT): P°PjP»PsPrPëPïC-PS PI PrPsP·C- 2,5-10 PjPi/PrPsP±Cf
(P±P»PsPeP°C,PsCĤ PeP°P»CHC†C-C"PIPëC... PeP°PSP°P»C-PI) P°P±Ps
P»C-P·PëPSPsPïCĤPëP» CÍ PrPsP·C- 10-40 PjPi/PrPsP±Cf (C-PSPiC-P±C-C,PsCĤ
PĤPµPα) CĬPe C,PµCĤP°PiC-CĬ PïPµCĤC€PsPiPs PIPëP±PsCĤCf C- C,C-P°P·PëPrPSPëPNë
PrC-CfCĤPµC,PëPe C...P»PsCĤC,P°P»C-PrPsPS PI PrPsP·C- 12,5-25 PjPi/PrPsP±Cf
PïCĤPë PiC-PiPµCĤC,PµPSP·C-C— PIC-Pr P»PµPiPePsPiPs PrPs PïPsPjC-CĤPSPsPiPs
CÍC,CfPïPµPSCĬ. P—P°PiP°P»PsPj 33357 PïP°C†C-C" PSC,C-PI P·
P°CĤC,PµCĤC-P°P»CHPSPsCĤ PiC-PiPµCĤC,PµPSP·C-C" CĤ CÍ PIC-C†C- PIC-Pr 55
CĤPsPeC-PI P±CfP»Pë CĤP°PSPPrPsPjC-P·PsPIP°PSC- C- P·PSP°C...PsPrPëP»PëCÍCĬ
PïC-Pr CÍPïPsCÍC,PµCĤPµP¶PµPSPSCĬPj PI CÍPµCĤPµPrPSCĤPsPjCf PïCĤPsC,CĬPiPsPj
4,9 CĤPsPeCf. PJ PïP°C†C-C" PSC,C-PI P±CfPI CĬPe PjC-PSC-PjCfPj PsPrPëPS
PrPsPrP°C,PePsPIPëPNë C,,P°PeC,PsCĤ CĤPëP·PëPeCf CĤPsP·PIPëC,PeCf P†PĬPŸ,
PIPëP»CĤC†P°CĤC†Pë PïPµCĤPµPSPµCÍPµPSC- CĤP°PSC-C€Pµ C-PSC,,P°CĤPeC,
PjC-PsPeP°CĤPrP° C†Pë C-PSCÍCfP»CHC, (P±C-P»CHC€Pµ 6 PjC-CÍCĬC†C-PI
PïPµCĤPµPr PIC-PrP±PsCĤPsPj) P°P±Ps PSP°CĬPIPSC-CÍC,CH C-PSC€PëC... PŸPŸP—
P°C,PµCĤPsCÍPeP»PµCĤPsC,PëC†PSPsPiPs C...P°CĤP°PeC,PµCĤCf (PICÍCĤPsPiPs
51,5 %), C†CfPeCĤPsPIPëPNë PrC-P°P±PµC, 2 C,PëPïCf (36,1 %), CĤC-PIPµPSCĤ
P»PµP°P©-C...PsP»PµCÍC,PµCĤPëPSCf < 35 PjPi/PrP» (11,6 %),
PiC-PiPµCĤC,CĤPsC,,C-CĤ P»C-PIPsPiPs C€P»CfPSPsC†PeP°,
PrC-P°PiPSPsCÍC,PsPIP°PSCf P·P° PrPsPïPsPjPsPiPsCĤ
PµP»PµPeC,CĤPsPeP°CĤPrC-PsPiCĤP°C,,C-C— P°P±Ps
PµC...PsPeP°CĤPrC-PsPiCĤP°C,,C-C— (20,9 %), PeCfCĤC-PSPSCĬ PSP° C†P°CÍ
CfC†P°CÍC,C- PI PrPsCÍP»C-PrP¶PµPSPSC- (21,9 %).

PµPµCĤPIPëPSPSPsCĤ PeC-PSC†PµPIPsCĤ C,PsC†PePsCĤ PrPsCÍP»C-PrP¶PµPSPSCĬ
P±CfP»Ps PïPsC"PrPSP°PSPSCĬ P†PĬPŸ P· P»PµC,P°P»CHPSPëPj PSP°CÍP»C-PrPePsPj
P°P±Ps C-PSC,,P°CĤPeC, PjC-PsPeP°CĤPrP° P±PµP· P»PµC,P°P»CHPSPsPiPs

PSP°CÍP»C-PrPeCf. P—PSP°C†CfC%PëC... PIC-PrPjC-PSPSPsCÍC,PµPNë C%PsPrPs PïPµCĤPIPëPSPSPsC— PeC-PSC†PµPIPsC— C,PsC†PePë PrPsCÍP»C-PrP¶PµPSPSCĬ PjC-P¶ C,PµCĤP°PïC-C”CĤ P°PjP»PsPrPëPïC-PSPsPj C- C...P»PsCĤC,P°P»C-PrPsPSPsPj PSPµ P±CfP»Ps: PhP (RR 0,98 95 % P”P† (0,90–1,07) CĤ = 0,65. P©Ps CÍC,PsCÍCfC”C,CHCÍCĬ PIC,PsCĤPëPSPSPëC... PeC-PSC†PµPIPsC... C,PsC†PsPe PrPsCÍP»C-PrP¶PµPSPSCĬ, C†P°CÍC,PsC,P° CĤPsP·PIPëC,PeCf CÍPµCĤC†PµPIPsC— PSPµPrPsCÍC,P°C,PSPsCÍC,C- (PePsPjPïPsPSPµPSC, PePsPjP±C-PSPsPIP°PSPsC— PeC-PSC†PµPIPsC— C,PsC†PePë CÍPµCĤC†PµPIPs-CÍCfPrPëPSPSPsPiPs P·P°C...PIPsCĤCĤPIP°PSPSCĬ) P±CfP»P° P·PSP°C†PSPs PIPëC%PsCĤ CÍ PïCĤCfPïC-P°PjP»PsPrPëPïC-PSCf PïPsCĤC-PIPsCĬPSPs P· PïCĤCfPïPsCĤ C...P»PsCĤC,P°P»C-PrPsPSCf (10,2 % C- 7,7 %, PhP 1,38, 95 % P”P† [1,25–1,52] p < 0,001). PhPrPSP°Pe P·PSP°C†CfC%PëC... PIC-PrPjC-PSPSPsCÍC,PµPNë C%PsPrPs PïPsPeP°P·PSPëPeP° CÍPjPµCĤC,PSPsCÍC,C- P· P±CfPrCĤ-CĬPePsC— PïCĤPëC†PëPSPë PjC-P¶ C,PµCĤP°PïC-C”CĤ P°PjP»PsPrPëPïC-PSPsPj C- C...P»PsCĤC,P°P»C-PrPsPSPsPj PSPµ P±CfP»Ps (PhP 0,96, 95 % P”P† [0,89BĤ“1,02], p = 0,20).

P†PSCëP° C-PSC,,PsCĤPjP°C†C-CĬ

PŸPïC-P»CHPSPµ P·P°CÍC,PsCÍCfPIP°PSPSCĬ C-PSPiC-P±C-C,PsCĤC-PI PĥPµPα C,P° P±P»PsPeP°C,PsCĤC-PI CĤPµC†PµPïC,PsCĤC-PI P°PSPiC-PsC,PµPSP·PëPSCf II P±CfP»Ps PrPsCÍP»C-PrP¶PµPSPs CÍ PrPIPsC... CëPëCĤPsPePsPjP°CÍCëC,P°P±PSPëC..., CĤP°PSPrPsPjC-P·PsPIP°PSPëC..., PePsPSC,CĤPsP»CHPsPIP°PSPëC... PrPsCÍP»C-PrP¶PµPSPSCĬC... (ONTARGET (ONgoing Telmisartan Alone C- PI PePsPjP±C-PSP°C†C-C— P· Ramipril Global Endpoint Trial) C- VA NEPHRON-D (The Veterans Affairs Nephropathy in Diabetes)).

ONTARGET CĬPIP»CĬP»Ps CÍPsP±PsCĤ PrPsCÍP»C-PrP¶PµPSPSCĬ, PïCĤPsPIPµPrPµPSPµ P·P° CÍC†P°CÍC,CĤ PïP°C†C-C”PSC,C-PI C-P· CÍPµCĤC†PµPIPs-CÍCfPrPëPSPSPëPj P°P±Ps C†PµCĤPµP±CĤPsPIP°CÍPeCfP»CĬCĤPSPëPj P·P°C...PIPsCĤCĤPIP°PSPSCĬPj PI P°PSP°PjPSPµP·C- P°P±Ps C†CfPeCĤPsPIPëPj PrC-P°P±PµC,PsPj 2 C,PëPïCf, C%Ps CÍCfPïCĤPsPIPsPrP¶CfC”C,CHCÍCĬ PsP·PSP°PeP°PjPë CÍCĤP°P¶PµPSPSCĬ PsCĤPiP°PSCf-PjC-CëPµPSC-. VA NEPHRON-D CĬPIP»CĬP»Ps CÍPsP±PsCĤ PrPsCÍP»C-PrP¶PµPSPSCĬ, PïCĤPsPIPµPrPµPSPµ P·P° CÍC†P°CÍC,CĤ PïP°C†C-C”PSC,C-PI P· C†CfPeCĤPsPIPëPj PrC-P°P±PµC,PsPj 2 C,PëPïCf C,P° PrC-P°P±PµC,PëC†PSPsCĤ PSPµC,,CĤPsPïP°C,C-C”CĤ. P†C- PrPsCÍP»C-PrP¶PµPSPSCĬ PSPµ PIPëCĬPIPëP»Pë P·PSP°C†PëPjPsPiPs CÍPïCĤPëCĬC,P»PëPIPsPiPs PIPïP»PëPICf PSP° CĤPµP·CfP»CHC,P°C, PSPëCĤPePsPIPëC... C,P°/P°P±Ps CÍPµCĤC†PµPIPs-CÍCfPrPëPSPSPëC... P·P°C...PIPsCĤCĤPIP°PSCĤ C- PSP° CÍPjPµCĤC,PSC-CÍC,CH PIC-Pr PSPëC..., C,PsPrC- CĬPe CÍ PïPsCĤC-PIPsCĬPSPSC- P· PjPsPSPsC,PµCĤP°PïC-C”CĤ CÍPïPsCÍC,PµCĤC-PiP°PICÍCĬ PïC-PrPIPëC%PµPSPëPñë CĤPëP·PëPe CĤPsP·PIPëC,PeCf PïC-PïPµCĤPeP°P»C-C”PjC-C—, PiPsCÍC,CĤPsPiPs CÍCĤP°P¶PµPSPSCĬ PSPëCĤPsPe C,P°/P°P±Ps PïC-PïPsC,PsPSC-C—. P—PIP°P¶P°CĤC†Pë PSP° CÍC...PsP¶C-CÍC,CH C,,P°CĤPjP°PePsPrPëPSP°PjC-C†PSPëC... PIP»P°CÍC,PëPIPsCÍC,PµPNë, C†C- CĤPµP·CfP»CHC,P°C,Pë C,P°PePsP¶ P·P°CÍC,PsCÍPsPIPsC- PrP»CĬ C-PSCëPëC... C-PSPiC-P±C-C,PsCĤC-PI PĥPµPα C,P° P±P»PsPeP°C,PsCĤC-PI CĤPµC†PµPïC,PsCĤC-PI P°PSPiC-PsC,PµPSP·PëPSCf II.

PŸPĭC-P»CHPSPµ P·P°CĭC,PsCĭCřPIP°PSPSCĭ C-PSPiC-P±C-C,PsCĭC-PI PĥPµPα C,P° P±P»PsPeP°C,PsCĭC-PI CĭPµC†PµPĭC,PsCĭC-PI P°PSPiC-PsC,PµPSP·PĕPSCř II PĭCĭPsC,PĕPĭPsPeP°P·P°PSPs PĭP°C†C-C”PSC,P°Pj P· PrC-P°P±PµC,PĕC†PSPsCĭ PSPµC,,CĭPsPĭP°C,C-C”Cĭ.

ALTITUDE (Aliskiren Trial in Type 2 Diabetes Using Cardiovascular and Renal Disease Endpoints) Bĭ“ C†Pµ PrPsCĭP»C-PrP¶PµPSPSCĭ, PĭCĭPsPIPµPrPµPSPµ PrP»Cĭ PIPĕCĭPIP»PµPSPSCĭ PĭPsP·PĕC,PĕPIPSPsPiPs PµC,,PµPeC,Cř PIC-Pr PrPsPrP°PIP°PSPSCĭ P°P»C-CĭPeC-CĭPµPSCř PrPs CĭC,P°PSPPrP°CĭC,PSPsC— C,PµCĭP°PĭC-C— C-PSPiC-P±C-C,PsCĭP°PjPĕ PĥPµPα P°P±Ps P±P»PsPeP°C,PsCĭP°PjPĕ CĭPµC†PµPĭC,PsCĭC-PI P°PSPiC-PsC,PµPSP·PĕPSCř II PĭP°C†C-C”PSC,P°Pj P· C†CřPeCĭPsPIPĕPj PrC-P°P±PµC,PsPj 2 C,PĕPĭCř C,P° C...CĭPsPSC-C†PSPĕPj P·P°C...PIPSCĭPĭP°PSPSCĭPj PSPĕCĭPsPe, CĭPµCĭC†PµPIPSCĭCřPrPĕPSPSPĕPj P·P°C...PIPSCĭPĭP°PSPSCĭPj P°P±Ps C,PĕPj, C...C,Ps PjP°C” PsP±PĕPrPIP° P·P°C...PIPSCĭPĭP°PSPSCĭ. PĭPµ PrPsCĭP»C-PrP¶PµPSPSCĭ P±CřP»Ps PĭCĭPĕPĭPĕPSPµPSPs PrPsCĭC,CĭPsPePsPIPs Cř P·PIBĭ™ CĭP·PeCř P· PĭC-PrPIPĕC%PµPSPĕPj CĭPĕP·PĕPePsPj PSPµP±P°P¶P°PSPĕC... PSP°CĭP»C-PrPeC-PI. PŸPjPµCĭC,CH PIC-Pr CĭPµCĭC†PµPIPSCĭCřPrPĕPSPSPĕC... P·P°C...PIPSCĭPĭP°PSPSCĭ C- PIPĕPĭP°PrPePĕ PIPĕPSPĕPePSPµPSPSCĭ C-PSCĭCřP»CHC,Cř P±CřP»Pĕ C†P°CĭC,C-CĕPĕPjPĕ PI PiCĭCřPĭC-, C%Ps PĭCĭPĕPNĕPjP°P»P° P°P»C-CĭPeC-CĭPµPS, PSC-P¶ Cř PiCĭCřPĭC-, C%Ps PĭCĭPĕPNĕPjP°P»P° PĭP»P°C†PµP±Ps, C- PĭPsPIC-PrPsPjP»PµPSPSCĭ PĭCĭPs PSPµP±P°P¶P°PSC- CĭPIPĕC%P° C- CĭPµCĭPµPsP·PSC- PSPµP±P°P¶P°PSC- CĭPIPĕC%P° (PiC-PĭPµCĭPeP°P»C-C”PjC-Cĭ, P°CĭC,PµCĭC-P°P»CHPSP° PiC-PĭPsC,PµPSP·C-Cĭ C- PĭPsCĭCřCĕPµPSPSCĭ C,,CřPSPeC†C-C— PSPĕCĭPsPe) P±CřP»Pĕ C†P°CĭC,C-CĕPĕPjPĕ PI PiCĭCřPĭC-, CĭPeP° PĭCĭPĕPNĕPjP°P»P° P°P»C-CĭPeC-CĭPµPS, PSC-P¶ Cř PiCĭCřPĭC-, CĭPeP° PĭCĭPĕPNĕPjP°P»P° PĭP»P°C†PµP±Ps.

PαP°CĭPjP°PePsPeC-PSPµC,PĕPeP°.

PµC-CĭP»Cĭ PĭPµCĭPsCĭP°P»CHPSPsPiPs P·P°CĭC,PsCĭCřPIP°PSPSCĭ PĭCĭPµPĭP°CĭP°C,Cř PŸPµPIC-PeP°Cĭ PjP°PeCĭPĕPjP°P»CHPSP° PePsPSC†PµPSC,CĭP°C†C-Cĭ PsP»PjPµCĭP°CĭC,P°PSCř PjPµPrPsPeCĭPsPjC-P»Cř C,P° P°PjP»PsPrPĕPĭC-PSCř Cř PĭP»P°P·PjC- PeCĭPsPIC- PrPsCĭCĭPiP°C”C,CHCĭCĭ C†PµCĭPµP· 1,5Bĭ”2 C,P° 6Bĭ”8 PiPsPrPĕPS PIC-PrPĭPsPIC-PrPSPs. PĕPIPĕPrPeC-CĭC,CH C,P° CĭC,CřPĭC-PSCĭ PICĭPjPsPeC,CřPIP°PSPSCĭ PrPIPsC... PrC-CĭC†PĕC... CĭPµC†PsPIPĕPS PĭCĭPµPĭP°CĭP°C,Cř PŸPµPIC-PeP°Cĭ PIC-PrPĭPsPIC-PrP°C” C—C...PSC-PNĕ CĕPIPĕPrPePsCĭC,C- C,P° PICĭPjPsPeC,CřPIP°PSPSCĭ PĭCĭPĕ P·P°CĭC,PsCĭCřPIP°PSPSC- PsPeCĭPµPjPs. P’C-PsPrPsCĭC,CřPĭPSC-CĭC,CH PsP»PjPµCĭP°CĭC,P°PSCř PjPµPrPsPeCĭPsPjC-P»Cř C,P° P°PjP»PsPrPĕPĭC-PSCř Cř CĭPeP»P°PrC- P»C-PeP°CĭCĭCHPePsPiPs P·P°CĭPsP±Cř PŸPµPIC-PeP°Cĭ PSPµ P·P°P»PµP¶PĕC,CH PIC-Pr PĭCĭPĕPNĕPsPjCř C—P¶C-.

PĥP»PjPµCĭP°CĭC,P°PSCř PjPµPrPsPeCĭPsPjC-P» (PrC-CĭC†P° CĭPµC†PsPIPĕPSP° PŸPµPIC-PeP°CĭCř)

P’CĭPjPsPeC,CřPIP°PSPSCĭ C,P° CĭPsP·PĭPsPrC-P»

PhP»PjPµCΓP°CḡC,P°PSCr PjPµPrPsPeCΓPsPjC-P» Bḡ“ C†Pµ PīCḡPsP»C-PePë. P’C-PS C€PIPëPrPePs PīPµCḡPµC,PIPsCḡCḡC”C,CHCΓCḡ Cḡ C,,P°CḡPjP°PePsP»PsPiC-C†PSPs P°PeC,PëPIPSPëPNë PjPµC,P°P±PsP»C-C, PsP»PjPµCΓP°CḡC,P°PS PīC-Pr PrC-C”Cḡ PµCΓC,PµCḡP°P· Cḡ CΓP»PëP·PsPIC-PNë PsP±PsP»PsPSC†C- PePëC€PµC†PSPëPeCḡ C,P° PI PīPsCḡC,P°P»CHPSC-PNë PeCḡPsPIC- PīC-Pr C†P°CΓ PICΓPjPsPeC,CḡPIP°PSPSCḡ Cḡ C,CḡP°PIPSPsPjCḡ C,CḡP°PeC,C-. PJ PīP»P°P·PjC- PeCḡPsPIC- P°P±Ps Cḡ PīCḡPsPrCḡPeC,P°C... PIPëPIPµPrPµPSPSCḡ PSPµPīPµCḡPµC,PIPsCḡPµPSPëPNë PsP»PjPµCΓP°CḡC,P°PSCr PjPµPrPsPeCΓPsPjC-P» P°P±Ps P±PsPePsPIPëPNë P»P°PSC†CḡPi PjPµPrPsPeCΓPsPjC-P»CHPSPsC— PiCḡCḡPīPë PSPµ PIPëCḡPIP»CḡP»PëCḡCḡ. PŸPµCḡPµPrPSCḡ P°P±CḡPsP»CḡC,PSP° P±C-PsPrPsCḡC,CḡPīPSC-CḡC,CH PsP»PjPµCΓP°CḡC,P°PSCr Cḡ PIPëPīP»CḡPrC- C,P°P±P»PµC,PsPe CḡC,P°PSPsPIPëP»P° 25,6 %.

PŸPµCḡPµPrPSCḡ PjP°PeCḡPëPjP°P»CHPSP° PePsPSC†PµPSC,CḡP°C†C-Cḡ (PŸ_{max}) PsP»PjPµCΓP°CḡC,P°PSCr Cḡ PīP»P°P·PjC- PeCḡPsPIC- PrPsCḡCḡPīP°C”C,CHCΓCḡ PīCḡPëP±P»PëP·PSPs C†PµCḡPµP· 2 PīPsPrPëPSPë PīC-CḡP»Cḡ PīPµCḡPsCḡP°P»CHPSPsPiPs P·P°CḡC,PsCḡCḡPIP°PSPSCḡ. PḡPsPSC†PµPSC,CḡP°C†C-Cḡ PsP»PjPµCΓP°CḡC,P°PSCr Cḡ PīP»P°P·PjC- PeCḡPsPIC- P·P±C-P»CHC€CḡC”C,CHCΓCḡ PīCḡPëP±P»PëP·PSPs P»C-PSC-PNëPSPs P·C- P·P±C-P»CHC€PµPSPSCḡPj PsPrPSPsCḡP°P·PsPIPsc— PrPsP·Pë PrPs 80 PjPi.

P†P¶P° C†PëPSPëC,CH PjC-PSC-PjP°P»CHPSCr PrC-Cḡ PSP° P±C-PsPrPsCḡC,CḡPīPSC-CḡC,CH PsP»PjPµCΓP°CḡC,P°PSCr, C,PsPjCḡ PsP»PjPµCΓP°CḡC,P°PSCr PjPµPrPsPeCΓPsPjC-P» PjPsP¶PSP° P·P°CḡC,PsCḡPsPICrPIP°C,Pë PSPµP·P°P»PµP¶PSPs PIC-Pr C—C— PīCḡPëPNëPsPjCḡ.

PḡP»C-PSC-C†PSPs P·PSP°C†Cḡ%PsC— CḡC-P·PSPëC†C- Cḡ C,,P°CḡPjP°PePsPeC-PSPµC,PëC†C- PsP»PjPµCΓP°CḡC,P°PSCr P·P°P»PµP¶PSPs PIC-Pr CḡC,P°C,C- PīP°C†C-C”PSC,P° PSPµ PIPëCḡPIP»PµPSPs.

PhP»PjPµCΓP°CḡC,P°PS P°PeC,PëPIPSPs P·PīBḡ™ CḡP·CḡC”C,CHCΓCḡ P· P±C-P»PeP°PjPë PeCḡPsPIC- (99,7 %), PsPrPSP°Pe CḡPëP·PëPe PeP»C-PSC-C†PSPs P·PSP°C†Cḡ%PsC— PePsPSPeCḡCḡPµPSC,PSPsC— PIP·P°C”PjPsPrC-C— P· C-PSC€PëPjPë PīCḡPµPīP°CḡP°C,P°PjPë, C%Ps P°PeC,PëPIPSPs P·PīBḡ™ CḡP·CḡCḡC,CHCΓCḡ P· P±C-P»PeP°PjPë PeCḡPsPIC-, PrPsCḡC,P°C,PSCHPs PSPëP·CHPePëPNë. PµC-PrC,PIPµCḡPrP¶PµPSPSCḡPj C†CHPsPiPs C” PIC-PrCḡCḡC,PSC-CḡC,CH C,P°PePsC— PIP·P°C”PjPsPrC-C— PjC-P¶ PsP»PjPµCΓP°CḡC,P°PSCr PjPµPrPsPeCΓPsPjC-P»PsPj C,P° PIP°CḡC,,P°CḡPëPSPsPj. PhP»PjPµCΓP°CḡC,P°PS PSPµP·PSP°C†PSPsCḡ PjC-CḡPsCḡ P·PīBḡ™ CḡP·CḡC”C,CHCΓCḡ P· PeP»C-C,PëPSP°PjPë PeCḡPsPIC-. PŸPµCḡPµPrPSC-PNë PsP±Bḡ™ C”Pj CḡPsP·PīPsPrC-P»Cḡ PīC-CḡP»Cḡ PIPSCrC,CḡC-C€PSCḡPsPIPµPSPSPsPiPs P·P°CḡC,PsCḡCḡPIP°PSPSCḡ PSPµPIPëCḡPsPePëPNë (16–29 P»).

PḡPµC,P°P±PsP»C-P·Pj C,P° PIPëPIPµPrPµPSPSCḡ

P—P°PīP°P»CHPSPëPNë PīP»P°P·PjPsPIPëPNë PeP»C-CḡPµPSCḡ PsP»PjPµCΓP°CḡC,P°PSCr P·P°P·PIPëC†P°PNë CḡC,P°PSPsPIPëC,CH 1,3 P»/PīPsPrPëPSCḡ (PePsPµC,,C-C†C-C”PSC,

PIP°СҒС-Р°С†С-С— 19 %) С- PIC-PrPSPsCГPSPs PSpµPIPµP»PëPePëPNë
PïPsСҒС-PIPSCЦPSPs P· PïPµC†С-PSPePsPIPëPj PeСҒPsPIPсC,PsPePsPj
(PïСҒPëP±P»PëP·PSPs 90 P»/PiPsPrPëPSCГ).

PµС-СГР»СЦ PsPrPSPsСҒP°P·PsPIPsPiPs PïPµСҒPsСҒP°P»СҒPSPsPiPs
P·P°СГС,PsCГCГPIP°PSPSCЦ PsP»PjPµCГP°СҒС,Р°PSCГ PjPµPrPsPeCГPsPjC-P»CГ,
PjC-C†PµPSPsPiPs C-P·PsC,PsPiPsPj 14PŸ, 10BҒ“16 % СҒP°PrC-PsP°PeC,PëPIPSPsC—
СҒPµC†PsPIPëPSPë CГPïPsCГC,РµСҒС-РiP°P»PsCГCЦ PI CГPµC†С- (P±С-P»СҒС€P°
C†P°СГС,PëPSP° PïСҒPsC,СЦPiPsPj 24 PiPsPrPëPS PïC-СГР»СЦ
P·P°СГС,PsCГCГPIP°PSPSCЦ), P° СҒPµC€C,P° СҒP°PrC-PsP°PeC,PëPIPSPsC—
СҒPµC†PsPIPëPSPë PIPëPIPsPrPëP»P°СГСҒ C-P· PeP°P»PsPj. PкP° PsCГPSPsPIC-
CГPëCГC,РµPjPSPsC— PrPsCГC,CГPïPSPsCГC,C-, C%°Ps CГPeP»P°PrP°С” 25,6 %,
PjPsP¶PSP° СҒPsP·СҒP°С...CГPIP°С,Pë, C%°Ps P°P±CГPsСҒP±PsPIP°PSPëPNë
PsP»PjPµCГP°СҒС,Р°PS PIPëPIPsPrPëC,СҒCГCЦ СЦPe PSPëСҒPeP°PjPë (PïСҒPëP±P»PëP·PSPs
40 %), C,P°Pe C- PiPµPïP°С,PsP±С-P»C-P°СҒPSPsСҒ CГPëCГC,РµPjPsСҒ (PïСҒPëP±P»PëP·PSPs
60 %). PjCГCЦ PIPëCЦPIP»PµPSP° СҒP°PrC-PsP°PeC,PëPIPSC-CГC,СҒ P±CГP»P°
C-PrPµPSC,PëC,,C-PePsPIP°PSP° СЦPe PsP»PjPµCГP°СҒС,Р°PS. P†PSC€PëC...
P·PSP°C†CГC%°PëC... PjPµC,P°P±PsP»C-C,C-PI P·PSP°PNëPrPµPSPs PSpµ P±CГP»Ps.
PъPëC€PePsPIPs-PiPµC†С-PSPePsPIP° СҒPµC†PëСҒPeCГP»СЦC†C-СЦ PsP»PjPµCГP°СҒС,Р°PSCГ
PjC-PSC-PjP°P»СҒPSP°. PҒCГPeC-P»СҒPëPë P±С-P»СҒС€P° C†P°СГС,PëPSP°
PsP»PjPµCГP°СҒС,Р°PSCГ PIPëPIPsPrPëC,СҒCГCЦ P· P¶PsPIC†СҒ, C,Ps PNëPsPiPs
P·P°СГС,PsCГCГPIP°PSPSCЦ C...PIPsСҒPëPj C-P· P±С-P»C-P°СҒPSPsСҒ
PsP±CГC,СҒCГPeC†C-C”СҒ PïСҒPsC,PëPiPsPeP°P·P°PSPs (PrPëPI. СҒPsP·PrC-P»
B«PµСҒPsC,PëPïPsPeP°P·P°PSPSCЦB»).

PŸPµСҒPjC-PSP°P»СҒPSPëPNë PïPµСҒС-PsPr PSP°PïC-PIPIPëPIPµPrPµPSPSCЦ
PsP»PjPµCГP°СҒС,Р°PSCГ PïC-СГР»СЦ P±P°PiP°С,PsСҒP°P·PsPIPsPiPs
PïPµСҒPsСҒP°P»СҒPSPsPiPs P·P°СГС,PsCГCГPIP°PSPSCЦ PePsP»PëPIP°С”C,СҒCГCЦ
PIC-Pr 10 PrPs 15 PiPsPrPëPS. PŸC,Р°PS СҒС-PIPSPsPIP°PiPë
PrPsCГCЦPiP°С”C,СҒCГCЦ PïC-СГР»СЦ PïPµСҒС€PëC... PeC-P»СҒPëPsC... PrPsP·, P°
PïC-СГР»СЦ 14 PrC-P± P±P°PiP°С,PsСҒP°P·PsPIPsPiPs P·P°СГС,PsCГCГPIP°PSPSCЦ
PïPsPrP°P»СҒС€PsC— PeCГPjCГP»СЦC†C-C— PSpµ
CГPïPsCГC,РµСҒС-РiP°С”C,СҒCГCЦ. PкPëСҒPePsPIPëPNë PeP»C-СҒPµPSCГ
CГC,Р°PSPsPIPëPI PïСҒPëP±P»PëP·PSPs 0,5BҒ“0,7 P»/PiPsPrPëPSCГ C,P° PSpµ
P·P°P»PµP¶P°PI PIC-Pr PrPsP·Pë PïСҒPµPïP°СҒP°С,CГ.

P>C-PeP°СҒCГСҒPeC- PIP·P°С”PjPsPrC-C—

P>C-PeP°СҒCГСҒPePëPNë P·P°СГС-P± PePsP»PµCГPµPIPµP»P°Pj, C%°Ps P·PIBҒ™ СЦP·CГC”
P¶PsPIC†PSC- PePëCГP»PsC,Pë

PŸPïC-P»СҒPSPëPNë PïСҒPëPNëPsPj 40 PjPi PsP»PjPµCГP°СҒС,Р°PSCГ
PjPµPrPsPeCГPsPjC-P»CГ C- 3750 PjPi PePsP»PµCГPµPIPµP»P°PjCГ
PïC-PrСҒPsC...P»PsСҒPëPrCГ CГ P·PrPsСҒPsPIPëC... PrPsP±СҒPsPIPsP»СҒC†C-PI
PïСҒPëP·PIPsPrPëPI PrPs P·PSPëP¶PµPSPSCЦ C_{max} PsP»PjPµCГP°СҒС,Р°PSCГ PSP° 28 %
C- PrPs P·PSPëP¶PµPSPSCЦ AUC PsP»PjPµCГP°СҒС,Р°PSCГ PSP° 39 %. PҒPµPSC€PëPNë
PIPïP»PëPI, P·PSPëP¶PµPSPSCЦ C_{max} C- AUC PSP° 4 % C- 15 % PIC-PrPïPsPIC-PrPSPs,
CГPïPsCГC,РµСҒС-РiP°PICГCЦ, PePsP»Pë PsP»PjPµCГP°СҒС,Р°PSCГ
PjPµPrPsPeCГPsPjC-P» PïСҒPëP·PSP°C†P°P»Pë P·P° 4 PiPsPrPëPSPë PrPs

PiC7P8PN0PsPjCf PePsP»PmCfPmPIPmP»P°PjCf PiC-PrC7PsC...P»PsC7P8PrCf. P8P°Cf PSP°PiC-PIPIP8PIPMPrPmPSPSCf PsP»PjPmCfP°C7C,P°PSCf P·PSP8P¶CfPIP°PICfCfCf PSP° 50B7“52 % PSPmP·P°P»PmP¶PSPs PIC-Pr C,PsPiPs, PiC7P8P·PSP°CfP°P»P8CfCfCf PiC7PmPiP°C7P°C,P8 CfPiC-P»CHBPSPs CfP8 PiC7P8PN0PsPj PsP»PjPmCfP°C7C,P°PSCf PIC-PrP±CfPIP°PICfCfCf P·P° 4 PiPsPrP8PSP8 PrPs PiC7P8PN0PsPjCf PePsP»PmCfPmPIPmP»P°PjCf PiC-PrC7PsC...P»PsC7P8PrCf (PrP8PI. C7PsP·PrC-P» B«P·P·P°C”PjPsPrC-Cf P· C-PSC8P8PjP8 P»C-PeP°C7CfC7PeP8PjP8 P·P°CfPsP±P°PjP8 C,P° C-PSC8C- PIP8PrP8 PIP·P°C”PjPsPrC-PN0B»).

P8PjP»PsPrP8PiC-PS (PrC-CHCfP° C7PmCfPsPIP8PSP° P8PmPIC-PeP°C7C8)

P°CfPjPsPeC,CfPIP°PSPSCf C,P° C7PsP·PiPsPrC-P»

PmC-CfP»Cf PiPmC7PsC7P°P»CHBPSPsPiPs P·P°CfC,PsCfCfPIP°PSPSCf Cf C,PmC7P°PiPmPIC,P8CfPSP8C... PrPsP·P°C... P°PjP»PsPrP8PiC-PS PrPsP±C7Pm PICfPjPsPeC,CfC”C,CHCfCfCf C-P· PrPsCfCfPiPSPmPSPSCfPj PiC-PePsPiPsC— PePsPSCfPmPSC,C7P°CfC-C— Cf PeC7PsPIC- CfPmC7PmP· 6B7“12 PiPsPrP8PS PiC-CfP»Cf PiC7P8PN0PsPjCf. P8P±CfPsP»CHC,PSP° P±C-PsPrPsCfC,CfPiPSC-CfC,CH CfC,P°PSPsPIP8C,CH PiC7P8P±P»P8P·PSPs 64B7“80 %. P8P±B7™ C”Pj C7PsP·PiPsPrC-P»Cf CfC,P°PSPsPIP8C,CH PiC7P8P±P»P8P·PSPs 21 P»/PePi. P”PsCfP»C-PrP¶PmPSPSCf *in vitro* PiPsPeP°P·P°P»P8, C%0Ps PjP°PN0P¶Pm 97,5 % CfP8C7PeCfP»CHCHCfPsPiPs P°PjP»PsPrP8PiC-PSCf P·PIB7™ CfP·CfC”C,CHCfCfCf PiC7PsC,PmC—PSP°PjP8 PiP»P°P·PjP8.

PmC7P8PN0PsPj C—P¶C- PSP° PiC7PsCfPmCf PICfPjPsPeC,CfPIP°PSPSCf P°PjP»PsPrP8PiC-PSCf PSPm PIPiP»P8PIP°C”.

P8PmC,P°P±PsP»C-P·Pj C,P° PIP8PIPMPrPmPSPSCf

PmPmC7C—PsPr PSP°PiC-PIPIP8PIPMPrPmPSPSCf P· PiP»P°P·PjP8 PeC7PsPIC— C7P°P·PsPiPsC— PrPsP±PsPiPsC— PrPsP·P8 PePsP»P8PIP°C”C,CHCfCfCf PIC-Pr 35 PrPs 50 PiPsPrP8PS. P8PjP»PsPrP8PiC-PS P·PSP°CfPSPsC7 PjC-C7PsC7 PjPmC,P°P±PsP»C-P·CfC”C,CHCfCfCf Cf PiPmCfC—PSCfC— P· CfC,PiPsC7PmPSPSCfPj PSPmP°PeC,P8PIPSP8C... PjPmC,P°P±PsP»C-C,PI. PmC7P8P±P»P8P·PSPs 60 % PIPiPmPrPmPSPsC— PrPsP·P8 PIP8PIPsPrP8C,CHCfCfCf P·C— CfPmCfPmC7, PiC7P8P±P»P8P·PSPs 10 % P· CfPeP8C... CfC,P°PSPsPIP8C,CH P°PjP»PsPrP8PiC-PS Cf PSPmP·PjC—PSPmPSPsPjCf PIP8PiP»CfPrC—.

P8P»PjPmCfP°C7C,P°PSCf PjPmPrPsPeCfPsPjC-P» C,P° P°PjP»PsPrP8PiC-PS (PrC-CHCfC— C7PmCfPsPIP8PSCf P8PmPIC-PeP°C7Cf)

P8PeC7PmPjC- PiC7CfPiP8 PiP°CfC-C”PSC,C-PI

P”C-C,P8 (PIC-PePsPj PrPs 18 C7PsPeC-PI)

P”P°PSP8C... PiC7Ps C,,P°C7PjP°PePsPeC-PSPmC,P8PeCf Cf PrC-C,PmPN0 PSPmPjP°C”.

PmP°CfC-C”PSC,P8 P»C-C,PSCHBPsiPs PIC-PeC8 (PIC-PePsPj PIC-Pr 65 C7PsPeC-PI)

P”PsPiPmPrPmPSPs, C%0Ps PiC7P8 P°C7C,PmC7C-P°P»CHBPSC-PN0 PiC-PiPmC7C,PmPSP·C-C— AUC (PiP»PsC%0P° PiC-Pr PeC7P8PIPsC7

B«PePsPSC†PμPSC, CḡP°C†C-Cḡ-C†P°CḡB») PsP»PjPμCḡP°CḡC, P°PSCr Cḡ
CḡC-PIPSPsPIP°PḡPSPsPjCḡ CḡC, P°PSC- P·P±C-P»CHC€CḡPIP°P»P°CḡCḡ
PḡCḡPḡP±P»PḡP·PSPs PSP° 35 % Cḡ P»C-C, PSC-C... PḡP°C†C-C”PSC, C-PI (PIC-PePsPj
65ḡḡ”75 CḡPsPeC-PI) P· P°CḡC, PμCḡC-P°P»CHPSPsCḡ PiC-PḡPμCḡC, PμPSP·C-C”Cḡ
PḡPsCḡC-PIPSCḡPSPs P· PjPsP»PsPrC€PsCḡ PiCḡCḡPḡPsCḡ C, P° PḡCḡPḡP±P»PḡP·PSPs
PSP° 44 % Cḡ PrCḡPḡPμ P»C-C, PSC-C... PḡP°C†C-C”PSC, C-PI (≥ 75 CḡPsPeC-PI) (PrPḡPI.
CḡPsP·PrC-P» B«PḡPḡPsCḡC-P± P·P°CḡC, PsCḡCḡPIP°PSPSCḡ C, P° PrPsP·PḡB»). PḡPμ
PjPsPḡPSP° PḡPsCḡCḡPSPḡC, Pḡ PḡPsPjC-CḡPSPḡPj P·PSPḡPḡPμPSPSCḡPj
C,, CḡPSPeC†C-C— PSPḡCḡPsPe, P·P°P»PμPḡPSPḡPj PIC-Pr PIC-PeCḡ PI C†C-PNḡ
PiCḡCḡPḡC- PḡP°C†C-C”PSC, C-PI. PhPrPSP°Pe PrP»Cḡ PḡP°C†C-C”PSC, C-PI
P»C-C, PSCḡPsPiPs PIC-PeCḡ CḡPμPePsPjPμPSPPrCḡC”C, CHCḡCḡ C, PsPNḡ CḡP°PjPḡPNḡ
CḡPμPḡPḡPj PrPsP·CḡPIP°PSPSCḡ, C%ḡPs PNḡ PrP»Cḡ C-PSC€PḡC... PḡP°C†C-C”PSC, C-PI,
P°P»Pμ PiCḡPḡ C†CHPsPjCḡ CḡPμPePsPjPμPSPPrCḡC”C, CHCḡCḡ P·
PsP±PμCḡPμPḡPSC-CḡC, Cḡ PiC-PrPIPḡC%ḡCḡPIP°C, Pḡ PrPsP·Cḡ PiCḡPμPḡP°CḡP°C, Cḡ.

PḡP°Cḡ PrPsCḡCḡPiPSPμPSPSCḡ PjP°PeCḡPḡPjP°P»CHPSPsC—
PePsPSC†PμPSC, CḡP°C†C-C— P°PjP»PsPrPḡPiC-PSCr Cḡ PḡP»P°P·PjC- PeCḡPsPIC-
PsPrPSP°PePsPIPḡPNḡ Cḡ PḡP°C†C-C”PSC, C-PI CḡC, P°CḡC€PsPiPs C, P°
PjPsP»PsPrC€PsPiPs PIC-PeCḡ. Pj PḡP°C†C-C”PSC, C-PI P»C-C, PSCḡPsPiPs PIC-PeCḡ
CḡPḡPsCḡC, PμCḡC-PiP°C”C, CHCḡCḡ C, PμPSPPrPμPSC†C-Cḡ PrPs P·PjPμPSC€PμPSPSCḡ
PeP»C-CḡPμPSCḡCḡ P°PjP»PsPrPḡPiC-PSCr, C%ḡPs PḡCḡPḡP·PIPPrPḡC, CH PrPs
PḡC-PrPIPḡC%ḡPμPSPSCḡ AUC C, P° C, CḡPḡPIP°P»PsCḡC, C- PḡPμCḡC-PsPrCḡ
PSP°PiC-PIPIPḡPIPμPrPμPSPSCḡ. PμC-PrPIPḡC%ḡPμPSPSCḡ AUC C, P°
C, CḡPḡPIP°P»C-CḡC, CH PḡPμCḡC-PsPrCḡ PSP°PiC-PIPIPḡPIPμPrPμPSPSCḡ Cḡ
PḡP°C†C-C”PSC, C-PI C-P· P·P°CḡC, C-PNḡPSPsCḡ CḡPμCḡC†PμPIPpsCḡ
PSPμPrPsCḡC, P°C, PSC-CḡC, Cḡ PIC-PrPḡPsPIC-PrP°P»Pḡ PsC†C-PeCḡPIP°PSPSCḡPj
PrP»Cḡ C†C-C”C— PIC-PePsPIPpsC— PiCḡCḡPḡPḡ (PrPḡPI. CḡPsP·PrC-P»
B«PhCḡPsP±P»PḡPIPpsCḡC, C- P·P°CḡC, PsCḡCḡPIP°PSPSCḡB»).

PμPsCḡCḡC€PμPSPSCḡ C,, CḡPSPeC†C-C— PSPḡCḡPsPe

Pj PḡP°C†C-C”PSC, C-PI C-P· PḡPsCḡCḡC€PμPSPSCḡPj C,, CḡPSPeC†C-C— PSPḡCḡPsPe
P»PμPiPePsPiPs, PḡPsPjC-CḡPSPsPiPs C, P° C, CḡPḡPePsPiPs CḡC, CḡPḡPμPSCḡ AUC
PsP»PjPμCḡP°CḡC, P°PSCr Cḡ CḡC-PIPSPsPIP°PḡPSPsPjCḡ CḡC, P°PSC- P±CḡP»P° PIPḡC%ḡPsCḡ
PSP° 62 %, 82 % C, P° 179 % PIC-PrPḡPsPIC-PrPSPs, PḡPsCḡC-PIPSCḡPSPs P·C-
P·PrPsCḡPsPIPḡPjPḡ PrPsP±CḡPsPIPpsP»CHC†CḡPjPḡ (PrPḡPI. CḡPsP·PrC-P»Pḡ B«PḡPḡPsCḡC-P±
P·P°CḡC, PsCḡCḡPIP°PSPSCḡ C, P° PrPsP·PḡB», B«PhCḡPsP±P»PḡPIPpsCḡC, C-
P·P°CḡC, PsCḡCḡPIP°PSPSCḡB»).

PhPjP»PsPrPḡPiC-PS P·PSP°C†PSPsCḡ PjC-CḡPsCḡ PjPμC, P°P±PsP»C-P·CḡC”C, CHCḡCḡ
PrPs PSPμP°PeC, PḡPIPSPḡC... PjPμC, P°P±PsP»C-C, C-PI. 10 % CḡPμC†PsPIPḡPSPḡ
PIPḡPIPpsPrPḡC, CHCḡCḡ C-P· CḡPμC†PμCḡ Cḡ PSPμP·PjC-PSPμPSPsPjCḡ CḡC, P°PSC-.
P—PjC-PSP° PePsPSC†PμPSC, CḡP°C†C-C— P°PjP»PsPrPḡPiC-PSCr Cḡ PḡP»P°P·PjC-
PeCḡPsPIC- PSPμ PePsCḡPμP»CḡC”C, CHCḡCḡ P·C- CḡC, CḡPḡPμPSPμPj
PḡPsCḡCḡC€PμPSPSCḡ C,, CḡPSPeC†C-C— PSPḡCḡPsPe. PḡP°PePḡPj C... PIPsCḡPḡPj
P°PjP»PsPrPḡPiC-PS PjPsPḡPSP° PḡCḡPḡP·PSP°C†P°C, Pḡ Cḡ P·PIPḡC†P°PNḡPSPḡC...
PrPsP·P°C.... PhPjP»PsPrPḡPiC-PS PSPμ PIPḡPIPpsPrPḡC, CHCḡCḡ P·P°
PrPsPiPsPjPsPiPsCḡ PiPμPjPsPrC-P°P»C-P·Cḡ.

Р҃C-С҃P»C҃ PсPrPSPсC҃P°P·PсPIPsPiPs PиPμC҃҃PсC҃P°P»C҃҃PSPсPiPs
P·P°C҃C,PсC҃C҃PиP°PSPSC҃ P·PSP°C҃PμPSPSC҃ AUC PсP»PjPμC҃P°C҃C,P°PSC҃
P±C҃P»Pε PSP° 6 % C,P° PSP° 65 % PиPεC%PεPjPε C҃ C...PиPсC҃PεC... C-P·
PиPsC҃C҃CεPμPSPSC҃PjPε C,,C҃PSPeC҃C-C— PиPμC҃C-PSPePε P»PμPiPePsPiPs P°P±Ps
PиPsPjC-C҃҃PSPсPiPs C҃C,C҃PиPμPSC҃ PIC-PrPиPsPIC-PrPSPс PиPsC҃҃C-PIPSC҃PSPс P·C-
P·PrPsC҃҃PсPIPεPjPε PrPsP±C҃҃PсPIPsP»C҃҃C҃C҃PjPε. PкPμP·PIB҃҃™ C҃P·P°PSP°
C,,C҃P°PeC҃C-C҃ PсP»PjPμC҃P°C҃C,P°PSC҃ C҃PμC҃҃PμP· 2 PиPsPrPεPSPε PиC-С҃P»C҃
P·P°C҃C,PсC҃C҃PиP°PSPSC҃ C҃ P·PrPsC҃҃PсPIPεC... PrPsP±C҃҃PсPIPsP»C҃҃C҃C-PI, C҃
C...PиPсC҃PεC... P· P»PμPiPePεPj P°P±Ps PиPsPjC-C҃҃PSPεPj PиPsC҃҃C҃CεPμPSPSC҃Pj
C,,C҃PSPeC҃C-C— PиPμC҃C-PSPePε C҃C,P°PSPсPIPεP»P° 0,26 %, 0,34 % C,P° 0,41 %
PIC-PrPиPsPIC-PrPSPс. PμC҃҃Pε P±P°PiP°C,PсC҃҃P°P·PсPIPsPjC҃
P·P°C҃C,PсC҃C҃PиP°PSPSC- C҃PμC҃҃PμPrPSC҃ AUC PсP»PjPμC҃P°C҃C,P°PSC҃ C҃
C...PиPсC҃PεC... P· PиPsC҃҃C҃CεPμPSPSC҃Pj C,,C҃PSPeC҃C-C— PиPμC҃C-PSPePε
C҃PμC҃҃PμPrPSC҃҃PсPiPs C҃C,C҃PиPμPSC҃ C,C҃P҃PεPсC҃C,C- P±C҃P»P° PSP° 65 %
PиPεC%PсC҃, PSC-P҃ C҃ P·PrPsC҃҃PсPIPεC... PrPsP±C҃҃PсPIPsP»C҃҃C҃C-PI.
P҃PμC҃҃PμPrPSC” P·PSP°C҃PμPSPSC҃ P҃_max PсP»PjPμC҃P°C҃C,P°PSC҃ C҃
C...PиPсC҃PεC... P· PиPsC҃҃C҃CεPμPSPSC҃Pj C,,C҃PSPeC҃C-C— PиPμC҃C-PSPePε C,P°
P·PrPsC҃҃PсPIPεC... PrPsP±C҃҃PсPIPsP»C҃҃C҃C-PI P±C҃P»Pε PиPsPrC-P±PSC-.
P҃P»PjPμC҃P°C҃C,P°PSC҃ PjPμPrPsPeC҃PсPjC-P» PSPμ PсC҃C-PSC҃҃PиP°P»Pε C҃
C...PиPсC҃PεC... P· C,C҃P҃PεPεPj PиPsC҃҃C҃CεPμPSPSC҃Pj PиPμC҃C-PSPePε (PrPεPI.
C҃҃PсP·PrC-P»Pε B«P҃PиPsC҃C҃C-P± P·P°C҃C,PсC҃C҃PиP°PSPSC҃ C,P° PrPsP·PεB»,
B«P҃C҃PсP±P»PεPIPsC҃C҃C- P·P°C҃C,PсC҃C҃PиP°PSPSC҃B»).

P”PсC҃C,C҃PиPSC- P»PεCεPμ PrC҃P҃Pμ PсP±PjPμP҃PμPSC- PeP»C-PSC-C҃PSC-
PrP°PSC- C҃C,PсC҃C҃PиPSPс P·P°C҃C,PсC҃C҃PиP°PSPSC҃ P°PjP»PсPrPεPиC-PSC҃
C...PиPсC҃PεPj C-P· C,C҃P҃PεPεPjPε PиPsC҃҃C҃CεPμPSPSC҃PjPε C,,C҃PSPeC҃C-PMε
PиPμC҃C-PSPePε. PJ C...PиPсC҃PεC... P· PиPsC҃҃C҃CεPμPSPSC҃Pj C,,C҃PSPeC҃C-C—
PиPμC҃C-PSPePε C҃PиPsC҃C҃,PμC҃҃C-PиP°C”C,C҃҃C҃C҃ P·PSPεP҃PμPSPSC҃
PeP»C-C҃҃PμPSC҃C҃ P°PjP»PсPrPεPиC-PSC҃ C,P° PиPsPrPsPIP҃PμPSPSC҃
PиPμC҃҃C-PсPrC҃ PSP°PиC-PIIPεPIPμPrPμPSPSC҃, C%Pс PиC҃҃PεP·PIPsPrPεC,C҃҃ PrPs
P·P±C-P»C҃҃CεPμPSPSC҃ AUC PиC҃҃PεP±P»PεP·PSPс PSP° 40B҃҃”60 % (PrPεPI.
C҃҃PсP·PrC-P»Pε B«P҃PиPsC҃C҃C-P± P·P°C҃C,PсC҃C҃PиP°PSPSC҃ C,P° PrPsP·PεB»,
B«P҃C҃PсP±P»PεPIPsC҃C҃C- P·P°C҃C,PсC҃C҃PиP°PSPSC҃B»).

P”PsPeP»C-PSC-C҃PSC- PrP°PSC- P· P±PμP·PиPμPePε

P— PsPiP»C҃PPrC҃ PSP° PrPsPeP»C-PSC-C҃PSPεPMε PиC҃҃PсC,,C-P»C҃҃
C,PсPeC҃PεC҃PSPсC҃C,C- PePsP҃PSPсC— PrC-C҃҃C҃PсC— C҃҃PμC҃҃PсPIPεPSPε
PиC-PrPIPεC%PμPSPSC҃ C,PсPeC҃PεC҃PSPсC҃C,C- PrP»C҃
PePsPjP±C-PSPсPIP°PSPсPiPs PиC҃҃PμPиP°C҃P°C,C҃ PSPμ PсC҃C-PεC҃C”C,C҃҃C҃C҃,
PсC҃PεC-P»C҃҃PePε C҃C- C҃҃PμC҃҃PсPIPεPSPε PиPиP»PεPIP°C҃҃C,C҃҃ PSP° C҃҃C-P·PSC-
PсC҃҃PиP°PSPε: PсP»PjPμC҃P°C҃C,P°PSC҃ PjPμPrPsPeC҃PсPjC-P» PrC-C” PSP°
PSPεC҃҃PePε, P° P°PjP»PсPrPεPиC-PS - PSP° C҃PμC҃҃C҃Pμ.

PJ 3-PjC-C҃C҃C҃PSPсPjC҃ PrPsC҃P»C-PrP҃PμPSPSC- C,PсPeC҃PεC҃PSPсC҃C,C-
PePsPjP±C-PSPсPIP°PSPсPiPs PиC҃҃PμPиP°C҃P°C,C҃ PсP»PjPμC҃P°C҃C,P°PSC҃

PjPμPrPsPeCÍPsPjC-P»Cí/ P°PjP»PsPrPëPíC-PSCr PSP° C%oCíCĤP°C... PíCĤPë
P±P°PiP°C,PsCĤP°P·PsPIPsPjCí PíCĤPëPN°PsPjC- PICÍPμCĤPμPrPëPSCr P±CíP»Pë
PIC-PrP·PSP°C±PμPSC- C,P°PeC- P·PjC-PSPë: P·PSPëP¶PμPSPSCĬ
PμCĤPëC,CĤPsC†PëC,P°CĤPSPëC... PíP°CĤP°PjPμC,CĤC-PI C- P·PjC-PSPë PI
PSPëCĤPeP°C... (PsP±PëPrPIP° C†C- PμC,,PμPeC,Pë PjPsP¶CíC,CH P±CíC,Pë
PIPëPeP»PëPeP°PSC- PsP»PjPμCÍP°CĤC,P°PSCr PjPμPrPsPeCÍPsPjC-P»PsPj), P·PjC-PSPë
PI PePëC€PμC±PSPëPeCí (P·P±C-P»CHC€PμPSPSCĬ PíCĤPsCÍPIC-C,Cí C-
PrPëC,,CíP·PSPμ PíPsC,PsPIC%oPμPSPSCĬ CÍP»PëP·PsPIPsC— PsP±PsP»PsPSPëPë
PeP»CíP±PsPIPsC— C,P° PsP±PsPrPsPIPsC— PePëC€PePë), PSP°PrPSPëCĤPePsPIPëC...
P·P°P»PsP· (PiC-PíPμCĤC,CĤPsC,,C-Ĭ PeP»C-C,PëPS PeP»CíP±PsC±PePsPIPsC—
P·PsPSPë PePsCĤPë C- PIP°PeCíPsP»C-P·P°C†C-Ĭ PeP»C-C,PëPS PíCíC±PePsPIPsC—
P·PsPSPë), P° C,P°PePsP¶ PiC-PíPμCĤC,CĤPsC,,C-Ĭ PíCĤPsC,PsPe PjPsP»PsC±PSPëC...
P·P°P»PsP·, ĬPeP° PjPsP¶Pμ P±CíC,Pë PIPëPeP»PëPeP°PSP° P°PjP»PsPrPëPíC-PSPsPj.
PíC- P·PjC-PSPë PSPμ PrPsPíPsPIPsCĤCĤC,CH CĤP°PSC-C€Pμ PsC,CĤPëPjP°PSC-
PrP°PSC- PíCĤPs C,PsPeCÍPëC±PSC-CÍC,CH PsPeCĤPμPjPëC... PePsPjPíPsPSPμPSC,C-PI
PíCĤPμPíP°CĤP°C,Cí C- PSPμ CÍPIC-PrC±P°C,CH PíCĤPs PíPsCĬPICr PSPsPIPëC...
C,PsPeCÍPëC±PSPëC... PμC,,PμPeC,C-PI P°P±Ps PíCĤPs PSP°ĬPIPsC-CÍC,CH
CÍPëPSPμCĤPiC-C±PSPsC— C,PsPeCÍPëC±PSPsCÍC,C-.

*PĥP»PjPμCÍP°CĤC,P°PSCé PjPμPrPsPeCÍPsPjC-P» (PrC-CHC±P° CĤPμC±PsPIPëPSP°
PŸPμPIC-PeP°CĤCé)*

PJ PrPsCÍP»C-PrP¶PμPSPSCĬC... C...CĤPsPSC-C±PSPsC— C,PsPeCÍPëC±PSPsCÍC,C-
PSP° C%oCíCĤP°C... C- CÍPsP±P°PeP°C... PμC,,PμPeC,Pë PsP»PjPμCÍP°CĤC,P°PSCr
PjPμPrPsPeCÍPsPjC-P»Cí P±CíP»Pë CÍC...PsP¶C- P· C,P°PePëPjPë C-PSC€PëC...
P°PSC,P°PiPsPSC-CÍC,C-PI CĤPμC†PμPíC,PsCĤC-PI AT₁ C,P° C-PSPiC-P±C-C,PsCĤC-PI
PĥPμPα: PíC-PrPIPëC%oPμPSPSCĬ CĤC-PIPsCĬ CÍPμC±PsPIPëPSPë (BUN) C-
PeCĤPμP°C,PëPSC-PSCr PI PeCĤPsPIC-, P·PSPëP¶PμPSPSCĬ PjP°CÍPë CÍPμCĤC†Ĭ,
P·PSPëP¶PμPSPSCĬ PμCĤPëC,CĤPsC†PëC,P°CĤPSPëC... PíP°CĤP°PjPμC,CĤC-PI
(PePsPSC†PμPSC,CĤP°C†C-Ĭ PμCĤPëC,CĤPsC†PëC,C-PI C- PiPμPjPsPiP»PsP±C-PSCr,
PiPμPjP°C,PsPeCĤPëC,), PiC-CÍC,PsP»PsPiC-C±PSC- PsP·PSP°PePë CíCĤP°P¶PμPSPSCĬ
PSPëCĤPsPe (CĤPμPiPμPSPμCĤP°C,PëPIPsC- CíCĤP°P¶PμPSPSCĬ PSPëCĤPePsPIPsPiPs
PμPíC-C,PμP»C-CĤ, PíPsC,PsPIC%oPμPSPSCĬ P±P°P·P°P»CHPSPsC—
PjPμPjP±CĤP°PSPë, CĤPsP·C€PëCĤPμPSPSCĬ PeP°PSP°P»CHC†C-PI). PíC-
PíPsP±C-C±PSC- CĤPμP°PeC†C-C—, CÍPíCĤPëC±PëPSPμPSC-
C,,P°CĤPjP°PePsP»PsPiC-C±PSPsCĤ PrC-C”CĤ PsP»PjPμCÍP°CĤC,P°PSCr
PjPμPrPsPeCÍPsPjC-P»Cí, CÍPíPsCÍC,PμCĤC-PiP°P»PëCÍCĬ C,P°PePsP¶ Cí
PrPsPeP»C-PSC-C±PSPëC... PrPsCÍP»C-PrP¶PμPSPSCĬC... P· C-PSC€PëPjPë
P°PSC,P°PiPsPSC-CÍC,P°PjPë CĤPμC†PμPíC,PsCĤP° AT₁ C,P° C-PSPiC-P±C-C,PsCĤP°PjPë
PĥPμPα C- PjPsP¶CíC,CH P±CíC,Pë P·PjPμPSC€PμPSC- C€P»ĬĬC...PsPj
PíPμCĤPsCĤP°P»CHPSPsPiPs PíCĤPëPN°PsPjCí PSP°C,CĤC-CĤ C...P»PsCĤPëPrCí. PJ
PsP±PsC... PIPëPrC-PI C,PIP°CĤPëPS CÍPíPsCÍC,PμCĤC-PiP°P»Pë PíC-PrPIPëC%oPμPSCr
P°PeC,PëPIPsC-CÍC,CH CĤPμPSC-PSCr PI PíP»P°P·PjC- C,P°
PiC-PíPμCĤC,CĤPsC,,C-CĤ/PiC-PíPμCĤPíP»P°P·C-CĤ
CĤPeCÍC,P°PiP»PsPjPμCĤCíP»ĬĬCĤPSPëC... PeP»C-C,PëPS PSPëCĤPsPe. PíC-
P·PjC-PSPë, ĬPeC- C” C,PëPíPsPIPëPj PμC,,PμPeC,PsPj PeP»P°CÍCí
C-PSPiC-P±C-C,PsCĤC-PI PĥPμPα C,P° C-PSC€PëC... P°PSC,P°PiPsPSC-CÍC,C-PI

CḤPμC†PμPīC,PsCḤP° AT₁, PNePjPsPIC-CḤPSPs, PSPμ PjP°CḤC,Ch PeP»C-PSC-C†PSPsC—
P·PSP°C†CřC%PsCřC,C-.

PμPsPrC-P±PSPs PrPs C-PSC€PëC... P°PSC,P°PiPsPSC-CřC,C-PI CḤPμC†PμPīC,PsCḤP°
AT₁, PsP»PjPμCřP°CḤC,P°PSCř PjPμPrPsPeCřPsPjC-P» P·P±C-P»ChC€CřC”
C†P°CřC,PsC,Cř C...CḤPsPjPsCřPsPjPSPëC... CḤPsP·CḤPëPIC-PI Cř PeCřP»ChC,CřCḤC-
PeP»C-C,PëPS *in vitro*. PḥPrPSP°Pe PīPsPrC-P±PSC- PμC,,PμPeC,Pë P±CřP»Pë
PIC-PrC,PIPScḤPμPSC- Cř PeC-P»ChPePsC... PrPsCřP»C-PrPḡPμPSPSCC... *in vivo*,
PrPμ PsP»PjPμCřP°CḤC,P°PSCř PjPμPrPsPeCřPsPjC-P» P·P°CřC,PsCřPsPICřPIP°P»Pë PI
PrCřPḡPμ PIPëCřPsPePëC... PīPμCḤPsCḤP°P»ChPSPëC... PrPsP·P°C..., P°Pḡ PrPs
2000 PjPi/PePi. P—P°PiP°P»PsPj CḤPμP·CřP»ChC,P°C,Pë PICřPμP±C-C†PSPsPiPs
PrPsCřP»C-PrPḡPμPSPSCC PSP° PiPμPSPsC,PsPeCřPëC†PSC-CřC,Ch
PiPsPIPsCḤCIC,Ch PiCḤPs C,Pμ, C%Ps PiPμPSPsC,PsPeCřPëC†PSP° PrC-CIC
PsP»PjPμCřP°CḤC,P°PSCř PīCḤPë PeP»C-PSC-C†PSPsPjCř P·P°CřC,PsCřCřPIP°PSPSC-
PjP°P»PsPNePjPsPIC-CḤPSP°.

P’ CḤP°PjPeP°C... 2-CḤC-C†PSPsPiPs PrPsCřP»C-PrPḡPμPSPSCC PSP° C%CřCḤP°C...
P°P±Ps 6-PjC-CřCIC†PSPsPiPs PrPsCřP»C-PrPḡPμPSPSCC
PeP°PSC†PμCḤPsPiPμPSPSPScřC,C- PSP° C,CḤP°PSCřPiPμPSPSPëC... PjPëC€P°C... PSPμ
P±CřP»Ps PIPëCICPIP»PμPSPs PeP°PSC†PμCḤPsPiPμPSPSPëC...
PIP»P°CřC,PëPIPsCřC,PμPNe PsP»PjPμCřP°CḤC,P°PSCř PjPμPrPsPeCřPsPjC-P»Cř.

PJ PrPsCřP»C-PrPḡPμPSPSCC... C,PsPeCřPëC†PSPsCřC,C- C%PsPrPs
CḤPμPīCḤPsPrCřPeC,PëPIPSPëC... PsCḤPiP°PSC-PI Cř C%CřCḤC-PI
PsP»PjPμCřP°CḤC,P°PSCř PjPμPrPsPeCřPsPjC-P» PSPμ PIPīP»PëPIP°PI PSP°
C,,PμCḤC,PëP»ChPSC-CřC,Ch C- PSPμ C†PëPSPëPI C,PμCḤP°C,PsPiPμPSPSPsC—
PrC-C—. PīPe C- Cř CḤP°P·C- P·P°CřC,PsCřCřPIP°PSPSCC C-PSC€PëC...
P°PSC,P°PiPsPSC-CřC,C-PI CḤPμC†PμPīC,PsCḤC-PI P°PSPiC-PsC,PμPSP·PëPSCř II,
PīC-CřP»CIC PμPeCřPīPsP·PëC†C-C— PsP»PjPμCřP°CḤC,P°PSCř
PjPμPrPsPeCřPsPjC-P»Cř PIPëPḡPëPIP°PSPSCC PīPsC,PsPjCřC,PIP° P±CřP»P°
P·PSPëPḡPμPSP°, P° Cř CřP°PjPsPe, CIPeC- PsC,CḤPëPjCřPIP°P»Pë PīCḤPμPīP°CḤP°C,
PSP° PīC-P·PSC-C... C,PμCḤPjC-PSP°C... PIP°PiC-C,PsPScřC,C- C- PīC-Pr C†P°Cř
P»P°PeC,P°C†C-C—, PIC-PrP·PSP°C†P°P»P°CřCIC PrPëP»P°C,P°C†C-CIC
PSPëCḤPePsPIPëC... PjPëCřPsPe. PīPe C- C-PSC€C- PiC-PīPsC,PμPSP·PëPIPSC-
PīCḤPμPīP°CḤP°C,Pë, PsP»PjPμCřP°CḤC,P°PSCř PjPμPrPsPeCřPsPjC-P» P±CřPI
P±C-P»ChC€ C,PsPeCřPëC†PSPëPNe PrP»CIC PIP°PiC-C,PSPëC... PeCḤPsP»C-PI, PSC-Pḡ
PrP»CIC PIP°PiC-C,PSPëC... C%CřCḤC-PI, PīCḤPsC,Pμ PSPμ C†PëPSPëPI PīCḤPë
C†ChPsPjCř C,,PμC,PsC,PsPeCřPëC†PSPsC— PrC-C—.

PḥPjP»PsPrPëPiC-PS (PrC-CḤC†P° CḤPμC†PsPIPëPSP° PŸPμPIC-PeP°CḤCž)

P PμPīCḤPsPrCřPeC,PëPIPSP° C,PsPeCřPëC†PSC-CřC,Ch

P”PsCřP»C-PrPḡPμPSPSCC CḤPμPīCḤPsPrCřPeC,PëPIPSPsC— C,,CřPSPeC†C-C— PSP°
C%CřCḤP°C... C- PjPëC€P°C... PIPëCICPIPëP»Pë P·P°C,CḤPëPjPeCř PSP°CřC,P°PSPSCC
PīPsP»PsPiC-PI, P·P±C-P»ChC€PμPSPSCC C†P°CřCř PīPμCḤPμPNePjC-PI C-
P·PjPμPSC€PμPSPSCC PIPëPḡPëPIP°PSPSCC PīPsC,PsPjCřC,PIP° PīCḤPë
PīCḤPëPNePsPjC- PrPsP·, PīCḤPëP±P»PëP·PSPs PI 50 CḤP°P·C-PI P±C-P»ChC€PëC...,

PSC-P¶ PjP°PeCΓPëPjP°P»CHΨPSP° CΨPμPePsPjPμPSPPrPsPIP°PSP° PrPsP·P° PrP»CΠ P»CΨPrPëPSPë, CΠPeP° T'CΨCΓPSC,CrC"C,CΨCΓCΠ PSP° CΨPsP·CΨP°C...CrPSPeCΓ PSP° 1 PePi PjP°CΓPë C,C-P»P° (PjPi/PePi).

PμPsCΨCΓC€PμPSPSCΠ C,,PμCΨC,PëP»CHΨPSPsCΓC,C-

P'PiP»PëPICr PSP° C,,PμCΨC,PëP»CHΨPSC-CΓC,CΨ Cr C%CrCΨC-PI, CΠPeC-PsC,CΨPëPjCΓPIP°P»Pë P°PjP»PsPrPëPiC-PS (CΓP°PjC†C- PiCΨPsC,CΠPiPsPj 64 PrPSC-PI, CΓP°PjPePë P·P° 14 PrPSC-PI PrPs CΓPiP°CΨCΨPIP°PSPSCΠ) PI PrPsP·P°C... PrPs 10 PjPi/ PePi/ PrPsP±Cr (Cr 8 CΨP°P·C-PI (CΠPeC%Ps PjP°CΓP° C,C-P»P° PiP°C†C-C"ΨC,P° PrPsCΨC-PIPSCΨC" 50 PePi) PiPμCΨPμPIPëC%CrC" PjP°PeCΓPëPjP°P»CHΨPSCr CΨPμPePsPjPμPSPPrPsPIP°PSCr PrPsP·Cr PrP»CΠ P»CΨPrPëPSPë, C%Ps CΓC,P°PSPsPIPëC,CΨ 10 PjPi PI PiPμCΨPμCΨP°C...CrPSPeCΓ PSP° PjPi/Pj²), PIPëCΠPIP»PμPSPs PSPμ P±CrP»Ps. P' C...PsPrC- C-PSC€PsPiPs PrPsCΓP»C-PrP¶PμPSPSCΠ, PI CΠPePsPjCΓ CΓP°PjC†C- C%CrCΨC-PI PsC,CΨPëPjCΓPIP°P»Pë P°PjP»PsPrPëPiC-PSCr P±PμCΓPëP»P°C, PiCΨPsC,CΠPiPsPj 30 PrPSC-PI Cr PrPsP·P°C..., PiPsCΨC-PIPSCΠPSPSPëC... P· PrPsP·PsCΨ PrP»CΠ P»CΨPrPëPSPë PI PiPμCΨPμCΨP°C...CrPSPeCΓ PSP° PjPi/Pj², P±CrP»Ps PIC-PrP·PSP°C†PμPSPs P·PSPëP¶PμPSPSCΠ PePsPSC†PμPSC,CΨP°C†C-PMë C,,PsP»C-PeCΓP»PsCΓC,PëPjCΓP»CΨCΨC†PsPiPs PiPsCΨPjPsPSCr C- C,PμCΓC,PsCΓC,PμCΨPsPSCr PI PiP»P°P·PjC-, P° C,P°PePsP¶ P·PSPëP¶PμPSPSCΠ C%CrC-P»CHΨPSPsCΓC,C- CΓPiPμCΨPjPë, P·PjPμPSC€PμPSPSCΠ PeC-P»CHΨPePsCΓC,C- P·CΨC-P»PëC... CΓPiPμCΨPjP°C,PëPr C- PeP»C-C,PëPS PŸPμCΨC,PsP»C-.

PμP°PSC†PμCΨPsPiPμPSPμP·, PjCΓC,P°PiPμPSPμP·

P'PëPiCΨPsP±CrPIP°PSPSCΠ PSP° C%CrCΨP°C... C- PjPëC€P°C..., CΠPeC-PsC,CΨPëPjCΓPIP°P»Pë P°PjP»PsPrPëPiC-PS P· C—P¶PμCΨ PiCΨPsC,CΠPiPsPj PrPIPpsC... CΨPsPeC-PI Cr PePsPSC†PμPSC,CΨP°C†C-CΠC..., CΨPsP·CΨP°C...PsPIP°PSPëC... PrP»CΠ PIC-PrC,PIPsCΨPμPSPSCΠ PrPsP· 0,5, 1,25 C- 2,5 PjPi/PePi/PrPsP±Cr, PSPμ PIPëCΠPIPëP»Pë PsP·PSP°Pe PeP°PSC†PμCΨPsPiPμPSPSPsCΓC,C-. PκP°PMëPIPëC%P° PrPsP·P° (PrP»CΠ PjPëC€PμPMë PμPePIC-PIP°P»PμPSC,PSP° PjP°PeCΓPëPjP°P»CHΨPSC-PMë CΨPμPePsPjPμPSPPrPsPIP°PSC-PMë PrPsP·C- 10 PjPi PI PiPμCΨPμCΨP°C...CrPSPeCΓ PSP° PjPi/Pj², P° PrP»CΠ C%CrCΨC-PI – PI PrPIP° CΨP°P·Pë PIPëC%Pμ PjP°PeCΓPëPjP°P»CHΨPSPsC— CΨPμPePsPjPμPSPPrPsPIP°PSPsC— PrPsP·Pë) P±CrP»P° P±P»PëP·CHΨPeP° PrPs PjP°PeCΓPëPjP°P»CHΨPSPsC— PiPμCΨPμPSPsCΓPëPjPsC— PrPsP·Pë PrP»CΠ PjPëC€PμPMë, P°P»Pμ PSPμ PrP»CΠ C%CrCΨC-PI.

P"PsCΓP»C-PrP¶PμPSPSCΠ PjCΓC,P°PiPμPSPSPsCΓC,C- PSPμ PIPëCΠPIPëP»Pë PiPsPIBΨ™ CΠP·P°PSPëC... P· PiCΨPμPiP°CΨP°C,PsPj PμC,,PμPeC,C-PI PSP° CΨC-PIPSC-PiPμPSC-PI P°P±Ps C...CΨPsPjPsCΓPsPj.

PμP»C-PSC-C†PSC- C...P°CΨP°PeC,PμCΨPëCΓC,PëPePë.

PμPsPeP°P·P°PSPSCΠ.

P>C-PeCfPIP°PSPSCЦ PμCfPμPSC†C-P°P»CHbPSPsC— PiC-PiPμCfC,PμPSP·C-C—.

PŸPμPIC-PeP°Cf PiPsPeP°P·P°PSPëPNë PrPsCfPsCfP»PëPj PiP°C†C-C”PSC,P°Pj, Cí CЦPePëC... PjPsPSPsC,PμCfP°PiC-CЦ PsP»PjPμCfP°CfC,P°PSCr PjPμPrPsPeCfPsPjC-P»PsPj P°P±Ps P°PjP»PsPrPëPiC-PSPsPj PSPμ P·P°P±PμP·PiPμC†CfC” PSPμPsP±C...C-PrPSPsPiPs PePsPSC,CfPsP»Cf P°CfC,PμCfC-P°P»CHbPSPsPiPs C,PëCfPeCf (PrPëPI. CfPsP·PrC-P»Pë B«PŸPiPsCfC-P± P·P°CfC,PscfCfPIP°PSPSCЦ C,P° PrPsP·PëB», B«PαP°CfPjP°PePsPrPëPSP°PjC-PeP°B»).

PμCfPsC,PëPiPsPeP°P·P°PSPSCЦ.

PμC-PrPIPëC%PμPSP° C†CfC,P»PëPIC-CfC,CH PrPs PrC-CfC†PëC... CfPμC†PsPIPëPS, PiPsC...C-PrPSPëC... PrPëPiC-PrCfPsPiC-CfPëPrPëPSCr P°P±Ps P±CfPrCHb-CЦPePsC— P· PrPsPiPsPjC-P¶PSPëC... CfPμC†PsPIPëPS (PrPëPI. CfPsP·PrC-P» B«PŸPeP»P°PrB»).

- P°P°PiC-C,PSC-CfC,CH C,P° PiP»P°PSCrPIP°PSPSCЦ PIP°PiC-C,PSPsCfC,C- (PrPëPI. CfPsP·PrC-P»Pë B«PfcfPsP±P»PëPiPsCfC,C- P·P°CfC,PscfCfPIP°PSPSCЦB», B«P—P°CfC,PscfCfPIP°PSPSCЦ Cí PiPμCfC-PsPr PIP°PiC-C,PSPsCfC,C- P°P±Ps PiPsPrCfPIP°PSPSCЦ PiCfCfPrPrCfB»).

- PŸCЦP¶PeP° PiPμC†C-PSPePsPIP° PSPμPrPsCfC,P°C,PSC-CfC,CH C,P° PSPμPiCfPsC...C-PrPSC-CfC,CH P¶PsPIC†PSPëC... CëP»CЦC...C-PI (PrPëPI. CfPsP·PrC-P» B«PαP°CfPjP°PePsPeC-PSPμC,PëPeP°B»).

- PfcPrPSPsC†P°CfPSPμ P·P°CfC,PscfCfPIP°PSPSCЦ PiCfPμPiP°CfP°C,Cf PŸPμPIC-PeP°Cf C,P° P»C-PeP°CfCfCHbPePëC... P·P°CfPsP±C-PI, C%Ps PjC-CfC,CЦC,CH P°P»C-CfPeC-CfPμPS, PiCfPsC,PëPiPsPeP°P·P°PSPμ PiP°C†C-C”PSC,P°Pj P· C†CfPeCfPsPIPëPj PrC-P°P±PμC,Pspj P°P±Ps PiPsCfCfCëPμPSPSCЦPj C,,CfPSPeC†C-C— PSPëCfPsPe (PëPлPα < 60 PjP»/C...PI/1,73 Pj²) (PrPëPI. CfPsP·PrC-P»Pë B«P’P·P°C”PjPsPrC-CЦ P· C-PSCëPëPjPë P»C-PeP°CfCfCHbPePëPjPë P·P°CfPsP±P°PjPë C,P° C-PSCëC- PIPëPrPë PIP·P°C”PjPsPrC-PNëB», B«PαP°CfPjP°PePsPrPëPSP°PjC-PeP°B»).

PSPμCfPμP· PSP°CЦPIPSC-CfC,CH Cí CfPeP»P°PrC- P°PjP»PsPrPëPiC-PSCr PiCfPμPiP°CfP°C, PŸPμPIC-PeP°Cf C,P°PePsP¶ PiCfPsC,PëPiPsPeP°P·P°PSPëPNë PiP°C†C-C”PSC,P°Pj, Cí CЦPePëC...:

- C,CЦP¶PeP° P°CfC,PμCfC-P°P»CHbPSP° PiC-PiPsC,PμPSP·C-CЦ;

- CëPsPe (PIPeP»CfC†P°CfC†Pë PeP°CfPrC-PsPiPμPSPSPëPNë CëPsPe);

- PiPsCfCfCëPμPSPSCЦ PIC-PrC,PspëCf PeCfPsPIC- C-P· P»C-PIPsPiPs CëP»CfPSPsC†PeP° (PSP°PiCfPëPeP»P°Pr, PiCfPë CfC,PμPSPsP·C- P°PsCfC,Pë C,CЦP¶PePsPiPs CfC,CfPiPμPSCЦ);

- PiPμPjPsPrPëPSP°PjC-C†PSPs PSPμCfC,P°P±C-P»CHbPSP° CfPμCfC†PμPIP° PSPμPrPsCfC,P°C,PSC-CfC,CH PiC-CfP»CЦ PiPsCfC,CfPsPiPs C-PSC,,P°CfPeC,Cf PjC-PsPeP°CfPrP°.

B

P'P·P°C" PjPsPrC-CI P· C-PSC€PëPjPë P»C-PeP°CĤCÍCHPePëPjPë P·P°CÍPsP±P°PjPë C,P° C-PSC€C- PIPëPrPë PIP·P°C" PjPsPrC-PN.

PμPsC,PμPSC†C-PN°PSC- PIP·P°C" PjPsPrC-C—, PIPëPeP»PëPeP°PSC- P'PsC" PrPSP°PSPSCIPj P· P»C-PeP°CĤCÍCHPePëPj P·P°CÍPsP±PsPj PŸPμPIC-PeP°CĤ

PŸP»C-Pr P±CÍC,Pë PsP±PμCĤPμP¶PSPëPj PíCĤPë PsPrPSPsC±P°CÍPSPsPjCÍ P·P°CÍC,PsCÍCÍPIP°PSPSC-

P†PSC€C- PiC-PiPsC,PμPSP·PëPIPSC- P·P°CÍPsP±Pë

P"C-PiPsC,PμPSP·PëPIPSPëPN° PμC,,PμPeC, PíCĤPμPíP°CĤP°C,CÍ PŸPμPIC-PeP°CĤ PjPsP¶Pμ P±CÍC,Pë P'PsCÍPëP»PμPSPëPN° PíCĤPë PsPrPSPsC±P°CÍPSPsPjCÍ P·P°CÍC,PsCÍCÍPIP°PSPSC- C-PSC€PëC... PiC-PiPsC,PμPSP·PëPIPSPëC... P»C-PeP°CĤCÍCHPePëC... P·P°CÍPsP±C-PI (PSP°PíCĤPëPeP»P°Pr P°P»CHC,,P°-P±P»PsPeP°C,PsCĤC-PI, PrC-CÍCĤPμC,PëPeC-PI).

PμPsC,PμPSC†C-PN°PSC- PIP·P°C" PjPsPrC-C—, P'PsPIeĤ™ CIIP·P°PSC- P· PrC-CĤC±PsCĤ CĤPμC±PsPIPëPSPsCĤ PíCĤPμPíP°CĤP°C,Cž PŸPμPIC-PeP°CĤ PsP»PjPμCÍP°CĤC,P°PSCž PjPμPrPsPeCÍPsPjC-P»PsPj

PŸCÍPíCÍC,PSC" P·P°CÍC,PsCÍCÍPIP°PSPSCIP PSPμ CĤPμPePsPjPμPSPPrCÍC"C,CHCÍCI

P†PSPiC-P±C-C,PsCĤPë PĥPμPα, P±P»PsPeP°C,PsCĤPë CĤPμC†PμPíC,PsCĤC-PI P°PSPiC-PsC,PμPSP·PëPSCÍ P†P† P°P±Ps P°P»C-CÍPeC-CĤPμPS

P"°PSC- PeP»C-PSC-C±PSPëC... PIPëPíCĤPsP±CÍPIP°PSCĤ P'PsPeP°P·CÍCĤC,CH, C%°Ps P'PsPrPIC-PN°PSP° P±P»PsPeP°PrP° CĤPμPSC-PS-P°PSPiC-PsC,PμPSP·PëPS- P°P»CHPrPsCÍC,PμCĤPsPSPsPíPsC— CÍPëCÍC,PμPjPë (P PĥPĥPŸ), P'PsPIBĤ™ CIIP·P°PSP° P·C- CÍPíC-P»CHPSPëPj P·P°CÍC,PsCÍCÍPIP°PSPSCIPj C-PSPiC-P±C-C,PsCĤC-PI PĥPμPα C,P° P±P»PsPeP°C,PsCĤC-PI CĤPμC†PμPíC,PsCĤC-PI P°PSPiC-PsC,PμPSP·PëPSCÍ II P°P±Ps P°P»C-CÍPeC-CĤPμPSCÍ, PíCĤPëP·PIPPrPëC,CH PrPs PíC-PrPIPëC%°PμPSPSCIP C±P°CÍC,PsC,Pë PSPμP±P°P¶P°PSPëC... CIPIPëC%°, C,P°PePëC... CIPe P°CĤC,PμCĤC-P°P»CHPSP° PiC-PiPsC,PμPSP·C-CI, PiC-PiPμCĤPeP°P»C-C" PjC-CI C- P·PSPëP¶PμPSPSCIP C,,CÍPSPeC†C-C— PSPëCĤPsPe (PI C,PsPjCÍ C±PëCÍP»C- PíPsCÍC,CĤP° PSPëCĤPePsPIP° PSPμPrPsCÍC,P°C,PSC-CÍC,CH), PI P'PsCĤC-PIPSCIPSPSC- C-P· P·P°CÍC,PsCÍCÍPIP°PSPSCIPj PsPrPSPsPiPs P·P°CÍPsP±CÍ, CIPePëPN° PrC-C" PSP° P PĥPĥPŸ (PrPëPI. CĤPsP·PrC-P»Pë B«PμCĤPsC,PëP'PsPeP°P·P°PSPSCIPB», B«PĥCÍPsP±P»PëPIPSCÍC,C- P·P°CÍC,PsCÍCÍPIP°PSPSCIPB», B«PαP°CĤPjP°PePsPrPëPSP°PjC-PeP°B»).

PμCĤPμPíP°CĤP°C,Pë, C%°Ps PIPíP»PëPIP°CĤC,CH PSP° CĤC-PIPμPSCĤ PeP°P»C-CĤ

PĥPrPSPsC±P°CÍPSPμ P·P°CÍC,PsCÍCÍPIP°PSPSCIP P· PeP°P»C-PN°P·P±PμCĤC-PiP°CĤC±PëPjPë PrC-CÍCĤPμC,PëPeP°PjPë, PeP°P»C-C" PIPëPjPë PrPsP±P°PIPeP°PjPë, P·P°PjC-PSPSPëPeP°PjPë CÍPsP»C-, C%°Ps PjC-CÍC,CIIC,CH PeP°P»C-PN°, P°P±Ps C-PSC€PëPjPë P»C-PeP°CĤCÍCHPePëPjPë P·P°CÍPsP±P°PjPë, CIPeC- PjPsP¶CÍC,CH PíC-PrPIPëC%°PëC,Pë CĤC-PIPμPSCĤ PeP°P»C-CĤ (PSP°PíCĤPëPeP»P°Pr, PiPμPíP°CĤPëPS, C-PSPiC-P±C-C,PsCĤPë PĥPμPα), PjPsP¶Pμ

PiCtjPep·PIPμCfC, Pē PrPs P·P±C-P»CHCεPμPSPSCl PePsPSCtPμPSC, CtjP°CtC-C—
PeP°P»C-Ct PI CfPēCtjPsPIP°C, CtC- PeCtjPsPIC- (PrPēPI. CtjPsP·PrC-P»
B«PhCfPsP±P»PēPIPsCfC, C- P·P°CfC, PsCfCfPIP°PSPSClB»). PμCtjPē
PiCtjPep·PSP°CtPμPSPSC- PiCtjPμPiP°CtjP°C, C-PI, C%oPs PIPiP»PēPIP°CtC, CH PSP°
CtjC-PIPμPSCt PeP°P»C-Ct, PI PiPsC”PrPSP°PSPSC- P· PiCtjPμPiP°CtjP°C, PsPj
PŸPμPIC-PeP°Ct CtjPμPePsPjPμPSPPrCfC”C, CHCfCfC PePsPSC, CtjPsP»CH
CfPēCtjPsPIP°C, PePsPIPSC— PePsPSCtPμPSC, CtjP°CtC-C— PeP°P»C-Ct.

PμCtjPμPiP°CtjP°C, Pē P»C-C, C-Ct

PμCtjPē PsPrPSPsCtP°CfPSPsPjCf P·P°CfC, PsCfCfPIP°PSPSC- P»C-C, C-Ct P·
C-PSPiC-P±C-C, PsCtjP°PjPē
P°PSPiC-PsC, PμPSP·PēPSPiPμCtjPμC, PIPsCtjCtjPIP°P»CHPSPsPiPs C,, PμCtjPjPμPSC, Cf
C- CtjC-PrPePs P· P°PSC, P°PiPsPSC-CfC, P°PjPē CtjPμCtPμPiC, PsCtjC-PI
P°PSPiC-PsC, PμPSP·PēPSCf II CfPiPsCfC, PμCtjC-PiP°P»PsCfCfC PsP±PsCtjPsC, PSPμ
P·P±C-P»CHCεPμPSPSCl CfPēCtjPsPIP°C, PePsPIPSC— PePsPSCtPμPSC, CtjP°CtC-C—
P»C-C, C-Ct C- C, PsPeCfPēCtPSPsCfC, C- PnPsPiPs PiCtjPμPiP°CtjP°C, C-PI. Pj
P·PiBtj™ ClP·PeCf P· CtPēPj PSPμ CtjPμPePsPjPμPSPPrCfC”C, CHCfCfC
PsPrPSPsCtP°CfPSPμ P·P°CfC, PsCfCfPIP°PSPSCl PiCtjPμPiP°CtjP°C, Cf
PŸPμPIC-PeP°Ct C, P° PiCtjPμPiP°CtjP°C, C-PI P»C-C, C-Ct (PrPēPI. CtjPsP·PrC-P»
B«PhCfPsP±P»PēPIPsCfC, C- P·P°CfC, PsCfCfPIP°PSPSClB»). PμCtjPē
PSPμPsP±C...C-PrPSPsCfC, C- PsPrPSPsCtP°CfPSPsPiPs P·P°CfC, PsCfCfPIP°PSPSCl
PiCtjPμPiP°CtjP°C, Cf PŸPμPIC-PeP°Ct C, P° PiCtjPμPiP°CtjP°C, C-PI P»C-C, C-Ct
CtjPμPePsPjPμPSPPrCfC”C, CHCfCfC CtjPμPiCfP»ClCtjPSPēPn PePsPSC, CtjPsP»CH
CtjC-PIPSCl P»C-C, C-Ct PI CfPēCtjPsPIP°C, CtC-.

PŸPiC-P»CHPSPμ P·P°CfC, PsCfCfPIP°PSPSCl PIPēPjP°PiP°C”
PsP±PμCtjPμP¶PSPsCfC, C-

PkPμCfC, PμCtjPsC—PrPSC- PiCtjPsC, PēP·P°PiP°P»CHPSC- P·P°CfCfPsP±Pē (PkPμP—P—),
PIPeP»CtCtP°CtCtPē CfPμP»PμPeC, PēPIPSC- C-PSPiC-P±C-C, PsCtjPē PjPhP”-2,
P°CtPμC, PēP»CfP°P»C-CtPēP»PsPICf PePēCfP»PsC, Cf (> 3 Pi/PrPsP±Cf) C-
PSPμCfPμP»PμPeC, PēPIPSC- PkPμP—P—

PμCtjPē PiCtjPep·PSP°CtPμPSPSC- P°PSC, P°PiPsPSC-CfC, C-PI
P°PSPiC-PsC, PμPSP·PēPSCf II CfPiC-P»CHPSPs P· PkPμP—P— PjPsP¶P»PēPIPμ
PiPsCfP»P°P±P»PμPSPSCl PiC-PiPsC, PμPSP·PēPIPSPsPiPs PμC,, PμPeC, Cf. PnCtjC-Pj
C, PsPiPs, PsPrPSPsCtP°CfPSPμ P·P°CfC, PsCfCfPIP°PSPSCl P°PSC, P°PiPsPSC-CfC, C-PI
P°PSPiC-PsC, PμPSP·PēPSCf II C- PkPμP—P— PjPsP¶Pμ PiC-PrPIPēC%oCfPIP°C, Pē
CtjPep·PēPe PiPsPiC-CtjCεPμPSPSCl C,, CfPSPeCtC-C— PSPēCtjPsPe C-
PiCtjPep·PIPsPrPēC, Pē PrPs P·P±C-P»CHCεPμPSPSCl CfPēCtjPsPIP°C, PePsPIPSC—
PePsPSCtPμPSC, CtjP°CtC-C— PeP°P»C-Ct. PŸPsPjCf PiCtjPē C, P°PeC-Pn
CfPiC-P»CHPSC-Pn C, PμCtjP°PiC-C— CtjPμPePsPjPμPSPPrCfC”C, CHCfCfC
CtjPμPiCfP»ClCtjPSPs PsCtC-PSCtjPIP°C, Pē C,, CfPSPeCtC-Ct PSPēCtjPsPe, P°
C, P°PePsP¶ CfC, PμP¶PēC, Pē P·P° PSP°P»PμP¶PSPsCt PiC-PrCtjP°C, P°CtC-C”Ct
PiP°CtC-C”PSC, P°.

P»C-PeP°CtjCfCHPePēPn P·P°CfC-P± PePsP»PμCfPμPIPμP»P°Pj, C%oPs

P·PIBTM C_{II}P·C_rC” P[¶]PsPIC_‡PSC- PePëC_IP»PsC,Pë

PhPrPSPsC_‡P[°]C_IPSP_μ P·P[°]C_IC,P_sC_IC_rPIP[°]PSPSC_{II} PePsP»P_μC_IP_μPIP_μP»P[°]P_jC_r
PiC-PrC_{II}PsC...P»PsC_{II}PëPrC_r, C_%Ps P·PIBTM C_{II}P·C_rC” P[¶]PsPIC_‡PSC- PePëC_IP»PsC,Pë,
P·P_jP_μPSC€C_rC” C_IPëC_IC,P_μP_jPSPëPN_ë PIP_iP»PëPI C- P_iC-PePsPIC_r
PePsPSC_‡P_μPSC,C_{II}P[°]C_‡C-C_{II} PsP»P_jP_μC_IP[°]C_{II}C,P[°]PSC_r PI P_iP»P[°]P·P_jC-, P[°] C,P[°]PePsP[¶]
P·PSPëP[¶]C_rC” P_iP_μC_{II}C-PsPr PSP[°]P_iC-PIPIPëPIP_μPrP_μPSPSC_{II}. P_μC_{II}PëPN_ëPsP_j
PsP»P_jP_μC_IP[°]C_{II}C,P[°]PSC_r P_jP_μPrPsPeC_IP_sP_jC-P»C_r C_{II}Pe P_jC-PSC-P_jC_rP_j P·P[°] 4
PiPsPrPëPSPë PrPs P_iC_{II}PëPN_ëPsP_jC_r PePsP»P_μC_IP_μPIP_μP»P[°]P_jC_r
PiC-PrC_{II}PsC...P»PsC_{II}PëPrC_r P·PSPëP[¶]C_rPIP[°]PI P_μC,,P_μPeC, P»C-PeP[°]C_{II}C_IC_{II}PePsC—
PIP·P[°]C”P_jPsPrC-C—. P_YP»C-Pr C_{II}PsP·PiP»C_{II}PSC_rC,Pë P_jPsP[¶]P»PëPIC-C_IC,C_{II}
P_iC_{II}PëPN_ëPsP_jC_r PsP»P_jP_μC_IP[°]C_{II}C,P[°]PSC_r P_jP_μPrPsPeC_IP_sP_jC-P»C_r C_{II}Pe
P_jC-PSC-P_jC_rP_j P·P[°] 4 PiPsPrPëPSPë PrPs P_iC_{II}PëPN_ëPsP_jC_r PePsP»P_μC_IP_μPIP_μP»P[°]P_jC_r
PiC-PrC_{II}PsC...P»PsC_{II}PëPrC_r (PrPëPI. C_{II}PsP·PrC-P»
B«P^αP[°]C_{II}P_jP[°]PePsPeC-PSP_μC,PëPeP[°]B»).

P”PsPrP[°]C,PePsPIP[°] C-PSC,,PsC_{II}P_jP[°]C_‡C-C_{II}

P’C-PrP·PSP[°]C_‡P_μPSPs P_iPsP_jC-C_{II}PSP_μ P·PSPëP[¶]P_μPSPSC_{II}
P±C-PsPrPsC_IC,C_rP_iPSPsC_IC,C- PsP»P_jP_μC_IP[°]C_{II}C,P[°]PSC_r P_jP_μPrPsPeC_IP_sP_jC-P»C_r
P_iC-C_IP»C_{II} P»C-PeC_rPIP[°]PSPSC_{II} P[°]PSC,P[°]C_‡PëPrP[°]P_jPë (P_jP[°]P_iPSC-C_{II} C,P[°]
P[°]P»C_{II}P_jC-PSC-C_{II} PiC-PrC_{II}PsPeC_IPëPrP[°]P_jPë).

PhP»P_jP_μC_IP[°]C_{II}C,P[°]PSC_r P_jP_μPrPsPeC_IP_sP_jC-P» P·PSP[°]C_‡C_rC_%P_μ PSP_μ
PIP_iP»PëPIP[°]C” PSP[°] C,,P[°]C_{II}P_jP[°]PePsPeC-PSP_μC,PëPeC_r C-
C,,P[°]C_{II}P_jP[°]PePsPrPëPSP[°]P_jC-PeC_r PIP[°]C_{II}C,,P[°]C_{II}PëPSC_r P[°]P±Ps
C,,P[°]C_{II}P_jP[°]PePsPeC-PSP_μC,PëPeC_r PrPëPiPsPeC_IPëPSC_r. PhPrPSPsC_‡P[°]C_IPSP_μ
P·P[°]C_IC,P_sC_IC_rPIP[°]PSPSC_{II} PsP»P_jP_μC_IP[°]C_{II}C,P[°]PSC_r P_jP_μPrPsPeC_IP_sP_jC-P»C_r P·
P_iC_{II}P[°]PIP[°]C_IC,P[°]C,PëPSPsP_j PSP_μ P_iC_{II}PëP·PIPsPrPëC,C_{II} PrPs PeP»C-PSC-C_‡PSPs
P·PSP[°]C_‡C_rC_%PëC... P·P_jC-PS C,,P[°]C_{II}P_jP[°]PePsPeC-PSP_μC,PëPePë C_‡PëC...
P_iC_{II}P_μP_iP[°]C_{II}P[°]C,C-PI C_r P·PrPsC_{II}PsPIPëC... PrPsP±C_{II}PsPIP_sP»C_{II}C_‡C-PI.

P_kP_μ PIPëC_{II}PIP»P_μPSPs PeP»C-PSC-C_‡PSPs P·PSP[°]C_‡C_rC_%PsC—
C-PSPiC-P±C_rC_{II}C_‡PsC— PrC-C— PsP»P_jP_μC_IP[°]C_{II}C,P[°]PSC_r PSP[°] C,,P_μC_{II}P_jP_μPSC,Pë
1A1/2, 2A6, 2C8/9, 2C19, 2D6, 2E1 C,P[°] 3A4 C_‡PëC,PsC...C_{II}PsP_jC_r P 450 C_r
P»C_{II}PrPëPSPë *in vitro*, C- PIC-PrP·PSP[°]C_‡P_μPSPëPN_ë P_jC-PSC-P_jP[°]P»C_{II}PSPëPN_ë P[°]P±Ps
PSC_rP»C_{II}PsPIPëPN_ë C-PSP_rC_rPeC_rC_{II}C_‡PëPN_ë P_μC,,P_μPeC, C_%PsPrPs
P[°]PeC,PëPIPsC_IC,C- C_‡PëC,PsC...C_{II}PsP_jC_r P 450 C_r C_%C_rC_{II}C-PI. P_YP[°]PePëP_j
C_‡PëPSPsP_j, PeP»C-PSC-C_‡PSPs P·PSP[°]C_‡C_rC_%PëC... PIP·P[°]C”P_jPsPrC-PN_ë P_jC-P[¶]
PsP»P_jP_μC_IP[°]C_{II}C,P[°]PSPsP_j C- P»C-PeP[°]C_{II}C_IC_{II}PePëP_jPë P·P[°]C_IPsP±P[°]P_jPë,
P_jP_μC,P[°]P±PsP»C-P·P_j C_{II}PePëC... P·PrC-PN_ëC_IPSC_{II}C”C,C_{II}C_IC_{II}
PIPëC_%P_μP_iP_μC_{II}P_μP»C-C_‡P_μPSPëP_jPë C,,P_μC_{II}P_jP_μPSC,P[°]P_jPë PiC_{II}C_rP_iPë
C_‡PëC,PsC...C_{II}PsP_jC-PI P450, P_jPsP[¶]PSP[°] PSP_μ PsC_‡C-PeC_rPIP[°]C,Pë.

*P_μPsC,P_μPSC_‡C-PN_ëPSC- PIP·P[°]C”P_jPsPrC-C—, P_iPsPIBTM C_{II}P·P[°]PSC- P· PrC-C_{II}C_‡PsC_{II}
C_{II}P_μC_‡PsPIPëPSPsC_{II} P_iC_{II}P_μP_iP[°]C_{II}P[°]C,C_é P_YP_μPIC-PeP[°]C_{II} P[°]P_jP»PsPrPëPiC-PSPsP_j*

*P’P_iP»PëPI C-PSC€PëC... P»C-PeP[°]C_{II}C_IC_{II}PePëC... P·P[°]C_IPsP±C-PI PSP[°] PrC-C_{II}
P[°]P_jP»PsPrPëPiC-PSC_é*

P†PSPiC-P±C-C,PsCḡPë CYP3A4

PḡCḡPë PsPrPSPsC†P°C†PSPsPjC† P·P°C†C,PsC†C†PIP°PSPSC- P°PjP»PsPrPëPiC-PSC† P·
C†PëP»CHḡPSPëPjPë P°P±Ps PiPsPjC-CḡPSPëPjPë C-PSPiC-P±C-C,PsCḡP°PjPë CYP3A4
(C-PSPiC-P±C-C,PsCḡP°PjPë PiCḡPsC,PḡP°P·, P°P·PsP»CHḡPSPëPjPë
C,,C†PSPiC-C†PëPrP°PjPë, PjP°PeCḡPsP»C-PrP°PjPë C,PëPiC†
PḡCḡPëC,CḡPsPjC-C†PëPSC† P°P±Ps PeP»P°CḡPëC,CḡPsPjC-C†PëPSC†,
PIPḡCḡP°PiP°PjC-P»C† P°P±Ps PrPëP»C,C-P°P·PḡPjC†) PIPiP»PëPI P°PjP»PsPrPëPiC-PSC†
PjPsPḡPḡ P·PSP°C†PSPs PiPsC†PëP»CḡPIP°C,PëC†Cḡ, C%Ps C,P°PePsPḡ PjPsPḡPḡ
PiCḡPëP·PIPḡC†C,Pë PrPs PiC-PrPIPëC%PḡPSPSCḡ CḡPëP·PëPeC†
PIPëPSPëPePSPḡPSPSCḡ PiC-PiPsC,PḡPSP·C-C—. PḡP»C-PSC-C†PSC- PiCḡPsCḡPIPë
C,P°PePëC... C,,P°CḡPjP°PePsPeC-PSPḡC,PëC†PSPëC... P·PjC-PS PjPsPḡC†C,CHḡ P±C†C,Pë
P±C-P»CHḡC€ PIPëCḡP°PḡPḡPSPëPjPë C† PiP°C†C-C”PSC,C-PI P»C-C,PSCHḡPsPiPs
PIC-PeC†. P†C†PSC†C” PiC-PrPIPëC%PḡPSPëPḡ CḡPëP·PëPe PiC-PiPsC,PḡPSP·C-C—.
P PḡPePsPjPḡPSP†C†C”C,CHḡC†Cḡ CḡPḡC,PḡP»CHḡPSPḡ PsP±C†C,PḡPḡPḡPSPSCḡ
PiP°C†C-C”PSC,C-PI. PḡPsPḡPḡ P±C†C,Pë PiPsC,CḡC-P±PSPḡ PeP»C-PSC-C†PSPḡ
C†PiPsC†C,PḡCḡPḡPḡPSPSCḡ C- PePsCḡPḡPeC†C-Cḡ PrPsP·Pë.

P†PSP†C†PeC,PsCḡPë CYP3A4

PḡCḡPë PsPrPSPsC†P°C†PSPsPjC† P·P°C†C,PsC†C†PIP°PSPSC- PIC-PrPsPjPëC...
C-PSP†C†PeC,PsCḡC-PI CYP3A4 PePsPSC†PḡPSC,CḡP°C†C-Cḡ P°PjP»PsPrPëPiC-PSC† C†
PiP»P°P·PjC- PjPsPḡPḡ P·PjC-PSCḡPIP°C,PëC†Cḡ. PḡPsPjC† C†P»C-Pr
PePsPSC,CḡPsP»CḡPIP°C,Pë P°CḡC,PḡCḡC-P°P»CHḡPSPëPḡ C,PëC†Pe C,P°
CḡPḡPiC†P»CḡPIP°C,Pë PrPsP·C† CḡPe PiC-Pr C†P°C†, C,P°Pe C- PiC-C†P»Cḡ
C†C†PiC†C,PSCHḡPsC— C,PḡCḡP°PiC-C—, PsC†PsP±P»PëPIPs C† CḡP°P·C-
C†C†PiC†C,PSCHḡPsC— C,PḡCḡP°PiC-C— C-P· C†PëP»CHḡPSPëPjPë
C-PSP†C†PeC,PsCḡP°PjPë CYP3A4 (C,P°PePëPjPë CḡPe CḡPëC,,P°PjPiC-C†PëPS,
P·PIC-CḡPsP±C-Pḡ P·PIPëC†P°PḡPSPëPḡ).

PḡCḡPëPḡPsPj P°PjP»PsPrPëPiC-PSC† CḡP°P·PsPj P· PiCḡPḡPḡPiC,,CḡC†C,PsPj P°P±Ps
PiCḡPḡPḡPiC,,CḡC†C,PsPIPëPj C†PsPePsPj PSPḡ CḡPḡPePsPjPḡPSP†C†C”C,CHḡC†Cḡ,
PsC†PeC-P»CHḡPePë C† PrPḡCḡPePëC... PiP°C†C-C”PSC,C-PI PjPsPḡPḡ
PiC-PrPIPëC%C†PIP°C,PëC†Cḡ P±C-PsPrPsC†C,C†PiPSC-C†C,CHḡ PiCḡPḡPiP°CḡP°C,C†,
C%Ps PiCḡPsCḡPIP»CḡC”C,CHḡC†Cḡ PI PiPsC†PëP»PḡPSPSC- PḡPsPiPs
PiC-PiPsC,PḡPSP·PëPIPSPsC— PrC-C—.

P”P°PSC,CḡPsP»PḡPS (C-PSC,,C†P·C-Cḡ)

PḡC-Pr C†P°C† PrPsC†P»C-PrC-PI PSP° P»P°P±PsCḡP°C,PsCḡPSPëC...
C,PIP°CḡPëPSP°C... PiC-C†P»Cḡ PiCḡPëPḡPsPjC† PIPḡCḡP°PiP°PjC-P»C† C-
PIPSC†C,CḡC-C€PSCHḡPsPIPḡPSPSPsPiPs PIPḡPrPḡPSPSCḡ PrP°PSC,CḡPsP»PḡPSC†
C†PiPsC†C,PḡCḡC-PiP°P»PëC†Cḡ C,,C-P±CḡPëP»CḡC†C-Cḡ C€P»C†PSPsC†PeC-PI C-
C†PḡCḡC†PḡPIPs-C†C†PrPëPSPSPëPḡ PePsP»P°PiC† P· P»PḡC,P°P»CHḡPSPëPj
PSP°C†P»C-PrPePsPj C† P·Piḡḡ CḡP·PeC† P· CḡPsP·PIPëC,PePsPj
PiC-PiPḡCḡPeP°P»C-C”PjC-C—. P— PsPiP»CḡPrC† PSP° CḡPëP·PëPe CḡPsP·PIPëC,PeC†
PiC-PiPḡCḡPeP°P»C-C”PjC-C— C† PiP°C†C-C”PSC,C-PI, C†C...PëP»CHḡPSPëC... PrPs
PIPëPSPëPePSPḡPSPSCḡ P·P»PsCḡPeC-C†PSPsC— PiC-PiPḡCḡC,PḡCḡPjC-C—, P°

C,P°PePsP¶ PSP° C,P»C- C,PµCḡP°PīC-C— P·P»PsCḡPeC-CḡPSPsC—
PīC-PīPµCḡC,PµCḡPjC-C— CḡPµPePsPjPµPSPPrCḡC”C,CḡCḡCḡ CḡPSPēPeP°C,Pē
PsPrPSPsC‡P°CḡPSPsPiPs PīCḡPēPN₆PsPjCḡ P±P»PsPeP°C,PsCḡC-PI
PeP°P»CḡC†C-C”PIPēC... PeP°PSP°P»C-PI, C,P°PePēC... CḡPe P°PjP»PsPrPēPīC-PS.

*P’PīP»PēPI P°PjP»PsPrPēPīC-PSCḡ PSP° PrC-Cḡ C-PSCēPēC... P»C-PeP°CḡCḡCḡPePēC...
P·P°CḡPsP±C-PI*

P”C-PīPsC,PµPSP·PēPIPSPēPN₆ PµC,,PµPeC, P°PjP»PsPrPēPīC-PSCḡ PrPsPrP°C”C,CḡCḡCḡ
PrPs PiC-PīPsC,PµPSP·PēPIPSPsPiPs PµC,,PµPeC,Cḡ C-PSCēPēC... PīCḡPµPīP°CḡP°C,C-PI,
C%Ps P·PSPēP¶CḡCḡC,Cḡ P°CḡC,PµCḡC-P°P»CḡPSPēPN₆ C,PēCḡPe.

Pj C...PsPrC- PeP»C-PSC-C‡PSPēC... PrPsCḡP»C-PrP¶PµPSCḡ
P»C-PeP°CḡCḡCḡPePēC... PIP·P°C”PjPsPrC-PN₆ P°PjP»PsPrPēPīC-PS PSPµ
PIPīP»PēPIP°PI PSP° C,,P°CḡPjP°PePsPeC-PSPµC,PēPeCḡ P°C,PsCḡPIP°CḡC,P°C,PēPSCḡ,
PrPēPiPsPeCḡPēPSCḡ P°P±Ps PIP°CḡC,,P°CḡPēPSCḡ.

PŸPēPjPIP°CḡC,P°C,PēPS

PḡPrPSPsC‡P°CḡPSPēPN₆ PīCḡPēPN₆PsPj P±P°PiP°C,PsCḡP°P·PsPIPēC... PrPsP·
P°PjP»PsPrPēPīC-PSCḡ PīPs 10 PjPi C- CḡPēPjPIP°CḡC,P°C,PēPSCḡ PīPs 80 PjPi
PīCḡPēP·PIPPrPēPI PrPs P·P±C-P»CḡCēPµPSPSCḡ PIPīP»PēPICḡ
CḡPēPjPIP°CḡC,P°C,PēPSCḡ PSP° 77 % Cḡ PīPsCḡC-PIPSCḡPSPSC- C-P·
PīCḡPēPN₆PsPjPsPj P»PēCēPµ CḡPēPjPIP°CḡC,P°C,PēPSCḡ. P”PsP·P°
CḡPēPjPIP°CḡC,P°C,PēPSCḡ PrP»Cḡ PīP°C†C-C”PSC,C-PI, C%Ps PīCḡPēPN₆PjP°CḡC,Cḡ
P°PjP»PsPrPēPīC-PS, PSPµ PīPsPIPēPSPSP° PīPµCḡPµPIPēC%CḡPIP°C,Pē 20 PjPi PSP°
PrPsP±Cḡ.

PŷP°PeCḡPsP»C-PjCḡCḡ

P†CḡPSCḡC” CḡPēP·PēPe PīC-PrPIPēC%PµPSPSCḡ CḡC-PIPSC-PI
C,P°PeCḡPsP»C-PjCḡCḡCḡ PI PeCḡPsPIC- PīCḡPē PsPrPSPsC‡P°CḡPSPsPjCḡ
P·P°CḡC,PsCḡCḡPIP°PSPSC- P· P°PjP»PsPrPēPīC-PSPsPj. P©PsP± CḡPSPēPePSCḡC,Pē
C,PsPeCḡPēC‡PSPsCḡC,C- C,P°PeCḡPsP»C-PjCḡCḡCḡ PīCḡPē CḡCḡPīCḡC,PSCḡPsPjCḡ
P·P°CḡC,PsCḡCḡPIP°PSPSC- P°PjP»PsPrPēPīC-PSCḡ, PīPsC,CḡC-P±PµPS
CḡPµPīCḡP»CḡCḡPSPēPN₆ PjPsPSC-C,PsCḡPēPSPi CḡC-PIPSC-PI
C,P°PeCḡPsP»C-PjCḡCḡCḡ PI PeCḡPsPIC-, P° Cḡ CḡP°P·C- PSPµPsP±C...C-PrPSPsCḡC,C-
- PePsCḡPµPeC†C-Cḡ PrPsP·CḡPIP°PSPSCḡ.

P†PSPiC-P±C-C,PsCḡPē mTOR (PjC-CēPµPSC- CḡP°PīP°PjC-C†PēPSCḡ)

P†PSPiC-P±C-C,PsCḡPē mTOR, C,P°PeC- CḡPe CḡPēCḡPsP»C-PjCḡCḡ,
C,PµPjCḡPēCḡPsP»C-PjCḡCḡ C,P° PµPIPµCḡPsP»C-PjCḡCḡ, C” CḡCḡP±CḡC,CḡP°C,P°PjPē CYP3A.
PḡPjP»PsPrPēPīC-PS ḡḡ” CḡP»P°P±PePēPN₆ C-PSPiC-P±C-C,PsCḡ CYP3A. PµCḡPē
PsPrPSPsC‡P°CḡPSPsPjCḡ P·P°CḡC,PsCḡCḡPIP°PSPSC- C-PSPiC-P±C-C,PsCḡC-PI mTOR
P°PjP»PsPrPēPīC-PS PjPsP¶Pµ PīC-PrPIPēC%CḡPIP°C,Pē PIPīP»PēPI C-PSPiC-P±C-C,PsCḡC-PI
mTOR.

PḡPēPeP»PsCḡPīPsCḡPēPS

PμC-Pr C‡P°CÍ PíCĤPsCÍPíPμPeC, PëPIPSPsPiPs PeP»C-PSC-C‡PSPsPiPs
PrPsCÍP»C-PrP¶PμPSPSCĬ P·P° CíC‡P°CÍC, CĤ PíP°C†C-C" PSC, C-PI C-P·
C, CĤP°PSCÍPíP»P°PSC, P°C†C-C" CĤ PSPëCĤPePë CÍPíPsCÍC, PμCĤC-PiP°P»PsCÍCĤ
P·P±C-P»CĤC€PμPSPSCĬ PjC-PSC-PjP°P»CĤPSPsPiPs CĤC-PIPSCĬ
C†PëPeP»PsCÍPíPsCĤPëPSCÍ PI CÍPμCĤPμPrPSCĤPsPjCÍ PSP° 40 % PíCĤPë
PsPrPSPsC‡P°CÍPSPsPjCÍ P·P°CÍC, PsCÍCÍPIP°PSPSC- P№PsPiPs P·
P°PjP»PsPrPëPíC-PSPsPj. PŸCíPíCÍC, PSC" P·P°CÍC, PsCÍCÍPIP°PSPSCĬ
PíCĤPμPíP°CĤP°C, Cí PŸPμPIC-PeP°CĤ C, P° C†PëPeP»PsCÍPíPsCĤPëPSCÍ PjPsP¶Pμ
PíPsCÍPëP»PëC, Pë PíPíP»PëPI C†PëPeP»PsCÍPíPsCĤPëPSCÍ. PμCĤPë
CÍCíPíCíC, PSCĤPsPjCÍ P·P°CÍC, PsCÍCÍPIP°PSPSC- P· P°PjP»PsPrPëPíC-PSPsPj
CÍP»C-Pr CĤPsP·PiP»CĬPSCÍC, Pë PSPμPsP±C...C-PrPSC-CÍC, CĤ PePsPSC, CĤPsP»CĤ
P·P° PjC-PSC-PjP°P»CĤPSPëPj CĤC-PIPSPμPj C†PëPeP»PsCÍPíPsCĤPëPSCÍ PI
PeCĤPsPIC-, P° Cí CĤP°P·C- PíPsC, CĤPμP±Pë BĤ" P·PSPëP·PëC, Pë PrPsP·CÍ
C†PëPeP»PsCÍPíPsCĤPëPSCÍ.

PĤCÍPsP±P»PëPIPsCÍC, C- P·P°CÍC, PsCÍCÍPIP°PSPSCĬ.

PμP°C†C-C" PSC, Pë P· PiC-PíPsPIPsP»PμPjC-C" CĤ P°P±Ps PSPμCÍC, P°C‡PμCĤ PSP°C, CĤC-CĤ

Pj PíP°C†C-C" PSC, C-PI P· PiC-PíPsPIPsP»PμPjC-C" CĤ C, P°/P°P±Ps
PiC-PíPsPSP°C, CĤC-C" PjC-C" CĤ, C%Ps PíPëPSPëPeP»Pë PI CĤPμP·CíP»CĤC, P°C, C-
C-PSC, PμPSCÍPëPIPsPSC— C, PμCĤP°PíC-C— PrC-CíCĤPμC, PëPeP°PjPë, PsP±PjPμP¶PμPSPSCĬ
CÍPíPsP¶PëPIP°PSPSCĬ CÍPsP»C- P· C—P¶PμCĤ, PrC-P°CĤPμC— P°P±Ps P±P»CĤPIP°PSPSCĬ,
PjPsP¶Pμ PíPëPSPëPePSCÍC, Pë CÍPëPjPíC, PsPjP°C, PëC‡PSP° PiC-PíPsC, PμPSP·C-CĬ,
PsCÍPsP±P»PëPIPs PíC-CÍP»CĬ PíCĤPëPμPsPjCÍ PíPμCĤC€PsC— PrPsP·Pë.
P PμPePsPjPμPSPPrPsPIP°PSPs CíCÍCÍPSCÍC, Pë C†C- CÍC, P°PSPë PrPs PíPsC‡P°C, PeCí
P»C-PeCíPIP°PSPSCĬ PíCĤPμPíP°CĤP°C, PsPj PŸPμPIC-PeP°CĤ P°P±Ps PíCÍC, P°PSPsPíPëC, Pë
CĤPμC, PμP»CĤPSPμ CÍPíPsCÍC, PμCĤPμP¶PμPSPSCĬ P·P° CÍC, P°PSPsPj PíP°C†C-C" PSC, P° PSP°
PíPsC‡P°C, PeCí P»C-PeCíPIP°PSPSCĬ.

P†PSC€C- CÍC, P°PSPë, C%Ps CÍCÍPíCĤPsPIPsPrP¶C‡CĤC, CĤCÍCĬ CÍC, PëPjC‡P»CĬC†C-C" CĤ
CĤPμPSC-PS-P°PSPíC-PsC, PμPSP·PëPS-P°P»CĤPrPsCÍC, PμCĤPsPSPsPIPsC— CÍPëCÍC, PμPjPë

PμP°C†C-C" PSC, Pë, Cí CĬPePëC... CÍCíPrPëPSPSPëPμ C, PsPSCÍCÍ C- C, CíPSPeC†C-CĬ
PSPëCĤPsPe PíPμP»PëPePsCĤ PjC-CĤPsCĤ P·P°P»PμP¶P°C, CĤ PIC-Pr P°PeC, PëPIPsPSCÍC, C-
CĤPμPSC-PS-P°PSPíC-PsC, PμPSP·PëPS-P°P»CĤPrPsCÍC, PμCĤPsPSPsPIPsC— CÍPëCÍC, PμPjPë
(PSP°PíCĤPëPeP»P°Pr PíP°C†C-C" PSC, Pë P· C, CĬP¶PePsCĤ P·P°CÍC, C-PμPSPsCĤ
CÍPμCĤC†PμPIPsCĤ PSPμPrPsCÍC, P°C, PSC-CÍC, CĤ P°P±Ps P· C...PIPsCĤPsP±PsCĤ
PSPëCĤPsPe, PíPeP»CĤC‡P°CĤC‡Pë CÍC, PμPSPsP· PSPëCĤPePsPIPsC— P°CĤC, PμCĤC-C—),
PjPsP¶CíC, CĤ CĤPμP°PíCíPIP°C, Pë PSP° C-PSC€C- PíCĤPμPíP°CĤP°C, Pë, C%Ps
PíPíP»PëPIP°CĤC, CĤ PSP° C†CĤ CÍPëCÍC, PμPjCÍ (C, P°PeC- CĬPe P°PSC, P°PíPsPSC-CÍC, Pë
CĤPμC†PμPíC, PsCĤC-PI P°PSPíC-PsC, PμPSP·PëPSCÍ II), PíPsCÍC, CĤPsCĤ
PiC-PíPsC, PμPSP·C-C" CĤ, P°P·PsC, PμPjC-C" CĤ, PsP»C-PíCíCĤC-C" CĤ P°P±Ps CĤC-PrPePs -
PíPsCÍC, CĤPsCĤ PSPëCĤPePsPIPsCĤ PSPμPrPsCÍC, P°C, PSC-CÍC, CĤ.

P PμPSPsPIP°CÍPeC‡P»CĬCĤPSP° PiC-PíPμCĤC, PμPSP·C-CĬ

P—P°CÍC, PsCÍCÍPIP°PSPSCĬ PíCĤPμPíP°CĤP°C, C-PI, C%Ps PíPíP»PëPIP°CĤC, CĤ PSP°
CĤPμPSC-PS-P°PSPíC-PsC, PμPSP·PëPS-P°P»CĤPrPsCÍC, PμCĤPsPSPsPICÍ CÍPëCÍC, PμPjCÍ, Cí

C...PIPsCḥPëC... P· PrPIPsP±C-C‡PSPëPj C'c,PµPSPsP·PsPj PSPëCḥPePsPIPsC—
P°CḥC,PµCḥC-C— P°P±Ps C'c,PµPSPsP·PsPj P°CḥC,PµCḥC-C— C"PrPëPSPsC—
C,,C'PSPeC†C-PsPSCr'cḥC‡PsC— PSPëCḥPePë P'PsPIBḥ™ C'cP·P°PSPµ P· P'c-PrPIPëC%PµPSPëPj
CḥPëP·PëPePsPj C,C'P¶PePsC— P'c-P'PsC,PµPSP·C-C— C,P° PSPëCḥPePsPIPsC—
PSPµPrPsC'c,P°C,PSPsC'c,C-.

*PµPsCḥCáC€PµPSPSC'c C,,C'PSPeC†C-C— PSPëCḥPsPe C- C,CḥP°PSC'P'P»P°PSC,P°C†C-C'c
PSPëCḥPePë*

PµCḥPë P·P°C'c,PsC'cC'PIP°PSPSC- P'cḥPµP'P°CḥP°C,C' P'YµPIC-PeP°Cḥ
C...PIPsCḥPëPj C-P· P'PsCḥC'c€PµPSPSC'cPj C,,C'PSPeC†C-C— PSPëCḥPsPe
CḥPµPePsPjPµPSPr'c'c,CḥC'c'c' P'PµCḥC-PsPrPëC‡PSPëPñ PePsPSC,CḥPsP»Cḥ
CḥC-PIPsC'c PeP°P»C-Cḥ C,P° PeCḥPµP°C,PëPSC-PSC'c P' C'PëCḥPsPIP°C,C†C-
PeCḥPsPIC-. P—P°C'c,PsC'cC'PIP°PSPSC'c P'cḥPµP'P°CḥP°C,C' P'YµPIC-PeP°Cḥ PSPµ
CḥPµPePsPjPµPSPr'PsPIP°PSPs P'P°C†C-C"psc,P°Pj C-P· C,C'P¶PePëPj
P'PsCḥC'c€PµPSPSC'cPj C,,C'PSPeC†C-C— PSPëCḥPsPe (PeP»C-CḥPµPSC'c
PeCḥPµP°C,PëPSC-PSC'c < 20 PjP»/C...PI). P"PsC'cPIC-Pr P·P°C'c,PsC'cC'PIP°PSPSC'c
P'cḥPµP'P°CḥP°C,C' P'YµPIC-PeP°Cḥ P'P°C†C-C"psc,P°Pj, C'cPeC- PSPµC%PsPrP°PIPsP
P'PµCḥPµPSPµC'P»Pë C,CḥP°PSC'P'P»P°PSC,P°C†C-Cḥ PSPëCḥPePë, P°P±Ps
P'P°C†C-C"psc,P°Pj P· C,PµCḥPjC-PSP°P»CḥPSPsCḥ PSPëCḥPePsPIPsCḥ
PSPµPrPsC'c,P°C,PSC-C'c,Cḥ (PSP°P'cḥPëPeP»P°Pr, PeP»C-CḥPµPSC'c
PeCḥPµP°C,PëPSC-PSC'c < 12 PjP»/C...PI) PIC-PrC'cC'c,PSC-Pñ (PrPëPI. CḥPsP·PrC-P»Pë
B«P'Y'P'PsC'c-P± P·P°C'c,PsC'cC'PIP°PSPSC'cB», B«PαP°CḥPjP°PePsPeC-PSPµC,PëPeP°B»).

*PµPsPrPIC-PñPSP° P±P»PsPeP°PrP° CḥPµPSC-PS-P°PSPiC-PsC,PµPSP·PëPS-
P°P»CḥPrPsC'c,PµCḥPsPSPsPIPsC— C'PëC'c,PµPjPë (P PḥPḥP'Y)*

P,, C'PIC-PrC‡PµPSPSC'c, C%Ps C'P'c-P»CḥPSPµ P·P°C'c,PsC'cC'PIP°PSPSC'c
C-PSPiC-P±C-C,PsCḥC-PI PḥPµPα C,P° P±P»PsPeP°C,PsCḥC-PI CḥPµC†PµP'c,PsCḥC-PI
P°PSPiC-PsC,PµPSP·PëPSC'c II P°P±Ps P°P»C-C'PeC-CḥPµPSC'c P·P±C-P»CḥC€C'c" CḥPëP·PëPe
CḥPsP·PIPëC,PeC' P°CḥC,PµCḥC-P°P»CḥPSPsC— P'c-P'PsC,PµPSP·C-C—,
P'c-P'PµCḥPeP°P»C-C"pjc-C— C- P·PSPëP¶PµPSPSC'c C,,C'PSPeC†C-C— PSPëCḥPsPe (PI
C,PsPjC' C‡PëC'P»C- P'PsC'c,CḥPsC— PSPëCḥPePsPIPsC— PSPµPrPsC'c,P°C,PSPsC'c,C-).
P'YPsPjC' P'PsPrPIC-PñPSP° P±P»PsPeP°PrP° P PḥPḥP'Y PSP° C,P»C- C'P'c-P»CḥPSPsPiPs
P·P°C'c,PsC'cC'PIP°PSPSC'c C-PSPiC-P±C-C,PsCḥC-PI PḥPµPα C,P° P±P»PsPeP°C,PsCḥC-PI
CḥPµC†PµP'c,PsCḥC-PI P°PSPiC-PsC,PµPSP·PëPSC'c II P°P±Ps P°P»C-C'PeC-CḥPµPSC'c PSPµ
CḥPµPePsPjPµPSPr'c'c,CḥC'c'c' (PrPëPI. CḥPsP·PrC-P» B«P'P·P°C"pjsPrC-C'c P·
C-PSC€PëPjPë P»C-PeP°CḥC'cḥPePëPjPë P·P°C'cPsP±P°PjPë C,P° C-PSC€C- PIPëPrPë
PIP·P°C"pjsPrC-PñB»).

P'PeC%Ps C,PµCḥP°P'c-C'c P· P'PsPrPIC-PñPSPsCḥ P±P»PsPeP°PrPsCḥ C"
P°P±C'cPsP»CḥC,PSPs PSPµPsP±C...C-PrPSPsCḥ, C,Ps PIPsPSP° P'PsPIPëPSPSP°
P'cḥPsPIPsPrPëC,PëC'c'c' C,C-P»CḥPePë P'c-Pr PSP°P'P»C'cPrPsPj C,,P°C...C-PIC†C'c, P°
C,P°PePsP¶ PSP° C,P»C- CḥPµC,PµP»CḥPSPsPiPs C'P'PsC'c,PµCḥPµP¶PµPSPSC'c P·P°
C,,C'PSPeC†C-C"ḥ PSPëCḥPsPe, CḥC-PIPsPµPj PµP»PµPeC,CḥPsP»C-C,C-PI C-
P°CḥC,PµCḥC-P°P»CḥPSPsPiPs C,PëC'c'PeC'.

P'Y'P'c-P»CḥPSPµ P·P°C'c,PsC'cC'PIP°PSPSC'c C-PSPiC-P±C-C,PsCḥC-PI PḥPµPα C,P°
P±P»PsPeP°C,PsCḥC-PI CḥPµC†PµP'c,PsCḥC-PI P°PSPiC-PsC,PµPSP·PëPSC'c II
P'cḥPsC,PëP'PsPeP°P·P°PSPs P'P°C†C-C"psc,P°Pj P· PrC-P°P±PµC,PëC‡PSPsCḥ
PSPµC,,CḥPsP'P°C,C-C"ḥ.

PμPsCḡCáCεPμPSPSCḡ C,,CáPSPeC†C-C— PiPμC†C-PSPePë

PJ PiP°C†C-C" PSC,C-PI P· PiPsCḡC†CεPμPSPSCḡPj C,,C†PSPeC†C-C— PiPμC†C-PSPePë
P·P±C-P»CḡCεC†C"C,CḡC†CḡCḡ PμPeC†P†PsP·PëC†C-Cḡ P°PjP»PsPrPëPiC-PSC† C,P°
PsP»PjPμC†P°CḡC,P°PSC† PjPμPrPsPeC†PsPjC-P»C† (PrPëPI. CḡPsP·PrC-P»
B«P°P°CḡPjP°PePsPeC-PSPμC,PëPeP°B»). PŸPμPIC-PeP°Cḡ C†P»C-Pr P·
PsP±PμCḡPμP¶PSC-C†C,Cḡ PiCḡPëP·PSP°C†P°C,Pë PiP°C†C-C" PSC,P°Pj P·
PiPsCḡC†CεPμPSPSCḡPj C,,C†PSPeC†C-C— PiPμC†C-PSPePë P»PμPiPePsPiPs P°P±Ps
PiPsPjC-CḡPSPsPiPs C†C,C†PiPμPSCḡ. P" P»Cḡ PiP°C†C-C" PSC,C-PI P· PiPsPjC-CḡPSPëPj
PiPsCḡC†CεPμPSPSCḡPj C,,C†PSPeC†C-C— PiPμC†C-PSPePë PrPsP·P° PsP»PjPμC†P°CḡC,P°PSC†
PjPμPrPsPeC†PsPjC-P»C† PSPμ PiPsPiPëPSPSP° PiPμCḡPμPiPëC%°C†PiP°C,Pë 20 PjPi (PrPëPI.
CḡPsP·PrC-P» B«PŸPiPsC†C-P± P·P°C†C,PsC†C†PiP°PSPSCḡ C,P° PrPsP·PëB»).
PμP°C†C-C" PSC,P°Pj P· PiPsCḡC†CεPμPSPSCḡPj C,,C†PSPeC†C-C— PiPμC†C-PSPePë
PiCḡPëPμPsPj P°PjP»PsPrPëPiC-PSC† C†P»C-Pr PiPsC†PëPSP°C,Pë P· PSP°PμPSPëP¶C†PsC—
PrPsP·Pë C- PrPsC,CḡPëPjC†PiP°C,PëC†Cḡ PsP±PμCḡPμP¶PSPsC†C,C- CḡPe PSP°
PiPsC†P°C,PeC† P»C-PeC†PiP°PSPSCḡ, C,P°Pe C- PiCḡPë PiC-PrPiPëC%°PμPSPSC- PrPsP·Pë.
PμCḡPμPiP°CḡP°C, PŸPμPIC-PeP°Cḡ PiCḡPsC,PëPiPsPeP°P·P°PSPëPμ PiP°C†C-C" PSC,P°Pj P·
C,CḡP¶PePëPj PiPsCḡC†CεPμPSPSCḡPj C,,C†PSPeC†C-C— PiPμC†C-PSPePë (PrPëPI.
CḡPsP·PrC-P» B«PμCḡPsC,PëPiPsPeP°P·P°PSPSCḡB»).

P"C-PiPμCḡPeP°P»C-C"PjC-Cḡ

PİPe C- PiCḡPë P·P°C†C,PsC†C†PiP°PSPSC- C-PSCεPëC... P°PSC,P°PiPsPSC-C†C,C-PI
P°PSPiC-PsC,PμPSP·PëPSC† II C,P° C-PSPiC-P±C-C,PsCḡC-PI PḡPμPα, PiCḡPë
P»C-PeC†PiP°PSPSC- PiCḡPμPiP°CḡP°C,PsPj PŸPμPIC-PeP°Cḡ PjPsP¶Pμ PiPëPSPëPePSC†C,Pë
PiC-PiPμCḡPeP°P»C-C"PjC-Cḡ, PsC†PsP±P»PëPiPs PiCḡPë PiPsCḡC†CεPμPSPSC-
C,,C†PSPeC†C-C— PSPëCḡPsPe C,P°/P°P±Ps C†PμCḡC†PμPIC-Pμ PSPμPrPsC†C,P°C,PSPsC†C,C-
(PrPëPI. CḡPsP·PrC-P» B«P'P·P°C"PjPsPrC-Cḡ P· C-PSCεPëPjPë P»C-PeP°CḡC†CḡPePëPjPë
P·P°C†PsP±P°PjPë C,P° C-PSCεC- PiPëPrPë PiP·P°C"PjPsPrC-PμB»). PμP°C†C-C" PSC,P°Pj P·
C†C-C"C— PiCḡC†PiPë CḡPëP·PëPeC† CḡPμPePsPjPμPSP†C†C,CḡC†Cḡ C†P°C†C,PëPμ
PePsPSC,CḡPsP»Cḡ CḡC-PIPSCḡ C†PëCḡPsPiP°C,PePsPiPsPiPs PeP°P»C-Cḡ.

PμCḡPμPiP°CḡP°C, P· PsP±PμCḡPμP¶PSC-C†C,Cḡ PiCḡPëP·PSP°C†P°CḡC,Cḡ C†
PiPsC"PrPSP°PSPSC- P· PeP°P»C-C"PiPëPjPë PrPsP±P°PiPëP°PjPë,
PeP°P»C-PμP·P±PμCḡC-PiP°CḡC†PëPjPë PrC-C†CḡPμC,PëPeP°PjPë, P·P°PjC-PSPSPëPeP°PjPë
C†PsP»C-, C%°Ps PjC-C†C,CḡC,ḡ PeP°P»C-Pμ, P°P±Ps C-PSCεPëPjPë
P»C-PeP°CḡC†CḡPePëPjPë P·P°C†PsP±P°PjPë, CḡPeC- PjPsP¶C†C,Cḡ PiC-PrPiPëC%°PëC,Pë
CḡC-PIPμPSCḡ PeP°P»C-Cḡ (PiPμPiP°CḡPëPS C,PsC%°Ps); PiCḡPë C†CḡPsPjC†
CḡPμPePsPjPμPSP†C†C,CḡC†Cḡ CḡPμPiC†P»CḡCḡPSPëPμ PePsPSC,CḡPsP»Cḡ CḡC-PIPSCḡ
PeP°P»C-Cḡ PI PeCḡPsPIC-.

PμCḡPμPiP°CḡP°C,Pë P»C-C,C-Cḡ

PİPe C- PiCḡPë P·P°C†C,PsC†C†PiP°PSPSC- C-PSCεPëC... P°PSC,P°PiPsPSC-C†C,C-PI
P°PSPiC-PsC,PμPSP·PëPSC† II, PsPrPSPsC†P°C†PSPμ P·P°C†C,PsC†C†PiP°PSPSCḡ
PiCḡPμPiP°CḡP°C,C† PŸPμPIC-PeP°Cḡ C- PiCḡPμPiP°CḡP°C,C-PI P»C-C,C-Cḡ PSPμ
CḡPμPePsPjPμPSP†C†C,CḡC†Cḡ (PrPëPI. CḡPsP·PrC-P» B«P'P·P°C"PjPsPrC-Cḡ P·
C-PSCεPëPjPë P»C-PeP°CḡC†CḡPePëPjPë P·P°C†PsP±P°PjPë C,P° C-PSCεC- PiPëPrPë
PiP·P°C"PjPsPrC-PμB»).

*PŸC,PμPSPsP· C†C†C,Cḡ P°PsCḡC,Pë P°P±Ps PjC-C,CḡP°P»CḡPSPsPiPs PeP»P°PiP°PSP°;
PsP±C†C,CḡCáPeC,PëPiPSP° PiC-PiPμCḡC,CḡPsC,,C-C†PSP° PeP°CḡP†C-PsPjC-PsPiP°C,C-Cḡ*

PJ P·PIBTM CII·PeCf P· PSP°CIIPIPSC-CfC,Ch PI CfPeP»P°PrC- PiCTMPiP°CTMP°C,Cf PTMPiC-PeP°CTM P°PjP»PsPrPëPiC-PSCf, CII·Pe C- C°PsPrPs CfCfC-C... CfCfPrPëPSPsCTMPsP·CëPëCTMChPIP°P»CHPSPëC... P·P°CfPsP±C-PI, CTMPiPePsPjPTMPSPPrCfC°C,ChCfCII PrPsC,CTMPëPjCfPIP°C,PëCfCII PsCfPsP±P»PëPIPsC— PsP±PTMCTMPiPTMPSPScfC,C- PiCTMPë PiCTMPëP·PSP°C±PTMPSPSC- PiP°CfC-C°PSC,P°Pj P·C- CfC,PTMPSPsP·PsPj CfCfC,CII P°PsCTMC,Pë P°P±Ps PjC-C,CTMP°P»CHPSPsPiPs PeP»P°PiP°PSP° P°P±Ps PsP±CfC,CTMCfPeC,PëPIPSPsCh PiC-PiPTMCTMC,CTMPsC,,C-C±PSPsCh PeP°CTMPrC-PsPjC-PsPiP°C,C-C°Ch.

PTMPiCTMPiP°PSPSPëPTM P°P»CHPrPsCfC,PTMCTMPsPSC-P·Pj

PTMP°CfC-C°PSC,Pë P· PiPTMCTMPiP°PSPSPëPj P°P»CHPrPsCfC,PTMCTMPsPSC-P·PjPsPj P·P°P·PIPëC±P°PTM PSPTM CTMPiP°PiCfCTMC,Ch PSP° PiC-PiPsC,PTMPSP·PëPIPSC- PiCTMPiP°CTMP°C,Pë, C°Ps PiCTMPëPiPSC-C±CfCTMC,Ch CTMPiPSC-PS- P°PSPiC-PsC,PTMPSP·PëPSPsPICf CfPëCfC,PTMPjCf. PTMPsPjCf P·P°CfC,PsCfCfPIP°PSPSCII PTMPiC-PeP°CTMCf PSPTM CTMPiPePsPjPTMPSPPrPsIP°PSPs C,P°PePëPj PiP°CfC-C°PSC,P°Pj.

PTMPiCTMCfPTMPIP° PSPTMPrPsCfC,P°C,PSC-CfC,Ch

PJ P·PIBTM CII·PeCf P· PiCTMPëPiPSC-C±PTMPSPSCII Pj P°PSPiC-PsC,PTMPSP·PëPS- P°P»CHPrPsCfC,PTMCTMPsPSPsPIPsC— CfPëCfC,PTMPjPë PjPsPTMP»PëPIPTM PiPsCTMCfCëPTMPSPSCII C,,CfPSPeCfC-C— PSPëCTMPsPe Cf CfC...PëP»CHPSPëC... PrPs CfCHPsPiPs PiP°CfC-C°PSC,C-PI. Pj PiP°CfC-C°PSC,C-PI P· C,CII·PePsCTM CfPTMCTMCfPTMPIPsCTM PSPTMPrPsCfC,P°C,PSC-CfC,Ch, Cf CII·PePëC... PjPsPTMP»PëPIP° P·P°P»PTMPTMPSC-CfC,Ch PSPëCTMPePsPIPsC— C,,CfPSPeCfC-C— PIC-Pr P°PeC,PëPIPSPsCfC,C- CTMPiPSC-PS-P°PSPiC-PsC,PTMPSP·PëPS- P°P»CHPrPsCfC,PTMCTMPsPSPsPIPsC— CfPëCfC,PTMPjPë, P»C-PeCfPIP°PSPSCII C-PSPiC-P±C-C,PsCTMP°PjPë P°PSPiC-PsC,PTMPSP·PëPSPiPTMCTMPiC,PiPsCTMChPIP°P»CHPSPsPiPs C,,PTMCTMPjPTMPSC,Cf (PTMPTMα) C- P°PSC,P°PiPsPSC-CfC,P°PjPë CTMPiCfPTMPiC,PsCTMC-PI P°PSPiC-PsC,PTMPSP·PëPSCf PjPsPTMPTM CfCfPiCTMPsPIPsPrPTMCfPIP°C,PëCfCII PsP»C-PiCfCTMC-C°Ch C-/P°P±Ps PiCTMPsPiCTMPiCfCfC±PsCTM P°P·PsC,PTMPjC-C°Ch C- (CTMC-PrPePs) PiPsCfC,CTMPsCTM PSPëCTMPePsPIPsCTM PSPTMPrPsCfC,P°C,PSC-CfC,Ch C,P°/P°P±Ps CfPjPTMCTMC,Ch.

PTMP°CfC-C°PSC,P°Pj C-P· CfPTMCTMCfPTMPIPsCTM PSPTMPrPsCfC,P°C,PSC-CfC,Ch C,PTMCTMP°PiC-Ch CfP»C-Pr PiCTMPsPIPsPrPëC,Pë P· PsP±PTMCTMPiPTMPSC-CfC,Ch. Pj PrPsPIPsCfC,CTMPsPePsPIPsPjCf PiP»P°CfPTMP±Ps-PePsPSC,CTMPsP»CHPsPIP°PSPsPjCf PrPsCfP»C-PrPTMPTMPSPSC- P°PjP»PsPrPëPiC-PSCf Cf PiP°CfC-C°PSC,C-PI P· C,CII·PePsCTM CfPTMCTMCfPTMPIPsCTM PSPTMPrPsCfC,P°C,PSC-CfC,Ch (NYHA III C- IV) PeC-P»CHPeC-CfC,Ch PiPsPIC-PrPsPjP»PTMPSCTM PiCTMPs CTMPsP·PIPëC,PsPe PSP°P±CTMCII·PeCf P»PTMPiPTMPSC-PI Cf PiCTMCfPiC- P°PjP»PsPrPëPiC-PSCf P±CfP»P° P±C-P»CHCëPsCTM PI PiPsCTMC-PIPSCII·PSPSC- P· PiCTMCfPiPsCTM PiP»P°CfPTMP±Ps (PrPëPI. CTMPsP·PrC-P» B«PαP°CTMPjP°PePsPrPëPSP°PjC-PeP°B»). P°P»PsPeP°C,PsCTMPë PeP°P»CHCfC-C°PIPëC... PeP°PSP°P»C-PI, PiPeP»CHC±P°CTMC±Pë P°PjP»PsPrPëPiC-PS, CfP»C-Pr P· PsP±PTMCTMPiPTMPSC-CfC,Ch P·P°CfC,PsCfPsPICfPIP°C,Pë PiP°CfC-C°PSC,P°Pj C-P· P·P°CfC,C-PN°PSPsCTM CfPTMCTMCfPTMPIPsCTM PSPTMPrPsCfC,P°C,PSC-CfC,Ch, PsCfPeC-P»CHPePë CfC- PiCTMPiP°CTMP°C,Pë PjPsPTMCfC,Ch PiC-PrPIPëC°CfPIP°C,Pë CTMPëP·PëPe PiPëPSPëPePSPTMPSPSCII CII·PIPëC° P· P±PsPeCf CfPTMCTMCfPTMPIPs-CfCfPrPëPSPSPsC— CfPëCfC,PTMPjPë PI PjP°PN°P±CfC,PSCHPsPjCf C- CfPjPTMCTMC,PSC-CfC,Ch.

PTMPiCTMCfPTMPIP° PTMPSC,PTMCTMPsPiP°C,C-CII

PJ PrCfPTMPTM CTMC-PrPeC-CfPSPëC... PiPëPiP°PrPeP°C... PiPsPIC-PrPsPjP»CII·P»Pë PiCTMPs

C, CИP¶PeCí C... CЪPsPSC-C#PSCí PrC-P° CЪPµCЪ P·C- P·PSP° C#PSPsCЪ
PIC, CЪP° C, PsCЪ PjP° CГPë C, C-P»P°, C%Ps CЪPsP·PIPëP»P° CГCЦ C#PµCЪPµP·
PeC-P»CЪPeP° PjC-CГCЦC†C-PI P°P±Ps CЪPsPeC-PI PíC-CГP»CЦ PíPsC#P° C, PeCí
P»C-PeCíPIP°PSPSCЦ Cí PíP° C†C-C" PSC, C-PI, CИPeC- PíCЪPëP№PjP°P»Pë
PsP»PjPµCГP° CЪC, P°PS; PíCЪPëC#PëPSPsCЪ C—C— PIPëPSPëPePSPµPSPSCЦ C",
C-PjPsPIC-CЪPSPs, PjC-CГC†PµPIP° PIC-PrCГC, CЪPsC#PµPSP° CЪPµP°PeC†C-CЦ
PíC-PíPµCЪC#CíC, P»PëPIPsCГC, C-. P PµP·CíP»CЪC, P° C, Pë P±C-PsPíCГC-C—
CГP»PëP·PsPIPsC— PsP±PsP»PsPSPePë PePëC€PµC#PSPëPeCí Cí C, P°PePëC...
PíP° C†C-C" PSC, C-PI C#P° CГC, Ps PrPµPjPsPSCГC, CЪCíPIP°P»Pë P° C, CЪPsC,, C-CЪ
PePëC€PePsPIPëC... PIPsCЪCГPëPSPsPe. PíPeC%Ps C, P°PeC- CГPëPjPíC, PsPjPë
PIPëPSPëPeP° CЪC, CЪ Cí PíP° C†C-C" PSC, P° PíC-Pr C#P° CГ P»C-PeCíPIP°PSPSCЦ
PsP»PjPµCГP° CЪC, P°PSPsPj C, P° P·P° PIC-PrCГCíC, PSPsCГC, C- C-PSC€PëC...
CИPIPSPëC... PíCЪPëC#PëPS, CГP»C-Pr PSPµPíP°P№PSPs PIC-PrPjC-PSPëC, Pë
P·P° CГC, PsCГCíPIP°PSPSCЦ PsP»PjPµCГP° CЪC, P°PSCí C, P° PSPµ
CЪPsP·PíPsC#PëPSP° C, Pë P№PsPíPs PíCЪPëP№PsPj P·PSPsPICí. PíPeC%Ps
PrC-P° CЪPµCЦ PSPµ PjPëPSP° C" PíCЪPsC, CИPiPsPj C, PëP¶PSCЦ PíC-CГP»CЦ
PIC-PrPjC-PSPë PíCЪPµPíP° CЪP° C, Cí, CГP»C-Pr CЪPsP·PíP»CИPSCíC, Pë
PSPµPsP±C... C-PrPSC-CГC, CЪ PePsPSCГCíP»CЪC, P° C†C-C— Cí
PIC-PrPíPsPIC-PrPSPsPíPs CГPíPµC†C-P°P»C-CГC, P° (PSP° PíCЪPëPeP»P° Pr
PíP° CГC, CЪPsPµPSC, PµCЪPsP»PsPíP°).

PЪPSPiC-PsPSPµPICЪPsC, PëC#PSPëP№ PSP°P±CЪCИPe PePëC€PµC#PSPëPeCé

PµPsPIC-PrPsPjP»CИP»PsCГCЦ PíCЪPs PIPëPíP° PrPePë
P°PSPiC-PsPSPµPICЪPsC, PëC#PSPsPíPs PSP°P±CЪCИPeCí PePëC€PµC#PSPëPeCí Cí
PíP° C†C-C" PSC, C-PI, CИPeC- P·P° CГC, PsCГCíPIP°P»Pë P±P»PsPeP° C, PsCЪPë
CЪPµC†PµPíC, PsCЪC-PI P°PSPiC-PsC, PµPSP·PëPSCí II, PIPeP»CЪC#P° CЪC#Pë
PsP»PjPµCГP° CЪC, P°PS (PrPëPI. CЪPsP·PrC-P» B«PµPsP±C-C#PSC- CЪPµP°PeC†C-C—B»). Pj
C†PëC... PíP° C†C-C" PSC, C-PI CГPíPsCГC, PµCЪC-PíP°P»PëCГCЦ P±C-P»CЪ Cí P¶PëPIPsC, C-,
PSCíPrPsC, P°, P±P»CЪPIP°PSPSCЦ C, P° PrC-P° CЪPµCЦ. PŸPëPjPíC, PsPjPë PjPëPSP°P»Pë
PíC-CГP»CЦ PIC-PrPjC-PSPë P±P»PsPeP° C, PsCЪC-PI CЪPµC†PµPíC, PsCЪC-PI
P°PSPiC-PsC, PµPSP·PëPSCí P†P†. PíPeC%Ps PrC-P° PíPSPsCГC, PsPIP°PSPs
P°PSPiC-PsPSPµPICЪPsC, PëC#PSPëP№ PSP°P±CЪCИPe PePëC€PµC#PSPëPeCí, CГP»C-Pr
PíCЪPëPíPëPSPëC, Pë P·P° CГC, PsCГCíPIP°PSPSCЦ PsP»PjPµCГP° CЪC, P°PSCí C, P°
CЪPsP·PíPsC#P° C, Pë PIC-PrPíPsPIC-PrPSPëP№ PjPsPSC-C, PsCЪPëPSPi PrPs
PíPsPIPSsPíPs P·PSPëPePSPµPSPSCЦ CГPëPjPíC, PsPjC-PI.

P·C, PSC-C#PSC- PIC-PrPjC-PSPSPsCГC, C-

PíPe C- Cí CЪP°P·C- P·P° CГC, PsCГCíPIP°PSPSCЦ C-PSC€PëC... P°PSC, P°PíPsPSC-CГC, C-PI
CЪPµC†PµPíC, PsCЪC-PI P°PSPiC-PsC, PµPSP·PëPSCí II, PíC-PíPsC, PµPSP·PëPIPSPëP№
PµC,, PµPeC, PíCЪPµPíP° CЪP° C, Cí PŸPµPIC-PeP° CЪ Cí PíCЪPµPrCГC, P°PIPSPëPeC-PI
PSPµPíCЪPsC—PrPSPsC— CЪP° CГPë PjPsP¶Pµ P±CíC, Pë PrPµC%Ps PjPµPSC€PëPj, PSC-P¶ Cí
C-PSC€PëC... PíP° C†C-C" PSC, C-PI, PjPsP¶P»PëPIPs, PIPSP° CГP»C-PrPsPe P±C-P»CЪC€PsC—
PíPsC€PëCЪPµPSPsCГC, C- PSPëP·CЪPePsPíPs CЪC-PIPSЦЦ CЪPµPSC-PSCí PI C†C-P№
PíPsPíCíP»CЦC†C-C—.

PµP° C†C-C" PSC, Pë P»C-C, PSCHPsPíPs PIC-PeCé

РµС-РrPIPëC%oCíPIP°С, Pë PrPsP·Cí PíCĤPµPíP°CĤP°С, Cí PíCĤPë P»C-PeCíPIP°PSPSC-
PíP°C†C-C" PSC, C-PI P»C-C, PSCĤPsPiPs PIC-PeCí CÍP»C-Pr P· PsP±PµCĤPµP¶PSC-CÍС, CĤ
(PrPëPI. CĤPsP·PrC-P» B«P»P°CĤPjP°PePsPeC-PSPµC, PëPeP°B»).

P°PíC-C, PSC-CÍС, CĤ

PĤPSC, P°PiPsPSC-CÍС, Pë CĤPµC†PµPíC, PsCĤC-PI P°PSPiC-PsC, PµPSP·PëPSCí P†P†
PíCĤPsC, PëPíPsPeP°P·P°PSC- PíC-Pr C†P°CÍ PIP°PíC-C, PSPsCÍС, C-.

PíPeC%oPs PíCĤPsPrPsPIP¶PµPSPSC¶ PíC-PíPsC, PµPSP·PëPIPSPsC— C, PµCĤP°PíC-C—
PIPIP°P¶P°С, CĤCÍС¶ PSPµPsP±C...C-PrPSPëPj, Cí PíP°C†C-C" PSC, PsPe, C¶PeC-
PíP»P°PSCíCĤC, CĤ PIP°PíC-C, PSC-CÍС, CĤ, CÍP»C-Pr P·PjC-PSPëC, Pë
P°PSC, P°PiPsPSC-CÍС, Pë CĤPµC†PµPíC, PsCĤC-PI P°PSPiC-PsC, PµPSP·PëPSCí P†P† PSP°
P°P»CĤC, PµCĤPSP°С, PëPIPSCí P°PSC, PëPíC-PíPµCĤC, PµPSP·PëPIPSCí
C, PµCĤP°PíC-CĤ, C¶PeP° PjP°С" PrPsPIPµPrPµPSPëPµ PíCĤPsC,, C-P»CĤ
P±PµP·PíPµPePë PrP»C¶ P·P°CÍС, PsCÍСíPIP°PSPSC¶ PIP°PíC-C, PSPëPjPë. PíPeC%oPs
PIP°PíC-C, PSC-CÍС, CĤ PrC-P°PiPSPsCÍС, PsPIP°PSPs, P»C-PeCíPIP°PSPSC¶
P°PSC, P°PiPsPSC-CÍС, P°PjPë CĤPµC†PµPíC, PsCĤC-PI P°PSPiC-PsC, PµPSP·PëPSCí P†P†
CÍP»C-Pr PSPµPíP°PµPSPs PíCĤPëPíPëPSPëC, Pë C- PI CĤP°P·C- PíPsC, CĤPµP±Pë
CĤPsP·PíPsC†P°С, Pë P°P»CĤC, PµCĤPSP°С, PëPIPSCí C, PµCĤP°PíC-CĤ (PrPëPI.
CĤPsP·PrC-P»Pë B«PµCĤPsC, PëPíPsPeP°P·P°PSPSC¶B», B«P—P°CÍС, PsCÍСíPIP°PSPSC¶
Cí PíPµCĤC-PsPr PIP°PíC-C, PSPsCÍС, C- P°P±Ps PiPsPrCíPIP°PSPSC¶
PíCĤCíPrPrCĤB»).

P†PSC€Pµ

PíPe C- PíCĤPë P·P°CÍС, PsCÍСíPIP°PSPSC- P±CíPrCĤ-C¶PePëC...
PíC-PíPsC, PµPSP·PëPIPSPëC... P·P°CÍСPsP±C-PI, PSP°PrPjC-CĤPSPµ P·PSPëP¶PµPSPSC¶
P°CĤC, PµCĤC-P°P»CĤPSPsPiPs C, PëCíPeCí Cí PíP°C†C-C" PSC, C-PI P· C-€PµPjC-C†PSPsCĤ
C...PIPSCĤPsP±PsCĤ CÍPµCĤC†C¶ P°P±Ps P· PíPsCĤCíC€PµPSPSC¶Pj PjPsP·PePsPiPsPiPs
PeCĤPsPiPsPsP±C-PíCí PjPsP¶Pµ PíCĤPëP·PIPµCÍС, Pë PrPs C-PSC,, P°CĤPeC, Cí
PjC-PsPeP°CĤPrP° P°P±Ps C-PSCíCíP»CĤC, Cí.

PµCĤPµPíP°CĤP°С, PŸPµPIC-PeP°CĤ PjC-CÍС, PëC, CĤ PjPµPSC€Pµ 1 PjPjPsP»CĤ PSP°С, CĤC-CĤ
(23 PjPi) PSP° PsPrPSCí C, P°P±P»PµC, PeCí, PíPeCĤPëC, Cí PíP»C-PIPePsPiPsCĤ
PsP±PsP»PsPSPePsCĤ, PsC, P¶Pµ PIPIP°P¶P°С, CĤCÍС¶ PíCĤP°PeC, PëC†PSPs
PIC-P»CĤPSPëPj PIC-Pr PSP°С, CĤC-CĤ.

*P—P°CÍС, PsCÍСíPIP°PSPSC¶ C¿ PíPµCĤC-PsPr PIP°PíC-C, PSPsCÍС, C- P°P±Ps
PiPsPrC¿PIP°PSPSC¶ PíCĤC¿PrPrCĤ.*

PµCĤPµPíP°CĤP°С, PŸPµPIC-PeP°CĤ PíCĤPsC, PëPíPsPeP°P·P°PSPëPµ PIP°PíC-C, PSPëPj P°P±Ps
P¶C-PSPeP°Pj, C¶PeC- PíP»P°PSCíCĤC, CĤ PIP°PíC-C, PSC-CÍС, CĤ (PrPëPI. CĤPsP·PrC-P»Pë
B«PµCĤPsC, PëPíPsPeP°P·P°PSPSC¶B», B«PĤCÍPsP±P»PëPIPSCíС, C-
P·P°CÍС, PsCÍСíPIP°PSPSC¶B»).

P°PíC-C, PSC-CÍС, CĤ

P°P°PSC- PíCĤPs P·P°CÍС, PsCÍСíPIP°PSPSC¶ PíCĤPµPíP°CĤP°С, Cí PŸPµPIC-PeP°CĤ
PIP°PíC-C, PSPëPj P¶C-PSPeP°Pj PIC-PrCÍСíС, PSC-. P°PsCÍP»C-PrP¶PµPSPSC¶
C, PsPeCíPëC†PSPsCÍС, C- PŸPµPIC-PeP°CĤCí C%oPsPrPs

СҗPµPиСҗPµPrCfPeC, PëPIPSPëC... PsСҗPиP°PSC-PI PSP° C, PIP°СҗPëPSP°C... PSPµ
PиСҗPµPµPrPëP»PëCfCj.

*PhP»PjPµCfP°СҗC, P°PSCз PjPµPrPsPeCfPsPjC-P» (PrC-СҗC#P° СҗPµC#PsPIPëPSP°
PŸPµPIC-PeP°СҗCз)*

P—P°CfC, PsCfCfPIP°PSPSCj P°PSC, P°PiPsPSC-CfC, C-PI СҗPµCfPµPиC, PsСҗC-PI
P°PSPиC-PsC, PµPSP·PëPSCf P†P† PиСҗPµC, PëPиPsPeP°P·P°PSPs PIP°PиC-C, PSPëPj C-
P¶C-PSPeP°Pj, CjPeC- PиP»P°PSCfСҗC, CH PIP°PиC-C, PSC-CfC, CH (PrPëPI.
СҗPµP·PrC-P»Pë B«PµСҗPµC, PëPиPsPeP°P·P°PSPSCjB», B«PhCfPsP±P»PëPIPsCfC, C-
P·P°CfC, PsCfCfPIP°PSPSCjB»).

P·PиC-PrPµPjC-PsP»PsPиC-C#PSC- PrP°PSC- CfC, PsCfPsPIPSPs СҗPëP·PëPeCf
C, PµСҗP°C, PsPиPµPSPSPsC— PrC-C— C-PSPиC-P±C-C, PsСҗC-PI PµPµCf C P†
C, СҗPëPjPµCfC, СҗC- PIP°PиC-C, PSPsCfC, C- PSPµ PrP°СҗC, CH P·PjPsPиPë P·СҗPµP±PëC, Pë
PePsPSPeСҗPµC, PSC- PIPëCfPSPsPиPePë, PsPrPSP°Pe СҗPëP·PëPe PиPsPrC-P±PSPëC...
PµC, PµPeC, C-PI PиPsPSC-CfC, Cҗ PIPëPeP»СҗC#PëC, Pë PSPµ PjPsP¶PSP°. PµPsP¶PSP°
PиСҗPëPиCfCfC, PëC, Pë, C°%Ps C-CfPSCfC” P°PSP°P»PsPиC-C#PSPëPµ СҗPëP·PëPe
PIC-PrPSPsCfPSPs P°PSC, P°PiPsPSC-CfC, C-PI СҗPµCfPµPиC, PsСҗC-PI
P°PSPиC-PsC, PµPSP·PëPSCf II, PsCfPeC-P»CHPePë PePsPSC, СҗPµP»CHPsPIP°PSC-
PµPиC-PrPµPjC-PsP»PsPиC-C#PSC- PrPsCfP»C-PrP¶PµPSPSCj C†PëC... PиСҗPµPиP°СҗP°C, C-PI
PSPµ PиСҗPµPµPrPëP»PëCfCj. PµP°C†C-C”PSC, PeP°Pj, CjPeC- PиP»P°PSCfСҗC, CH
PIP°PиC-C, PSC-CfC, CH, СҗPµPePsPjPµPSPrCfC”C, CHCfCj PиPµСҗPµPµC, Pë PSP° C-PSC€C-
PиC-PиPsC, PµPSP·PëPIPSC- PиСҗPµPиP°СҗP°C, Pë, P±PµP·PиPµPeP° CjPePëC... PиСҗPë
PIP°PиC-C, PSPsCfC, C- PrPsPµPrPµPSP°. PиPeC°%Ps PIP°PиC-C, PSC-CfC, CH
PиC-PrC, PIPµСҗPrP¶CfC”C, CHCfCj PиC-Pr C#P°Cf P»C-PeCfPIP°PSPSCj
P°PSC, P°PiPsPSC-CfC, P°PjPë СҗPµCfPµPиC, PsСҗC-PI P°PSPиC-PsC, PµPSP·PëPSCf P†P†, C, P°PePµ
P»C-PeCfPIP°PSPSCj CfP»C-Pr PSPµPиP°PµPSPs PиСҗPëPиPëPSPëC, Pë C- PI СҗP°P·C-
PиPsC, СҗPµP±Pë PиСҗPëP·PSP°C#PëC, Pë P°P»CHC, PµСҗPSP°C, PëPIPSPµ P»C-PeCfPIP°PSPSCj.

Pj PrСҗCfPиPsPjCf C- C, СҗPµC, CHPsPjCf C, СҗPëPjPµCfC, СҗP°C... P°PSC, P°PiPsPSC-CfC, Pë
СҗPµCfPµPиC, PsСҗC-PI P°PSPиC-PsC, PµPSP·PëPSCf II C#PëPSCjC, CH C, PsPeCfPëC#PSCf
PrC-СҗC PSP° PиP»C-Pr (PиСҗPëPиPSC-C#PµPSPSCj C, C†PSPeC†C-C— PSPëСҗPµPe,
PsP»C-PиPsPиC-PrСҗP°PjPSC-PsPS, P·P°C, СҗPëPjPeP° PsCfPëC,, C-PeP°C†C-C— PeC-CfC, PsPe
C#PµСҗPµPиP°) C, P° PSPsPµPSP°СҗPµPrP¶PµPSPsPиPs (PSPëСҗPµPePsPIP°
PSPµPrPsCfC, P°C, PSC-CfC, CH, P°СҗC, PµСҗC-P°P»CHPSP° PиC-PиPsC, PµPSP·C-Cj,
PиC-PиPµСҗPµPeP°P»C-C”PjC-Cj) (PrPëPI. СҗPµP·PrC-P» B«P”PsPeP»C-PSC-C#PSC- PrP°PSC- P·
P±PµP·PиPµPePëB»).

Pj СҗP°P·C- PиСҗPëPµPsPjCf P°PSC, P°PiPsPSC-CfC, C-PI СҗPµCfPµPиC, PsСҗC-PI
P°PSPиC-PsC, PµPSP·PëPSCf II PI PrСҗCfPиPsPjCf P°P±Ps C, СҗPµC, CHPsPjCf
C, СҗPëPjPµCfC, СҗC- PSPµPsP±C...C-PrPSPëPµ PePsPSC, СҗPµP»CH C,, C†PSPeC†C-C—
PSPëСҗPµPe C, P° PиСҗPµCfPµCfCf PsCfPëC,, C-PeP°C†C-C— PeC-CfC, PsPe
C#PµСҗPµPиP° C PиP»PsPrP° P·P° PrPsPиPsPjPsPиPsСҗ PJP—P”. P—P° CfC, P°PSPsPj
PSPsPµPSP°СҗPµPrP¶PµPSPëC..., PjP°C, PµСҗC- CjPePëC... PиСҗPëPµPjP°P»Pë
P°PSC, P°PiPsPSC-CfC, Pë СҗPµCfPµPиC, PsСҗC-PI P°PSPиC-PsC, PµPSP·PëPSCf II,
CfP»C-Pr CfPиPsCfC, PµСҗC-PиP°C, Pë C°%PsPrPs PjPsP¶P»PëPIPsC—
P°СҗC, PµСҗC-P°P»CHPSPsC— PиC-PиPsC, PµPSP·C-C— (PrPëPI. СҗPµP·PrC-P»Pë
B«PµСҗPµC, PëPиPsPeP°P·P°PSPSCjB», B«PhCfPsP±P»PëPIPsCfC, C-
P·P°CfC, PsCfCfPIP°PSPSCjB»).

PħPjP»PsPrPëPiC-PS (PrC-CHC±P° CħPµC±PsPIPëPSP° PŸPµPIC-PeP°CħCž)

P°P°PSC-, PsC,CħPëPjP°PSC- PI C...PsPrC- CÍPiPsCÍC,PµCħPµP¶PµPSCħ P·P°
PsP±PjPµP¶PµPSPsCħ PeC-P»CHħPeC-CÍC,Cħ PIP°PiC-C,PSPëC... P¶C-PSPsPe, PSPµ
PiPsPeP°P·P°P»Pë, C%Ps P°PjP»PsPrPëPiC-PS P°P±Ps C-PSC€C-
P°PSC,P°PiPsPSC-CÍC,Pë PeP°P»CHC†C-C"PIPëC... CħPµC†PµPiC,PSCħC-PI
CÍPiCħPëC±PëPSC¶CħC,Cħ C€PeC-PrP»PëPIPëPNë PIPiP»PëPI PSP°
P·PrPSCħPsPIBħ™ C¶ PiP»PsPrP°. PħPrPSP°Pe PiCħPë C†CHħPsPjC† C" CħPëP·PëPe
PiPsPrPsPIP¶PµPSPSC¶ C,CħPëPIP°P»PsCÍC,C- PiPsP»PsPiC-PI.

P"PsPrCžPIP°PSPSC¶ PiCħCžPrPrCħ

PħP»PjPµCÍP°CħC,P°PS PIPëPrC-P»C¶C"C,CħCÍC¶ PI PjPsP»PsPePs
P»P°PeC,C†CħC±PëC... C%C†CħC-PI. PќPµPIC-PrPsPjPs, C±Pë PiCħPsPSPëPeP°C"
PsP»PjPµCÍP°CħC,P°PS C† PiCħC†PrPSPµ PjPsP»PsPePs C† P¶C-PSPsPe.
PħPjP»PsPrPëPiC-PS PµPeCÍPeCħPµC,C†C"C,CħCÍC¶ PI PiCħC†PrPSPµ PjPsP»PsPePs
P»CħPrPëPSPë. PšP°CÍC,PeC† PjP°C,PµCħPëPSCÍCħPePsC— PrPsP·Pë,
PsC,CħPëPjP°PSPsC— PSPµPjPsPIP»C¶Pj, P±C†P»Ps PsC†C-PSPµPSPs P·P°
PjC-P¶PePIP°CħC,PëP»CHħPSPëPj PrC-P°PiP°P·PsPSPsPj 3-7 % P· PjP°PeCÍPëPjC†PjPsPj
15 %. P'PiP»PëPI P°PjP»PsPrPëPiC-PSC† PSP° PsCħPiP°PSC-P·Pj PSPµPjPsPIP»C¶C,Pë
PSPµPIC-PrPsPjPëPNë.

PµC-Pr C±P°CÍ PiPsPrC†PIP°PSPSC¶ PiCħC†PrPrCħ P·P°CÍC,PsCÍC†PIP°PSPSC¶
PiCħPµPiP°CħP°C,C† PŸPµPIC-PeP°Cħ PSPµ CħPµPePsPjPµPSP†C†C"C,CħCÍC¶ C,P°
PSP°PrP°C"C,CħCÍC¶ PiPµCħPµPIP°PiP° P°P»CHħC,PµCħPSP°C,PëPIPSPëPj
C,PµCħP°PiC-C¶Pj C-P· PeCħP°C%Pµ PrPsPIPµPrPµPSPëPj PiCħPsC,,C-P»PµPj
P±PµP·PiPµPePë PiC-Pr C±P°CÍ PiPsPrC†PIP°PSPSC¶ PiCħC†PrPrCħ, PsCÍPsP±P»PëPIPs
PiC-Pr C±P°CÍ PiPsPrC†PIP°PSPSC¶ PiCħC†PrPrCħ PSPsPIPsPSP°CħPsPrP¶PµPSPëC...
C,P° PSPµPrPsPSPsC€PµPSPëC... PSPµPjPsPIP»C¶C,.

PαPµCħC,PëP»CHħPSC-CÍC,Cħ

PµPsPIC-PrPsPjP»C¶P»Pë PiCħPs PIPëPiP°PrPePë, PePsP»Pë C† PrPµC¶PePëC...
PiP°C†C-C"PSC,C-PI, C%Ps PiCħPëPNëPjP°CħC,Cħ P±P»PsPeP°C,PsCħPë
PeP°P»CHħC†C-C"PIPëC... PeP°PSP°P»C-PI, PIPëPSPëPeP°P»Pë PsP±PsCħPsC,PSC-
P±C-PsC...C-PjC-C±PSC- P·PjC-PSPë PI PiPsP»PsPIC†C- CÍPiPµCħPjP°C,PsP·PsC—PrP°.
PħP»C-PSC-C±PSPëC... PrP°PSPëC... C%PsPrPs PiPsC,PµPSC†C-PNëPSPsPiPs
PIPiP»PëPIC† P°PjP»PsPrPëPiC-PSC† PSP° C,,PµCħC,PëP»CHħPSC-CÍC,Cħ
PSPµPrPsCÍC,P°C,PSCħPs. Pj PrPsCÍP»C-PrP¶PµPSPSC- PSP° C%C†CħP°C... P±C†P»Ps
PIPëC¶PIP»PµPSPs PSPµP±P°P¶P°PSC† PrC-Cħ PSP° C,,PµCħC,PëP»CHħPSC-CÍC,Cħ
CÍP°PjC†C-PI (PrPëPI. CħPsP·PrC-P» B«P"PsPeP»C-PSC-C±PSC- PrP°PSC- P·
P±PµP·PiPµPePëB»).

*P—PrP°C,PSC-CÍC,Cħ PIPiP»PëPIP°C,Pë PSP° C€PIPëPrPeC-CÍC,Cħ CħPµP°PeC†C-C—
PiCħPë PePµCħCžPIP°PSPSC- P°PIC,PsC,CħP°PSCÍPiPsCħC,PsPj P°P±Ps C-PSC€PëPjPë
PjPµC...P°PSC-P·PjP°PjPë.*

*PŸPµPIC-PeP°Cħ PjPsP¶Pµ PSPµP·PSP°C±PSPs P°P±Ps PiPsPjC-CħPSPs PIPiP»PëPIP°C,Pë PSP°
P·PrP°C,PSC-CÍC,Cħ PePµCħC†PIP°C,Pë C,CħP°PSCÍPiPsCħC,PSPëPjPë P·P°CÍPsP±P°PjPë C,P°*

PsP±CÍP»CfPiPsPICfPIP°C,Pë C-PSC€C- PjPµC...P°PSC-P·PjPë.

P†PSPsPrC- CÍ PíP°C†C-C" PSC,C-PI, C%Ps PíCĤPëPNëPjP°CĤC,CH PíC-PíPsC,PµPSP·PëPIPSC- P·P°CÍPsP±Pë, PjPsP¶CfC,CH CÍPíPsCÍC,PµCĤC-PíP°C,PëCÍCĬ PíPsP»PsPIPSC- P±PsP»C-, P·P°PíP°PjPsCĤPsC†PµPSPSCĬ, PSCfPrPsC,P° C- PíC-PrPIPëC%PµPSP° CÍC,PsPjP»CĤPIP°PSC-CÍC,CH, PIPSP°CÍP»C-PrPsPe C†PsPiPs PjPsP¶Pµ P·PSPëP¶CfPIP°C,PëCÍCĬ CĤPµP°PeC†C-CĬ. PŸP»C-Pr PrPsC,CĤPëPjCfPIP°C,PëCÍCĬ PsP±PµCĤPµP¶PSPsCÍC,C-, PsCÍPsP±P»PëPIPs PSP° PíPsC†P°C,PeCf P»C-PeCfPIP°PSPSCĬ.

PŸPíPsCÍC-P± P·P°CÍC,PsCÍCžPIP°PSPSCĬ C,P° PrPsP·Pë.

P"PsCĤPsCÍP»C-

P PµPePsPjPµPSPPrPsPIP°PSP° PrPsP·P° PíCĤPµPíP°CĤP°C,Cf PŸPµPIC-PeP°CĤ -1 C,P°P±P»PµC,PeP° PSP° PrPsP±Cf.

PŸPµPIC-PeP°CĤ, 20 PjPi/5 PjPi, PjPsP¶PSP° PíCĤPëP·PSP°C†P°C,Pë PíP°C†C-C" PSC,P°Pj P·PSPµPrPsCÍC,P°C,PSCHPsCĤ CĤPµP°PeC†C-C"Ĥ PsCĤPíP°PSC-P·PjCf PSP° PjPsPSPsC,PµCĤP°PíC-CĤ PsP»PjPµCÍP°CĤC,P°PSCf PjPµPrPsPeCÍPsPjC-P»PsPj Pí PrPsP·C- 20 PjPi P°P±Ps P°PjP»PsPrPëPíC-PSPsPj Pí PrPsP·C- 5 PjPi.

PŸPµPIC-PeP°CĤ, 40 PjPi/5 PjPi, PjPsP¶PSP° PíCĤPëP·PSP°C†P°C,Pë PíP°C†C-C" PSC,P°Pj P·PSPµPrPsCÍC,P°C,PSCHPsCĤ CĤPµP°PeC†C-C"Ĥ PsCĤPíP°PSC-P·PjCf PSP° C,PµCĤP°PíC-CĤ PíCĤPµPíP°CĤP°C,PsPj PŸPµPIC-PeP°CĤ, 20 PjPi/5 PjPi.

PŸPµPIC-PeP°CĤ, 40 PjPi/10 PjPi, PjPsP¶PSP° PíCĤPëP·PSP°C†P°C,Pë PíP°C†C-C" PSC,P°Pj P·PSPµPrPsCÍC,P°C,PSCHPsCĤ CĤPµP°PeC†C-C"Ĥ PsCĤPíP°PSC-P·PjCf PSP° C,PµCĤP°PíC-CĤ PíCĤPµPíP°CĤP°C,PsPj PŸPµPIC-PeP°CĤ, 40 PjPi/5 PjPi.

P"Ps PíCĤPëP·PSP°C†PµPSPSCĬ PePsPjP±C-PSPsPIP°PSPsPiPs PíCĤPµPíP°CĤP°C,Cf P·C,,C-PeCÍPsPIP°PSPëPjPë PrPsP·P°PjPë P°PeC,PëPIPSPëC... PePsPjPíPsPSPµPSC,C-PI CĤPµPePsPjPµPSPPrCfC"C,CHCÍCĬ PíPsPµC,P°PíPSPëPNë PíC-PrP±C-CĤ PrPsP· C†PëC... PePsPjPíPsPSPµPSC,C-PI CĬPe PjPsPSPsPíCĤPµPíP°CĤP°C,C-PI. Pj CĤP°P·C- PSPµPsP±C...C-PrPSPsCÍC,C- PjPsP¶P»PëPIP° PíCĤCĬPjP° P·P°PjC-PSP° PjPsPSPsPíCĤPµPíP°CĤP°C,C-PI PSP° PePsPjP±C-PSPsPIP°PSPëPNë PíCĤPµPíP°CĤP°C,.

P" P»CĬ P·CĤCfC†PSPsCÍC,C- PíP°C†C-C" PSC,Pë, CĬPeC- PsC,CĤPëPjCfCĤC,CH PsP»PjPµCÍP°CĤC,P°PSCf PjPµPrPsPeCÍPsPjC-P» C- P°PjP»PsPrPëPíC-PS Cf PIPëPíP»CĬPrC- PsPeCĤPµPjPëC... C,P°P±P»PµC,PsPe, PjPsP¶CfC,CH P±CfC,Pë PíPµCĤPµPIPµPrPµPSC- PSP° C,P°P±P»PµC,PePë PŸPµPIC-PeP°CĤ, C%Ps PjC-CÍC,CĬC,CH C†C- PePsPjPíPsPSPµPSC,Pë PíP°PSP°P»PsPíC-C†PSPëC... PrPsP·P°C....

PµCĤPµPíP°CĤP°C, PŸPµPIC-PeP°CĤ PjPsP¶PSP° PíCĤPëPNëPjP°C,Pë PSPµP·P°P»PµP¶PSPs PIC-Pr PíCĤPëPNëPsPjCf C-P¶C-.

PµP°C†C-C" PSC,Pë P»C-C,PSCHPsPiPs PIC-PeCž (PIC-PePsPj PIC-Pr 65 CĤPsPeC-PI)

PíPe PíCĤP°PIPëP»Ps, PrP»CĬ PíP°C†C-C" PSC,C-PI P»C-C,PSCHPsPiPs PIC-PeCf PSPµ PíPsC,CĤC-P±PSP° P·PjC-PSP° CĤPµPePsPjPµPSPPrPsPIP°PSPsC— PrPsP·Pë PíCĤPµPíP°CĤP°C,Cf, PíCĤPsC,Pµ PíC-PrPIPëC%CfPIP°C,Pë PrPsP·Cf PíCĤPµPíP°CĤP°C,Cf CÍP»C-Pr P·PsP±PµCĤPµP¶PSC-CÍC,CĤ (PrPëPI. CĤPsP·PrC-P»Pë B«PĥCÍPsP±P»PëPIPsCÍC,C-

Р·°СГГ,PsСГГрРIP°PSPSCЦБ», В«РᄁР°СГГРjР°PePsPeC-PSPμC,PePeP°B»).

РμСГГPe P·P±C-P»СГГCεPμPSPSC- PrPsP·Pe PsP»PjPμCГГ°СГГC,Р°PSCr PjPμPrPsPeCГГPsPjC-P»Cr PrPs PjP°PeCГГPePjP°P»СГГPSPSc— (40 PjPi PSP° PrPsP±Cr) Cr PIP°CГГ-C”PSC,Р° CГГP»C-Pr СГГPμC,РμP»СГГPSPs PePsPSC,СГГPsP»СГГPIP°C,Pe P°СГГC,РμСГГC-P°P»СГГPSPePNε C,PeCГГPe.

РμPsСГГCáCεPμPSPSCЦ C,,CáPSPeCГГC-C— PSPeСГГPsPe

РᄁР°PeCГГPePjP°P»СГГPSP° PrPsP·P° PsP»PjPμCГГ°СГГC,Р°PSCr PjPμPrPsPeCГГPsPjC-P»Cr PrP»CЦ PIP°CГГ-C”PSC,C-PI P· PIPsСГГCрCεPμPSPSCЦPj C,,CрPSPeCГГC-C— PSPeСГГPsPe PIC-Pr P»PμPiPePsPiPs PrPs PIPsPjC-СГГPSPsPiPs CГГC,CрPμPμPSCЦ (PeP»C-СГГPμPSCГГ PeCГГPμP°C,PePSC-PSCr 20-60 PjP»/C...PI) CГГC,Р°PSPsPIPeC,СГГ 20 PjPi PsPrPePS СГГP°P· PSP° PrPsP±Cr, PsCГГPeC-P»СГГPePe PrPsCГГPIC-Pr P·P°CГГC,PsCГГCрРIP°PSPSCЦ P±C-P»СГГCε PIPeCГГPsPePeC... PrPsP· CГГC-PNε PiCГГCрPiC- PIP°CГГ-C”PSC,C-PI PsP±PjPμPμPμPSPePNε. PŸPμPIC-PeP°СГГ PSPμ СГГPμPePsPjPμPSPPrCГГ”C,СГГCГГCЦ PIP°CГГ-C”PSC,Р°Pj P· PIPsСГГCрCεPμPSPSCЦPj C,,CрPSPeCГГC-C— PSPeСГГPsPe C,CЦPμPePsPiPs CГГC,CрPμPμPSCЦ (PeP»C-СГГPμPSCГГ PeCГГPμP°C,PePSC-PSCr < 20 PjP»/C...PI) (PrPePI. СГГPsP·PrC-P»Pe В«PᄁCГГPsP±P»PePIPсГГC,C- P·P°CГГC,PsCГГCрРIP°PSPSCЦБ», В«РᄁР°СГГРjР°PePsPeC-PSPμC,PePeP°B»).

РμСГГPe P·P°CГГC,PsCГГCрРIP°PSPSC- PиСГГPμPIP°СГГP°C,Cр PIP°CГГ-C”PSC,Р°Pj P· PIPsСГГCрCεPμPSPSCЦPj C,,CрPSPeCГГC-C— PSPeСГГPsPe PIPsPjC-СГГPSPsPiPs CГГC,CрPμPμPSCЦ СГГPμPePsPjPμPSPPrCГГ”C,СГГCГГCЦ PePsPSC,СГГPsP»СГГPIP°C,Pe PePsPSCГГPμPSC,СГГP°CГГ-CГГ PeP°P»C-СГГ C,Р° PeCГГPμP°C,PePSC-PSCr.

РμPsСГГCáCεPμPSPSCЦ C,,CáPSPeCГГC-C— PμC±C-PSPePe

РμСГГPμPIP°СГГP°C, PŸPμPIC-PeP°СГГ P· PsP±PμСГГPμPμPSC-CГГC,СГГ PиСГГPeP·PSP°C±P°СГГC,СГГ PIP°CГГ-C”PSC,Р°Pj P· PIPsСГГCрCεPμPSPSCЦPj C,,CрPSPeCГГC-C— PμC±C-PSPePe PIC-Pr P»PμPiPePsPiPs PrPs PIPsPjC-СГГPSPsPiPs CГГC,CрPμPμPSCЦ (PrPePI. СГГPsP·PrC-P»Pe В«PᄁCГГPsP±P»PePIPсГГC,C- P·P°CГГC,PsCГГCрРIP°PSPSCЦБ», В«РᄁР°СГГРjР°PePsPeC-PSPμC,PePeP°B»). РμСГГPe PIPsPjC-СГГPSPsPjCr PIPsСГГCрCεPμPSPSC- C,,CрPSPeCГГC-C— PμC±C-PSPePe PsP»PjPμCГГ°СГГC,Р°PSCr PjPμPrPsPeCГГPsPjC-P» PиСГГPeP·PSP°C±P°СГГC,СГГ Cr PIPsC±P°C,PePsPIC-PNε PrPsP·C- 10 PjPi PsPrPePS СГГP°P· PSP° PrPsP±Cr. РᄁР°PeCГГPePjP°P»СГГPSP° PrPsP·P° PrP»CЦ C,Р°PePeC... PIP°CГГ-C”PSC,C-PI PSPμ PIPsPIPePSPSP° PμСГГPμPIPeC%CrPIP°C,Pe 20 PjPi PsPrPePS СГГP°P· PSP° PrPsP±Cr. Pj СГГP°P·C- CГГCрPиCрC,PSCᄁPsC— C,РμСГГP°PiC-C— PrC-CрСГГPμC,PePeP°PjPe C- (P°P±Ps) C-PSCεPePjPe PiC-PiPsC,РμPSP·PePIPSPePjPe PиСГГPμPIP°СГГP°C,Р°PjPe PIP°CГГ-C”PSC,Р°Pj P· PIPsСГГCрCεPμPSPSCЦPj C,,CрPSPeCГГC-C— PμC±C-PSPePe СГГPμPePsPjPμPSPPrCГГ”C,СГГCГГCЦ СГГPμC,РμP»СГГPSPμ CГГPиPsCГГC,РμСГГPμPμPμPSPSCЦ P·P° СГГC-PIPSμPj P°СГГC,РμСГГC-P°P»СГГPSPsPiPs C,PeCГГPeCr C- C,,CрPSPeCГГC-C”СГГ PSPeСГГPsPe. P”PsCГГPIC-Pr P·P°CГГC,PsCГГCрРIP°PSPSCЦ PsP»PjPμCГГ°СГГC,Р°PSCr PjPμPrPsPeCГГPsPjC-P»Cr PIP°CГГ-C”PSC,Р°Pj P· PIPsСГГCрCεPμPSPSCЦPj C,,CрPSPeCГГC-C— PμC±C-PSPePe C,CЦPμPePsPiPs CГГC,CрPμPμPSCЦ PIC-PrCГГCрC,PSC-PNε.

PiPe C- PиСГГPe P·P°CГГC,PsCГГCрРIP°PSPSC- C-PSCεPeC... P°PSC,Р°PiPsPSC-CГГC,C-PI PeP°P»СГГCГГ-CГГ, Cr PIP°CГГ-C”PSC,C-PI P· PrPeCГГC,,CрPSPeCГГC-C”СГГ PμC±C-PSPePe PIPsPrPsPIPиCрC”C,СГГCГГCЦ C±P°CГГ PSP°PiC-PIPIPePμPrPμPSPSCЦ P°PjP»PsPrPePiC-PSCr; СГГPμPePsPjPμPSPPr°CГГ-C— P· PrPsP·CрРIP°PSPSCЦ PSPμ PICГГC,Р°PSPsPIP»PμPSC-. PŸPsPjCr C,Р°PePePj PIP°CГГ-C”PSC,Р°Pj PиСГГPμPIP°СГГP°C, PŸPμPIC-PeP°СГГ CГГP»C-Pr PиСГГPeP·PSP°C±P°C,Pe P· PsP±PμСГГPμPμPSC-CГГC,СГГ. РᄁР°СГГРjР°PePsPeC-PSPμC,PePeP° P°PjP»PsPrPePiC-PSCr Cr PIP°CГГ-C”PSC,C-PI P· PIPsСГГCрCεPμPSPSCЦPj C,,CрPSPeCГГC-C—

PiPμC±C-PSPePë C, CIP¶¶PePsPiPs CÍC, CíPiPμPSCÍ PSpμ PIPëPIC±PμPSP°. PμC¶PëPNëPsPj P°PjP»PsPrPëPiC-PSCr PiP°C†C-C" PSC, P°Pj P· PiPsC¶CíC€PμPSPSCÍPj C,, CíPSPeC†C-C— PiPμC±C-PSPePë C, CIP¶¶PePsPiPs CÍC, CíPiPμPSCÍ CÍP»C-Pr PiPsC±PëPSP° C, Pë P· PSP°PNëPSPëP¶C±PsC— PrPsP·Pë C- PiPsPIC-P»CHPSPs C—C— PiC-PrPIPëC% CíPIP° C, Pë. PÿPμPIC-PeP°C¶ PiC¶PsC, PëPiPsPeP°P·P°PSPëPNë PiP°C†C-C" PSC, P°Pj P· PiPsC¶CíC€PμPSPSCÍPj C,, CíPSPeC†C-C— PiPμC±C-PSPePë C, CIP¶¶PePsPiPs CÍC, CíPiPμPSCÍ (PrPëPI. C¶PsP·PrC-P» B«PμC¶PsC, PëPiPsPeP°P·P°PSPSCÍB»).

PÿPiPsCÍC-P± P·P°CÍC, PsCÍC¿PIP°PSPSCÍ

PÿP°P±P»PμC, PePë PiC¶PsPePsPIC, CíC¶C, CH C†C-P»PëPjPë, P·P°PiPëPIP°C¶C±Pë PrPsCÍC, P°C, PSC¶PsC¶ PeC-P»CHPeC-CÍC, C¶ C¶C-PrPëPSPë (PSP°PiC¶PëPeP»P°Pr CÍPeP»CÍPSPePsC¶ PIPsPrPë). PÿP°P±P»PμC, PePë PSPμ CÍP»C-Pr C¶PsP·P¶PsPICrPIP° C, Pë. PμC¶PμPiP°C¶P° C, C¶PμPePsPjPμPSPPrCíC" C, CHCÍCÍ PiC¶PëPNëPjP° C, Pë C%PsPrPSCÍ PI PsPrPëPS C- C, PsPNë P¶Pμ C±P°CÍ.

P" C-C, Pë.

P'PμP·PiPμPeP° C, P° PμC,, PμPeC, PëPIPSC-CÍC, CH P·P°CÍC, PsCÍCíPIP°PSPSCÍ PiC¶PμPiP°C¶P° C, Cí PÿPμPIC-PeP°C¶ PrC-C, CÍPj (PIC-PePsPj PrPs 18 C¶PsPeC-PI) PSPμ PrPsCÍP»C-PrP¶CíPIP°P»PëCÍCH. P" P°PSC- PIC-PrCÍCíC, PSC-.

PμPμC¶PμPrPsP·C¿PIP°PSPSCÍ.

PÿPëPjPiC, PsPjPë

P'PëPiP°PrPeC-PI PiPμC¶PμPrPsP·CíPIP°PSPSCÍ PiC¶PμPiP°C¶P° C, Cí PÿPμPIC-PeP°C¶ PSPμ P·P°C,, C-PeCÍPsPIP°PSPs. PkP°PNëP±C-P»CHC€ PIC-C¶PsPiC-PrPSPëPjPë PμC,, PμPeC, P°PjPë PiPμC¶PμPrPsP·CíPIP°PSPSCÍ PsP»PjPμCÍP°C¶C, P°PSCr PjPμPrPsPeCÍPsPjC-P»Cí C" PiC-PiPsC, PμPSP·C-CÍ C- C, P°C...C-PeP°C¶PrC-CÍ; C, P°PePsP¶ PjPsP¶Pμ CÍPiPsCÍC, PμC¶C-PiP° C, PëCÍCÍ P±C¶P°PrPëPeP°C¶PrC-CÍ PI C¶P°P·C- PiP°C¶P°CÍPëPjPiP° C, PëC±PSPsC— CÍC, PëPjCíP»CÍC†C-C— (P±P»CíPeP°C¶C±PsPiPs PSPμC¶PIP°). PμPμC¶PμPrPsP·CíPIP°PSPSCÍ P°PjP»PsPrPëPiC-PSCr PjPsP¶Pμ PiC¶PëP·PIPμCÍC, Pë PrPs PSP°PrPjC-C¶PSPsPiPs C¶PsP·C€PëC¶PμPSPSCÍ PiPμC¶PëC,, PμC¶PëC±PSPëC... CÍCíPrPëPS, C%Ps CÍCíPiC¶PsPiPsPrP¶CíC" C, CHCÍCÍ PIPëC¶P°P¶PμPSPëPj P·PSPëP¶PμPSPSCÍPj P°C¶C, PμC¶C-P°P»CHPSPsPiPs C, PëCÍPeCí, C-, PjPsP¶P»PëPIs, PrPs C¶PμC,, P»PμPeC, PsC¶PSPsC— C, P°C...C-PeP°C¶PrC-C—. P,, PiPsPIC-PrPsPjP»PμPSPSCÍ PiC¶Ps C, C¶PëPIP°P»Cí C, CIP¶¶PeCí PiPμPSPμC¶P°P»C-P·PsPIP°PSCr PiC-PiPsC, PμPSP·C-C¶, P°P¶ PrPs C€PsPeCí C-P· P»PμC, P°P»CHPSPëPj PSP°CÍP»C-PrPePsPj.

P C-PrPePs PiPsPIC-PrPsPjP»CÍC" C, CHCÍCÍ PiC¶Ps PSPμPeP°C¶PrC-PsPiPμPSPSPëPNë PSP°P±C¶CÍPe P»PμPiPμPSC¶ CÍPe PSP°CÍP»C-PrPsPe PiPμC¶PμPrPsP·CíPIP°PSPSCÍ P°PjP»PsPrPëPiC-PSCr, C%Ps PjPsP¶Pμ PiC¶PsCÍPIP»CÍC, PëCÍCÍ P· PIC-PrCÍC, C¶PsC±PμPSPëPj PiPsC±P° C, PePsPj (24B¶"48 PiPsPrPëPS PiC-CÍP»CÍ PiC¶PëPNëPsPjCí) C, P° PiPsC, C¶PμP±CíC" C€C, CíC±PSPsC— PIPμPSC, PëP»CÍC†C-C— P»PμPiPμPSC¶. P P°PSPSC- C¶PμP°PSC-PjP°C†C-PNëPSC- P·P°C...PsPrPë (PIPeP»C¶C±P°C¶C±Pë PiPμC¶PμPIP°PSC, P°P¶PμPSPSCÍ C¶C-PrPëPSPsC¶) PrP»CÍ PiC-PrC, C¶PëPjP°PSPSCÍ PiPμC¶C,, CíP·C-C— C, P° CÍPμC¶C†PμPIPsPiPs PIPëPePëPrCí PjPsP¶CíC, CH C¶PsP·PiP»CÍPrP° C, PëCÍCÍ CÍPe PiC¶PsPiPsPeCíC¶C- C,, P°PeC, PsC¶Pë.

P>C-PeCáPIP°PSPSCII

PİPeC%Ps PiCḡPµPiP°CḡP°C, P±CfPI PiCḡPēPNēPSCIIIC,PēPNē PSPµC%PsPrP°PIPSPs, PİPsPeP°P°PSPs PiCḡPsPjPēPIP°PSPSCII CēP»CfPSPeP°. Pj P·PrPsCḡPsPIPēC... PrPsP±CḡPsPIPsP»CHC†C-PI PiCḡPēP·PSP°C†PµPSPSCII P°PeC,PēPIPsPIP°PSPsPiPs PICfPiC-P»P»CII PSPµPiP°PNēPSPs P°P±Ps PiCḡPsC,CIIPIPsPj 2 PiPsPrPēPS PiC-CfP»CII PiCḡPēPNēPsPjCf P°PjP»PsPrPēPiC-PSCf PICfPµCḡPµPrPēPSCf C-CfC,PsC,PSPs P·PSPēP¶CfC” PICfPjPsPeC,CfPIP°PSPSCII C†C-C”C— CḡPµC†PsPIPēPSPē.

PµCḡPē PIPēPSPēPePSPµPSPSC- PeP»C-PSC-C†PSPs P·PSP°C†CfC%PsC— PiC-PiPsC,PµPSP·C-C— PIPSP°CfP»C-PrPsPe PiPµCḡPµPrPsP·CfPIP°PSPSCII PiCḡPµPiP°CḡP°C,PsPj PŸPµPIC-PeP°Cḡ PiPsC,CḡC-P±PSP° P°PeC,PēPIPSP° PiC-PrC,CḡPēPjPeP° CfPµCḡC†PµPIPs-CfCfPrPēPSPSPsC— CfPēCfC,PµPjPē, PIPeP»CḡC†P°CḡC†Pē CḡPµC,PµP»CHPSPēPNē PjPsPSC-C,PsCḡPēPSPi C,,CfPSPeC†C-C— CfPµCḡC†CII C- P»PµPiPµPSC-PI, PiCḡPēPiC-PrPSCIIIC,Pµ PiPsP»PsP¶PµPSPSCII PSPēP¶PSC-C... PeC-PSC†C-PIPsPe, PePsPSC,CḡPsP»CH PsP±bḡ™ C”PjCf C†PēCḡPeCfP»CḡCḡC†PsC— PeCḡPsPIC- C,P° PrC-CfCḡPµP·Cf. P”P»CII PIC-PrPSPsPIP»PµPSPSCII CfCfPrPēPSPSPsPiPs C,PsPSCfCfCf C,P° P°CḡC,PµCḡC-P°P»CHPSPsPiPs C,PēCfPeCf PjPsP¶Pµ P±CfC,Pē PePsCḡPēCfPSPēPj P·P°CfC,PsCfCfPIP°PSPSCII CfCfPrPēPSPsP·PICfP¶CfPIP°P»CHPSPēC... P·P°CfPs±C-PI PiCḡPē PIC-PrCfCfC,PSPsCfC,C- PiCḡPsC,PēPiPsPeP°P°PSCḡ. P”P»CII CfCfCfPSPµPSPSCII P±P»PsPeP°PrPē PeP°P»CHC†C-C”PIPēC... PeP°PSP°P»C-PI CḡPµPePsPjPµPSPPrCfC”C,CHCfCII PeP°P»CHC†C-Cḡ PiP»CḡPePsPSP°C, PIPSCfC,CḡC-CēPSCḡPsPIPµPSPSPs.

PḡCfPeC-P»CHPePē P°PjP»PsPrPēPiC-PS P·PSP°C†PSPsCḡ PjC-CḡPsCḡ P·PiBḡ™ CII P·CfC”C,CHCfCII P· P±C-P»PeP°PjPē PeCḡPsPIC-, PNēPsPiPs PIPēPIPµPrPµPSPSCII P·P° PrPsPiPsPjPsPiPsCḡ PrC-P°P»C-P·Cf PjP°P»PsPNēPjPsPIC-CḡPSPµ. P†PSC,,PsCḡPjP°C†C-CII PiCḡPs PjPsP¶P»PēPIC-CfC,CH PIPēPIPµPrPµPSPSCII PsP»PjPµCfP°CḡC,P°PSCf P·P° PrPsPiPsPjPsPiPsCḡ PrC-P°P»C-P·Cf PIC-PrCfCfC,PSCII.

PµPsP±C-C†PSC- CḡPµP°PeC†C-C—.

PkP°PNēC†P°CfC,C-CēPēPjPē PiPsP±C-C†PSPēPjPē CḡPµP°PeC†C-CII PjPē, C%Ps PIPēPSPēPeP°P»Pē PiC-Pr C†P°Cf P·P°CfC,PsCfCfPIP°PSPSCII PiCḡPµPiP°CḡP°C,Cf PŸPµPIC-PeP°Cḡ, P±CfP»Pē PiPµCḡPēC,,PµCḡPēC†PSC- PSP°P±CḡCII PePē (11,3 %), PiPsP»PsPIPSPēPNē P±C-P»CH (5,3 %) C,P° P·P°PiP°PjPsCḡPsC†PµPSPSCII (4,5 %).

PµPsP±C-C†PSC- CIIPIPēC%P°, C%Ps CfPiPsCfC,PµCḡC-PiP°P»PēCfCII PiC-Pr C†P°Cf PeP»C-PSC-C†PSPēC... PrPsCfP»C-PrP¶PµPSCḡ C,P° PiCḡPsC,CIIPIPsPj PiC-CfP»CII CḡPµC”CfC,CḡP°C†C-PNēPSPēC... PrPsCfP»C-PrP¶PµPSCḡ P· P±PµP·PiPµPePē, P° C,P°PePsP¶ PiPsP±C-C†PSC- CḡPµP°PeC†C-C—, PiCḡPs CIIPeC- P±CfP»Pē CfPiPsPSC,P°PSPSC- PiPsPIC-PrPsPjP»PµPSPSCII, PiCḡPµPrCfC,P°PIP»PµPSPs Cf PSPēP¶C†PµPSP°PIPµPrPµPSC-PNē C,P°P±P»PēC†C-. PḡCḡC-Pj C†CHPsPiPs, Cf C,P°P±P»PēC†C- PSP°PIPµPrPµPSPs PiPsP±C-C†PSC- CḡPµP°PeC†C-C—, C%Ps CfPiPsCfC,PµCḡC-PiP°P»PēCfCII PSP° C,P»C- P·P°CfC,PsCfCfPIP°PSPSCII PePsP¶PSPsPiPs P· PrC-CḡC†PēC... PePsPjPiPsPSPµPSC,C-PI PiCḡPµPiP°CḡP°C,Cf PsPeCḡPµPjPs (PsP»PjPµCfP°CḡC,P°PSCf PjPµPrPsPeCfPsPjC-P»Cf C,P° P°PjP»PsPrPēPiC-PSCf) P· CfCḡP°C...CfPIP°PSPSCII Pj C—C...PSCḡPsPiPs PICfC,P°PSPsPIP»PµPSPsPiPs PiCḡPsC,,C-P»Cḡ P±PµP·PiPµPePē.

P”P»CII PeP»P°CfPēC,,C-PeP°C†C-C— C†P°CfC,PsC,Pē PIPēPSPēPePSPµPSPSCII

$PiPsP \pm C - C \neq PSP \ddot{e} C \dots$ $C \ddot{H} P \mu P^\circ P e C \ddot{t} C - P N \ddot{e}$ $PIP \ddot{e} P e P s C \ddot{H} P \ddot{e} C \ddot{t} C, P s P I C \ddot{t} P I P^\circ P \gg P^\circ C \ddot{t} C \ddot{t} C, P^\circ P e P^\circ$
 $C, P \mu C \ddot{H} P j C - P S P s P \gg P s P i C - C \ddot{t} C: P r C \ddot{t} P \mu C \neq P^\circ C \ddot{t} C, P s (\geq 1/10); C \neq P^\circ C \ddot{t} C, P s (\geq 1/100 P r P s < 1/10);$
 $P S P \mu C \neq P^\circ C \ddot{t} C, P s (P I C - P r \geq 1/1000 P r P s < 1/100); C \ddot{H} C - P r P e P s (P I C - P r \geq 1/10000 P r P s < 1/1000);$
 $P r C \ddot{t} P \mu C \ddot{H} C - P r P e P s (< 1/10000); C \neq P^\circ C \ddot{t} C, P s C, P^\circ P S P \mu P I C - P r P s P j P^\circ (C \neq P^\circ C \ddot{t} C, P s C, C \ddot{t}$
 $P I C \ddot{t} C, P^\circ P S P s P I P \ddot{e} C, P \ddot{e} P S P \mu P j P s P \mu P \gg P \ddot{e} P I P s P \cdot P^\circ P S P^\circ C \ddot{t} P I P S P \ddot{e} P j P \ddot{e} P r P^\circ P S P \ddot{e} P j P \ddot{e}.$

MedDRA PhC \ddot{H} PiP \ddot{e} PSP \ddot{e} C, P \circ C \ddot{t} P \ddot{e} C \ddot{t} C, P μ PjP \ddot{e}	P μ P s P \pm C - C \neq P S P \ddot{e} P r C - C \ddot{t}	P \ddot{S} P \circ C \ddot{t} C, P s C, P \circ P μ P s P j P \pm C - P S P \circ C \ddot{t} C - C \ddot{t} P s P \gg P j P μ C \ddot{t} P \circ C $\ddot{H} C, P^\circ$ P S C \ddot{t} C, P \circ P \circ P j P \gg P s P r P \ddot{e} P i C - P S C \ddot{t}	PhP \gg PjP μ C \ddot{t} P \circ C $\ddot{H} C, P^\circ$ PS	P μ PjP \gg P s P r P \ddot{e} P i C - P S
P - P \pm PsPeC \ddot{e} PeC \ddot{H} PsPIC- C, P \circ P \gg C-PjC, P \circ C, P \ddot{e} C \neq PSPsC - C \ddot{t} P \ddot{e} C \ddot{t} C, P μ PjP \ddot{e}	P \cdot P μ PN \ddot{e} PePsC \neq P \ddot{e} C, PsPiP μ PSC-C \ddot{t}		PkP μ C \neq P \circ C \ddot{t} C, Ps	P \circ C \neq P μ P μ C $\ddot{H} C - P r P e P s$
P - P \pm PsPeC \ddot{e} C-PjC \neq PSPSPsC - C \ddot{t} P \ddot{e} C \ddot{t} C, P μ PjP \ddot{e}	P μ P \gg P μ C \ddot{H} PiC-C \neq PSP \circ C \ddot{H} P μ P \circ PeC \ddot{t} C-C \ddot{t} /P i C - P i P μ C \ddot{H} C \neq C \ddot{t} C, P \gg P \ddot{e} P i C - C \ddot{t} C, C \ddot{H} P r P s P i C \ddot{H} P μ P i P \circ C \ddot{H} P \circ C, C \ddot{t}	P C - P r P e P s		P \circ C \neq P μ P μ C $\ddot{H} C - P r P e P s$
P μ PsC \ddot{H} C \ddot{e} C \ddot{e} P μ PSPSC \ddot{t} P \cdot P \pm PsPeC \ddot{e} C...P \circ C \ddot{H} C \neq C \ddot{t} P i P \circ PSPSC \ddot{t} C, P \circ P s P \pm P j C - P S C \ddot{e} C \ddot{H} P μ C \neq PsPiP \ddot{e} PS	P \circ C - P i P μ C \ddot{H} P i P \gg C - P e P μ P j C - C \ddot{t} P \circ C - P i P μ C \ddot{H} P e P \circ P \gg C - C \cdot P j C - C \ddot{t} P \circ C - P i P μ C \ddot{H} C, C \ddot{H} P \ddot{e} P i P \gg C - C \neq P μ C \ddot{H} P \ddot{e} P r P μ P j C - C \ddot{t} P \circ C - P i P μ C \ddot{H} C \ddot{H} P e P e P μ P j C - C \ddot{t}	PkP μ C \neq P \circ C \ddot{t} C, Ps	P C - P r P e P s	P \circ C \neq P μ P μ C $\ddot{H} C - P r P e P s$
P - P \pm PsPeC \ddot{e} P i C \ddot{t} P \ddot{e} C...C - P e P \ddot{e}	P \ddot{Y} P i P \gg C \ddot{t} C, P \circ P S C - C \ddot{t} C, C \ddot{H} C \ddot{t} P i C - P r P s P j P s C \ddot{t} C, C - P \circ P μ P i C \ddot{H} P μ C \ddot{t} C - C \ddot{t} P \circ P μ P \cdot C \ddot{t} P s P S P S C \ddot{t} P \circ C \ddot{H} P \circ C, C - P i P \gg P \ddot{e} P i C - C \ddot{t} C, C \ddot{H} P - P S P \ddot{e} P μ P S P S C \ddot{t} P \gg C - P \pm C - P r P s P - P j C - P S P \circ P S P \circ C \ddot{t} C, C \ddot{H} P s C \ddot{t} (C \ddot{t} C, P s P j C \ddot{t} C \neq P \ddot{e} C \ddot{t} P \gg C - C, C \ddot{H} P \ddot{e} P i P s P i P \circ)	PkP μ C \neq P \circ C \ddot{t} C, Ps		P C - P r P e P s PkP μ C \neq P \circ C \ddot{t} C, Ps PkP μ C \neq P \circ C \ddot{t} C, Ps PkP μ C \neq P \circ C \ddot{t} C, Ps
P - P \pm PsPeC \ddot{e} PSP μ C \ddot{H} P i P s P i P s C - C \ddot{t} P \ddot{e} C \ddot{t} C, P μ PjP \ddot{e}	P - P \circ P i P \circ P j P s C \ddot{H} P s C \neq P μ P S P S C \ddot{t} P \circ P \ddot{e} C \ddot{t} P i P μ P i P - C - C \ddot{t} P \circ P s P \gg P s P i P S P \ddot{e} P \ddot{e} P \pm C - P \gg C \ddot{H}	P \ddot{S} P \circ C \ddot{t} C, P s	P \ddot{S} P \circ C \ddot{t} C, P s	P \ddot{S} P \circ C \ddot{t} C, P s PkP μ C \neq P \circ C \ddot{t} C, Ps P \ddot{S} P \circ C \ddot{t} C, P s P \ddot{S} P \circ C \ddot{t} C, P s (P s C \ddot{t} P s P \pm P \gg P \ddot{e} P i P s P S P \circ P i P s C \neq P \circ C, P e C \ddot{t} P \gg C - P e C \ddot{t} P i P \circ P S P S C \ddot{t}) P \circ C \neq P μ P μ C $\ddot{H} C - P r P e P s$
P - P \pm PsPeC \ddot{e} PsC \ddot{H} P i P \circ P S C - P i P \cdot PsC \ddot{H} C \ddot{e}	P μ PsC \ddot{H} C \ddot{e} P μ P S P S C \ddot{t} P \cdot PsC \ddot{H} C \ddot{t} (C \ddot{t} C, P s P j C \ddot{t} C \neq P \ddot{e} C \ddot{t} P \gg C - P r P \ddot{e} P i P \gg P s P i C - C \ddot{t})			P \ddot{S} P \circ C \ddot{t} C, P s
P - P \pm PsPeC \ddot{e} PsC \ddot{H} P i P \circ P S C - P i C \ddot{t} P \gg C \ddot{e} C...C \ddot{e} C - C \ddot{H} C - P i P S P s P i P \circ P i P \ddot{e}	P \ddot{E} C \ddot{t} P j P i P i C \ddot{t} C...P \circ C... P \circ P μ C \ddot{H} C, P \ddot{e} P i P s	PkP μ C \neq P \circ C \ddot{t} C, Ps	PkP μ C \neq P \circ C \ddot{t} C, Ps	PkP μ C \neq P \circ C \ddot{t} C, Ps
P - P \pm PsPeC \ddot{e} C \ddot{t} P μ C \ddot{H} C \ddot{t} C \ddot{t}	P \ddot{Y} C, P μ P S P s P e P \circ C \ddot{H} P r C - C \ddot{t}	B	PkP μ C \neq P \circ C \ddot{t} C, Ps	PkP μ C \neq P \circ C \ddot{t} C, Ps (C \ddot{t} C, P s P j C \ddot{t} C \neq P \ddot{e} C \ddot{t} P \gg C - P \cdot P \circ P i P s C \ddot{t} C, C \ddot{H} P μ P S P S C \ddot{t} C \ddot{t} C, P μ P S P s P e P \circ C \ddot{H} P r C - C \ddot{t})
	P μ C \ddot{H} P \ddot{e} C, P j C - C - (C \ddot{t} C, P s P j C \ddot{t} C \neq P \ddot{e} C \ddot{t} P \gg C - P \pm C \ddot{H} P \circ P r P \ddot{e} P e P \circ C \ddot{H} P r C - C \ddot{t} , C \ddot{e} P \gg C \ddot{t} P S P s C \neq P e P s P i P \circ C, P \circ C...C - P e P \circ C \ddot{H} P r C - C \ddot{t} , P j P \ddot{e} P i P s C, P \gg P \ddot{e} P i P \circ P \circ C \ddot{H} P \ddot{e} C, P j C - C \ddot{t})	B	B	PkP μ C \neq P \circ C \ddot{t} C, Ps
	P i P S C, P \circ C \ddot{H} P e C, P j C - P e P s P \circ C \ddot{H} P r P \circ	B	B	P \circ C \neq P μ P μ C $\ddot{H} C - P r P e P s$
	P μ PsC \ddot{t} P \ddot{e} P \gg P μ P S P μ C \ddot{t} P μ C \ddot{H} C \neq P μ P \ddot{e} C, C, C \ddot{t}	PkP μ C \neq P \circ C \ddot{t} C, Ps	B	P \ddot{S} P \circ C \ddot{t} C, Ps
	P \ddot{Y} P \circ C...C - P e P \circ C \ddot{H} P r C - C \ddot{t}	PkP μ C \neq P \circ C \ddot{t} C, Ps	B	B
P - P \pm PsPeC \ddot{e} C \ddot{t} C \ddot{e} P r P \ddot{e} PS	P μ C \ddot{H} C, P μ C \ddot{H} C - P \circ P \gg C \ddot{H} P S P \circ P i C - P i P s C, P μ P S P \circ C - C \ddot{t}	PkP μ C \neq P \circ C \ddot{t} C, Ps	P C - P r P e P s	PkP μ C \neq P \circ C \ddot{t} C, Ps
	P μ C \ddot{H} C, P s C \ddot{t} C, P \circ C, P \ddot{e} C \neq P S P \circ P i C - P i P s C, P μ P S P \circ C - C \ddot{t}	PkP μ C \neq P \circ C \ddot{t} C, Ps	B	B
	P μ C \ddot{H} P \ddot{e} P i P \gg P \ddot{e} P i P \ddot{e}	P C - P r P e P s	B	P \ddot{S} P \circ C \ddot{t} C, Ps
	P \circ P \circ C \ddot{t} P e C \ddot{t} P \gg C - C,	B	B	P \circ C \neq P μ P μ C $\ddot{H} C - P r P e P s$
P - P \pm PsPeC \ddot{e} PsC \ddot{H} P i P \circ P S C - P i P r P \ddot{e} C...P \circ P S P S C \ddot{t}	P \circ C \ddot{H} P s P S C...C - C, P μ P \circ C \ddot{e} P μ P \gg C \ddot{H}	B	P \ddot{S} P \circ C \ddot{t} C, Ps	B
	P \circ P \ddot{e} C \ddot{t} P i P S P s P μ	PkP μ C \neq P \circ C \ddot{t} C, Ps	P \ddot{S} P \circ C \ddot{t} C, Ps	PkP μ C \neq P \circ C \ddot{t} C, Ps
	P μ P \circ C \ddot{H} P \ddot{e} P S P i C - C,	B	P \ddot{S} P \circ C \ddot{t} C, Ps	B
	P P \ddot{e} P S C - C,	B	P \ddot{S} P \circ C \ddot{t} C, Ps	PkP μ C \neq P \circ C \ddot{t} C, Ps

P— P±PsPeCž C, CĤP° PIPSPc— CĤP° CĤC, PμPjPē	PhP±PrPsPjC—PSP°P>CH̄PSPēPNe P±C—P>CH̄	B	PSP° CĤC, Ps	PSP° CĤC, Ps
	PuPsCĤCfCēPμPSPSCĬ C, CfPSPeCĤC— PePēCēPμC±PSPēPeCf (Cf C, PsPjCf C±PēCĤP°>C— P·P° PIPsCĤ C, P° PrC—P° CĤPμCĬ)	B	B	PSP° CĤC, Ps
	P—P° PIPsCĤ	PkPμC±P° CĤC, Ps	B	B
	P° C—P° CĤPμCĬ	PkPμC±P° CĤC, Ps	PSP° CĤC, Ps	B
	PŸCfC...C—CĤC, CH̄ PĬ CĤPSc, C—	PkPμC±P° CĤC, Ps	B	PkPμC±P° CĤC, Ps
	P° PēCĤPμPμPĬCĤC—CĬ	PkPμC±P° CĤC, Ps	PSP° CĤC, Ps	PSP° CĤC, Ps
	P° P° CĤC, CĤPēC,	B	B	P° CĤPμPμ CĤC—PrPePs
	P° P° CĤC, CĤPScPμPSC, PμCĤPēC,	B	PSP° CĤC, Ps	B
	P° C—PμCĤPμP° P° P·C—CĬ CĬCĤPμPS	B	B	P° CĤPμPμ CĤC—PrPePs
	PkCfPrPsC, P°	PkPμC±P° CĤC, Ps	PSP° CĤC, Ps	PSP° CĤC, Ps
	PμP° PSPeCĤPμP° C, PēC,	B	B	P° CĤPμPμ CĤC—PrPePs
	P° C—P>CH̄ Cf PIPμCĤC...PSC—PNe C±P° CĤC, PēPSC— PĤPēPIPsC, P°	PkPμC±P° CĤC, Ps	B	B
	P° P>CĤPμP° PSPSCĬ	PkPμC±P° CĤC, Ps	PkPμC±P° CĤC, Ps	PkPμC±P° CĤC, Ps
PŸPĬCĤCf—PIPsPrC—P±PSP° PμPSC, PμCĤPScPμP° C, C—CĬ (PrPēPĬ. CĤPScP·PrC—P> B>PhCĤPScP±P>PēPIPsCĤC, C— P·P° CĤC, PsCĤCfPμP° PSPSCĬB>)	B	P° CĤPμPμ CĤC—PrPePs	B	
PhPSPiC—PsPSPμPĬCĤPSc, PēC±PSPēPNe PSP° P±CĤCĬPe PePēCēPμC±PSPēPeCf		P C—PrPePs		
P— P±PsPeCž PμPμC±C—PSPePē C, P° PĤPSPiC±PsPĤPĬC—PrPSPēC... CēP>CĬC...C—PĬ	PuC—PrPĤPēC° PμPSPSCĬ CĤC—PĬPSCĬ C, PμCĤPμPμPSC, C—PĬ PIPμC±C—PSPePē	B	PSP° CĤC, Ps	P° CĤPμPμ CĤC—PrPePs (Cf P±C—P>CH̄CēPScĤC, C— PSP° C, P>C— C...PsP>PμCĤC, P° P·Cf)
	P° PμPĬP° C, PēC,	B		P° CĤPμPμ CĤC—PrPePs
	P—PsPIC, CĬPSPēCĤCĬ	B		P° CĤPμPμ CĤC—PrPePs
	PhCfC, PsC—PjCfPSPSPēPNe PμPμPĬP° C, PēC, *		PkPμPIC—PrPsPjPs	
P— P±PsPeCž CēPeC—CĤPē C, P° C—C— PĬPsC...C—PrPSPēC...	PhP>PsPμPμCĤC—CĬ	B		PkPμC±P° CĤC, Ps
	PhPSPiC—PsPSPμPĬCĤPSc, PēC±PSPēPNe PSP° P±CĤCĬPe	B	P C—PrPePs	P° CĤPμPμ CĤC—PrPePs
	PhP>PμCĤPμC—C±PSPēPNe PrPμCĤPμP° C, PēC,	B	PkPμC±P° CĤC, Ps	B
	PμPSP>C—PjPScCĤC, PSP° PμCĤPēC, PμPjP°	B	B	P° CĤPμPμ CĤC—PrPePs
	P·PeP·P° PSC, PμPjP°	B	PkPμC±P° CĤC, Ps	PkPμC±P° CĤC, Ps
	P·PeCĤC, PsP>C—P° C, PēPĬPSPēPNe PrPμCĤPμP° C, PēC,	B	B	P° CĤPμPμ CĤC—PrPePs
	PuC—PrPĤPēC° PμPSPμ PIPsC, PsPĬPēPrC—P>PμPSPSCĬ	B	B	PkPμC±P° CĤC, Ps
	P±PsC, PsCĤPμPSCĤPēP±C—P>C—P·P° CĤC—CĬ	B	B	P° CĤPμPμ CĤC—PrPePs
	PŸPĬPμCĤP±C—PĤ	B	PkPμC±P° CĤC, Ps	PkPμC±P° CĤC, Ps
	P° PμPjPScCĤP° PĬC—C±PSC— PĬPēCĤPēPμP° PSPSCĬ	B	B	PkPμC±P° CĤC, Ps
	PkP° P±CĤCĬPe PμPĬC—PSPePμ	B	B	P° CĤPμPμ CĤC—PrPePs
	P° PēCĤPēPμP° PSPSCĬ	PkPμC±P° CĤC, Ps	PkPμC±P° CĤC, Ps	PkPμC±P° CĤC, Ps
	P—PjC—PSP° PePsP>CH̄PScCĤCf CēPeC—CĤPē	B	B	PkPμC±P° CĤC, Ps
	PŸPēPSPrCĤPScPj PŸC, C—PĬPμPSCĤP° — P° PĤPSPSCĤPSPSP°	B	B	P° CĤPμPμ CĤC—PrPePs
	PŸPScPēCĤPēC±PSPēPNe	B	B	PSP° CĤC, PsC, P° PSPμPĬC—PrPsPjP°
	PμPĬC—PrPμCĤPμP° P>CH̄PSPēPNe PSPμPeCĤPScP>C—P· PμCĤPScPĬPēPNe CĬPSPeP°	B	P C—PrPePs	PkPμC±P° CĤC, Ps
	P— P±PsPeCž PeC—CĤC, PePsPĬS- PjēPμ CĬP·PsPĬPsC— CĤPēCĤC, PμPjPē C, P° PjPSP>PsC±PSPēC... P·P° P>PsP·	PkP° P±CĤCĬPe PIPsPjC—P>PsPe	B	B
PhCĤC, CĤP° P>PĬC—CĬ		B	B	PkPμC±P° CĤC, Ps
PhCĤC, CĤPēC,		B	PSP° CĤC, Ps	B
P° C—P>CH̄ Cf CĤPĬPēPSC—		PkPμC±P° CĤC, Ps	PSP° CĤC, Ps	PkPμC±P° CĤC, Ps
PŸPĬP° P·Pj PjēPμ CĬP·C—PĬ		PkPμC±P° CĤC, Ps	P C—PrPePs	PSP° CĤC, Ps
PμC—P° P>PĬC—CĬ		B	PkPμC±P° CĤC, Ps	PkPμC±P° CĤC, Ps
P° C—P>CH̄ Cf PeC—PSCĤC—PĬPeP° C...		PkPμC±P° CĤC, Ps	B	B
P° C—P>CH̄ Cf PeC—CĤC, PēP° C...		B	PSP° CĤC, Ps	B
P—P±C—P>CH̄CēPμPSPSCĬ C±P° CĤC, PsC, Pē CĤPμC±PsPĬPēPĬCĤPēP° PSPSCĬ		B	B	PkPμC±P° CĤC, Ps
PuPsCĤCfCēPμPSPSCĬ		B	B	PkPμC±P° CĤC, Ps
CĤPμC±PsPĬPēPrC—P>PμPSPSCĬ	B	B	B	
PkC—PeC, CfCĤC—CĬ	B	B	PkPμC±P° CĤC, Ps	
PμPSP>P° PeC—CfCĤC—CĬ	PkPμC±P° CĤC, Ps	B	B	
PkPēCĤPēPScPĬP° PSPμPrPsCĤC, P° C, PSC—CĤC, CH̄	B	P C—PrPePs	B	
PĤPSC, PμPeCĤC—C— CĤPμC±PsPĬPēPĬC—PrPSPēC... CēP>CĬC...C—PĬ	B	PSP° CĤC, Ps	B	
P— P±PsPeCž CĤC, P° C, PμPĬPsC— CĤPēCĤC, PμPjPē C, P° PjPSP>PsC±PSPēC... P·P° P>PsP·	P·CĤPμPeC, PēP>CH̄PSP° PrPēCĤC, CfPSPeCĤC—CĬ/ C—PjPĬPsC, PμPSCĤC—CĬ	PkPμC±P° CĤC, Ps	B	PkPμC±P° CĤC, Ps
	P° C—PSPμPePsPjP° CĤC, C—CĬ	B	B	PkPμC±P° CĤC, Ps
	PhCĤC, PμPSC—CĬ	PkPμC±P° CĤC, Ps	PkPμC±P° CĤC, Ps	PSP° CĤC, Ps
	P° C—P>CH̄ Cf PrC—P>CĬPSCĤC— PĬCĤCfPrPSPSc— PeP>C—C, PePē	B	PSP° CĤC, Ps	PkPμC±P° CĤC, Ps
	PkP° P±CĤCĬPe PsP±P>PēC±C±CĬ	P C—PrPePs	PkPμC±P° CĤC, Ps	B
	P° C, PsPjP>CĤPĬP° PSC—CĤC, CH̄	PSP° CĤC, Ps	PSP° CĤC, Ps	PSP° CĤC, Ps
	P° CĤPēPĬPsPĬPsPrC—P±PSC— CĤC, P° PSPē	B	PSP° CĤC, Ps	B
	PŸPSPSP>PēPĬC—CĤC, CH̄	B	P C—PrPePs	B
	PkPμP·PrCfPμP° PSPSCĬ	B	PkPμC±P° CĤC, Ps	PkPμC±P° CĤC, Ps
	PkP° P±CĤCĬPe	PSP° CĤC, Ps	B	P° CĤPμPμ CĤC—PrPePs, Ps
	P° C—P>CH̄	B	PSP° CĤC, Ps	PkPμC±P° CĤC, Ps
	PμPμCĤPēC, PμCĤPēC±PSPēPNe PSP° P±CĤCĬPe	PSP° CĤC, Ps	PSP° CĤC, Ps	B
	PkP° P±CĤCĬPe PjēPμ CĬPēPēC... C, PēP° PSPēPSc	PSP° CĤC, Ps	B	B

<i>P PμP-CáP»CHC,P°C,Pë PrPsPrP°C,PePsPIPëC... PrPsCÍP»C-PrP¶PμPSCĤ</i>	<i>PμC-PrPIPëC%PμPSPSCĬ CĤC-PIPSCĬ PeCĤPμP°C,PëPSC-PSCf Cf PeCĤPsPIC-</i>	<i>PkPμC†P°CfC,Ps</i>	<i>P C-PrPePs</i>	<i>B</i>
	<i>PμC-PrPIPëC%PμPSPSCĬ CĤC-PIPSCĬ PeCĤPμP°C,PëPSC-PSC,,PsCfC,,PsPeC-PSP°P·Pë Cf PeCĤPsPIC-</i>	<i>B</i>	<i>PšP°CfC,Ps</i>	<i>B</i>
	<i>P—PSPë¶PμPSPSCĬ CĤC-PIPSCĬ PeP°P»C-CĤ Cf PeCĤPsPIC-</i>	<i>PkPμC†P°CfC,Ps</i>	<i>B</i>	<i>B</i>
	<i>PμC-PrPIPëC%PμPSPSCĬ CĤC-PIPSCĬ CfPμC†PsPIPëPSPë Cf PeCĤPsPIC-</i>	<i>B</i>	<i>PšP°CfC,Ps</i>	<i>B</i>
	<i>PμC-PrPIPëC%PμPSPSCĬ CĤC-PIPSCĬ CfPμC†PsPIPëC— PePëCfP»PsC,Pë Cf PeCĤPsPIC-</i>	<i>PkPμC†P°CfC,Ps</i>	<i>B</i>	<i>B</i>
	<i>PμC-PrPIPëC%PμPSPSCĬ CĤC-PIPSCĬ PIP°PjPjP°· PiP°CfC,P°PjC-P»C,CĤP°PSCfC,,PμCĤP°P·Pë Cf PeCĤPsPIC-</i>	<i>PkPμC†P°CfC,Ps</i>	<i>B</i>	<i>B</i>
	<i>P—PjPμPSCëPμPSPSCĬ PjP°CfPë C,C-P»P°</i>	<i>B</i>	<i>B</i>	<i>PkPμC†P°CfC,Ps</i>
	<i>P—P±C-P»CĤCëPμPSPSCĬPj PjP°CfPë C,C-P»P°</i>	<i>B</i>	<i>B</i>	<i>PkPμC†P°CfC,Ps</i>

*Pj Pïc-CfP»CĬCĤPμC”CfC,CĤP°C†C-PMëPSPëPMë PìPμCĤC-PsPr PìPsPIC-PrPsPjP»CĬP»PsCfCĬ PìCĤPs PIPëPìP°PrPePë PIPëPSPëPePSPμPSPSCĬ P°CfC,PsC-PjCfPSPSPsPiPs PìPμPìP°C,PëC,Cf P· P»P°C,PμPSC,PSPëPj PìPμCĤC-PsPrPsPj PIC-Pr PeC-P»CĤPePsC... PjC-CfCĬC†C-PI PrPs CĤPsPeC-PI, C%Ps P±CfP»Pë PsP±PsCĤPsC, PSPëPjPë Pïc-CfP»CĬ PIC-PrPjC-PSPë P·P°CfC,PsCfCfPIP°PSPSCĬ PsP»PjPμCfP°CĤC,P°PSCf.

PμPsPIC-PrPsPjP»CĬP»PsCfCĬ PìCĤPs PrPμPeC-P»CĤPeP° PIPëPìP°PrPeC-PI CĤP°P±PrPsPjC-PsP»C-P·Cf, CĬPePëPMë P·P° C†P°CfPsPj CĤPsP·PIPëC,PeCf P±CfPI PìPsPìP°CĬP°PSPëPMë P· PìCĤPëPMëPsPjPsPj P±P»PsPeP°C,PsCĤC-PI CĤPμC†PμPìC,PsCĤC-PI P°PSPìC-PsC,PμPSP·PëPSCf II. Pj PìP°C†C-C”PSC,C-PI, CĬPeC- PìCĤPëPMëPjP°P»Pë P°PjP»PsPrPëPìC-PS, PìPsPIC-PrPsPjP»CĬP»PsCfCĬ PìCĤPs PrPμPeC-P»CĤPeP° PIPëPìP°PrPeC-PI CĤPsP·PIPëC,PeCf PμPeCfC,CĤP°PìC-CĤP°PjC-PrPSPsPiPs CfPëPSPPrCĤPsPjCf.

PμPsPIC-PrPsPjP»PμPSPSCĬ PìCĤPs PìPsP±C-C†PSC- CĤPμP°PeC†C-C—

PμPsPIC-PrPsPjP»PμPSPSCĬ PìCĤPs PìPsP±C-C†PSC- CĤPμP°PeC†C-C— Pïc-CfP»CĬ CĤPμC”CfC,CĤP°C†C-C— P»C-PeP°CĤCfCĤPePsPiPs P·P°CfPsP±Cf PjP°C” PIP°P¶P»PëPìPμ P·PSP°C†PμPSPSCĬ. PìPμ PrP°C” P·PjPsPìCf PìCĤPsPIPsPrPëC,Pë PjPsPSC-C,PsCĤPëPSPi CfPìC-PIPIC-PrPSPsCëPμPSPSCĬ PePsCĤPëCfC,CĤ/CĤPëP·PëPe PìCĤPë P·P°CfC,PsCfCfPIP°PSPSC- C†CĤPsPiPs P»C-PeP°CĤCfCĤPePsPiPs P·P°CfPsP±Cf. PĤPμPrPëC†PSPëPj C,P° C,,P°CĤPjP°C†PμPIC,PëC†PSPëPj PìCĤP°C†C-PIPSPëPeP°Pj, P° C,P°PePsP¶ PìP°C†C-C”PSC,P°Pj P°P±Ps C—C...PSC-Pj P·P°PePsPSPSPëPj PìCĤPμPrCfC,P°PIPSPëPeP°Pj CfP»C-Pr PìPsPIC-PrPsPjP»CĬC,Pë PìCĤPs CfCfC- PIPëPìP°PrPePë Pïc-PrPsP·CĤCĤPIP°PSPëC... PìPsP±C-C†PSPëC... CĤPμP°PeC†C-PMë C,P° PIC-PrCfCfC,PSPsCfC,C- PμC,,PμPeC,PëPIPSPsCfC,C- P»C-PeP°CĤCfCĤPePsPiPs P·P°CfPsP±Cf C†PμCĤPμP· P°PIC,PsPjP°C,PëP·PsPIP°PSCf C-PSC,,PsCĤPjP°C†C-PMëPSCf CfPëCfC,PμPjCf P· C,,P°CĤPjP°PePsPSP°PiP»CĬPTrCf P·P° PìPsCfPëP»P°PSPSCĬPj: <https://aisf.dec.gov.ua>.

PŷPμCĤPjC-PS PìCĤPëPrP°C,PSPsCfC,C-. 5 CĤPsPeC-PI.

PjPjPsPIPë P·P±PμCĤC-PiP°PSPSCĬ.

PkPμ PìPsC,CĤPμP±CfC” CfPìPμC†C-P°P»CĤPSPëC... CfPjPsPI P·P±PμCĤC-PiP°PSPSCĬ.

P—P±PμCĤC-PiP°C,Pë Cf PSPμPrPsCfC,CfPìPSPsPjCf PrP»CĬ PrC-C,PμPMë PjC-CfC†C-.

PJPiP°PePsPIPεP°.

14 C,P°P±P»PμC,PsPe, PIPεCḡPëC,PëC... PïP»C-PIPePsPIPsCḡ PsP±PsP»PsPSPePsCḡ, Cí P±P»C-CÍC,PμCḡC-; PïPs 2 P±P»C-CÍC,PμCḡPë Cí PeP°CḡC,PsPSPSC-PNë PïP°C‡C‡C-.

PльP°C,PμPiPsCḡC-Cḡ PIC-PrPïCíCÍPeCí. P—P° CḡPμC‡PμPïC,PsPj.

P’PëCḡPsP±PSPëPe.

P”P°C-C‡C- PŸP°PSPεC-Ps P@CḡPsCíPï P“PjP±Pí.

PњC-CÍC‡PμP·PSP°C...PsPrP¶PμPSPSCḡ PIPëCḡPsP±PSPëPeC-PI C,P° P°PrCḡPμCÍP° PjC-CÍC‡Cḡ PïCḡPsPIP°PrP¶PμPSPSCḡ C—C... PrC-CḡP»CњPSPsCÍC,C-.

P>CíC-C,PïPsP»CњPrC€C,CḡP°CÍCÍPμ 1, PμC„P°C„C„PμPSC...PsC„PμPS Pњ.Pr.C-P»Pj, P’P°PIP°CḡC-Cḡ, 85276, PќC-PjPμC‡C‡PëPSP°.