

Р†РќРЎРЎР РЈРЉРІР†РЇ В

$$\begin{aligned} & \text{PrP} \gg \text{CII} \text{ PjP}_{\mu} \text{PrP}^{\text{C}} \pm \text{PSPsPiPs} \text{ P} \cdot \text{P}^{\circ} \text{Cf}^{\circ} \text{C}, \text{PsCf}^{\circ} \text{Cr}^{\circ} \text{PIP}^{\circ} \text{PSPSCII} \text{ P} \gg \text{C-PeP}^{\circ} \text{Cf}^{\circ} \text{CHbPePsPiPs} \\ & \text{P} \cdot \text{P}^{\circ} \text{Cf}^{\circ} \text{PsP} \pm \text{Cr}^{\circ} \end{aligned}$$

PḥPḱPḥP Pḥ P•P>P>P†PḡPŷPḥ

(ANORO ELLIPTA)

$P \dot{\vee} P \in P \gg P^\circ P \Gamma$:

$$\text{PrC}-\text{CHC}\equiv\text{C}-\text{CH}_2\text{PuC}\equiv\text{PsPIP}\ddot{\text{P}}\text{SP}\ddot{\text{P}}\text{: CrPjPuPeP}\gg\text{C}-\text{PrP}\ddot{\text{P}}\text{PSC}-\text{CrPj}, \text{PIC}-\text{P}\gg\text{P}^\circ\text{PSC}, \text{PuCH}_2\text{PsP}\gg;$$

1 PrPsP·P°, C%Ps PrPsCf'C,P°PIP»CfC"C,CfCf'Cl, PjC-Cf'C,PëC,CH 55 PjPePi
CfPjPpPeP»C-PrPëPSC-CfPjCr (Cr C„PsCfPjC- P±CfPjPjC-PrCr) C- 22 PjPePi
PIC-P»P°PSC,PuCfPjPsP»Cr (Cr C„PsCfPjC- C,CfPëC„PuPSP°C,P°C,Cr);

CtPm PIC-PrPiPsPIC-PrP°C" PiPsPiPmCTPmPrPSCbPs CTjPsP·PiPsPrC-P»PmPSC-PN₆ PrPsP·C- 74,2
 PjPePi CrPjPmPeP»C-PrPëPSC-CrPjCr P±CTjPsPjC-PrCr C- 25 PjPePi PIC-P»P°PSC,PmCTjPsP»Cr
 (Cr C_mPsCTjPjC- C,CTjPëC_mPmPSP°C,P°C,Cr);

$PrPsPiPsPjC-P\eta PSC-C\eta P\mu C\ddagger PsPIP\ddot{e}PSP\ddot{e}$: $P\gg P^{\circ}PeC, PsP\cdot P^{\circ}$, $PjPsPSPsPiC-PrC\eta P^{\circ}C,;$
 $PjP^{\circ}PiPSC-C\eta C\dot{\Gamma}C, P\mu P^{\circ}C\eta P^{\circ}C,.$

**Р>С-РєР°СЃСЃСЃРєР°СЃ С,,РѕСЃРјРјР°. РѓРѕСЃРјРѕСЃРєРєРѕРє РїР»»СЃСЃ
С-РѕРїР°Р°Р»»СЃСЃСЃСЃСЃСЃСЃ. В**

PhC1PSPsPIPSC- C,,C-P.PëPePs-C...C-PjC-C#PSC- PIP»P°C1C,PëPIPsC1C,C-:
PïP»P°C1C,PëPePsPIPëPN C-PSPiP°P»C1C,PsC1 P·C- C1PIC-C,P»Ps-C1C-C1PëPj
PePsC1PïC1C1PsPj, C#PμC1PIPsPSPëPj PePsPIPïP°C#PePsPj C1PsP·PïPëP»C1PIP°C#P° C,P°
P»C-C#PëP»CHbPSPëPePsPj PrPsP·, C1PiP°PePsPIP°PSPëPN PI P»PsC,PsPe P· C,,PsP»CHbPiPë,
C%Ps PjC-C1C,PëC,CH PïP°PePμC,PëPe P· PIPsP»PsPiPsPïPsPiP»PëPSP°C#PμPj. P·PsC,PsPe
P·P°PïP°C1PSPëPN PIC-PrC1PëPIPSPsC1 PeC1PëCëPePsC1. P†PSPiP°P»C1C,PsC1
PjC-C1C,PëC,CH PrPIC- C1C,C1C-C#PePë PïPs 30 C1C-PIPSPsPjC-C1PSPs
C1PsP·PiPsPrC-P»PμPSPëC... P±P»C-C1C,PμC1C-PI, PePsP¶PμPS P· C1PePëC... PjC-C1C,PëC,CH
P±C-P»PëPN PïPsC1PsCëPsPe.

P_αP°C₇PjP°PePsC,PμC₇P°PiPμPIC,PëC±PSP° PiC₇CfPiP°. P>C-PeP°C₇CfCHPeC-
P·P°CfPsP±Pë, C%oPs P·P°CfC,PscfPsPICfCHC,CHCfCЦ PiC₇Pë
PsP±CfC,C₇CfPeC,PëPIPSPëC... P·P°C...PIPSc₇CHPIP°PSPSCЦC... PrPëC...P°P»CHHPSPëC...
CëP»CЦC...C-Pl. PhPrC₇PμPSPμC₇PiC-C±PSC- PiC₇PμPiP°C₇P°C,Pë PI PePsPjP±C-PSP°C†C-C—
P· P°PSC,PëC...PsP»C-PSPμC₇PiC-C±PSPëPjPë PiC₇PμPiP°C₇P°C,P°PjPë, PIPeP»CHC±P°C₇C±Pë
PiPsC,C₇C-PNëPSC- PePsPjP±C-PSP°C†C-C— P· PePsC₇C,PëPePsCfC,PμC₇PsC—PrP°PjPë.

P'C-P»P°PSC,PμCḡPsP» C,P° CṛPjPμPeP»C-PrPëPSC-CṛPjCṛ P±CḡPsPjC-Pr.

PḡPsPr PḡPṽPḡ R03A L03.

PαP°CḡPjP°PePsP»PsPiC-C≠PSC- PIP»P°CṛC,PëPIPsCṛC,C-

PαP°CḡPjP°PePsPrPëPSP°PjC-PeP°

PḡPμC...P°PSC-P·Pj PrC-C—

PjPjPμPeP»C-PrPëPSC-CṛPj/PIC-P»P°PSC,PμCḡPsP» C" PePsPjP±C-PSP°CṛC-C"Ḥ
C-PSPiP°P»CḶCṛC-PNḡPSPsPiPs P°PSC,P°PiPsPSC-CṛC,P° PjCṛCṛPeP°CḡPëPSPsPiPs
CḡPμCṛPμPīC,PsCḡP° C,CḡPëPIP°P»PsC— PrC-C— C,P° P±PμC,P°₂-
P°PrCḡPμPSPμCḡPiC-C≠PSPsPiPs P°PiPsPSC-CṛC,P° C,CḡPëPIP°P»PsC— PrC-C—. PḡC-CṛP»CḶ
PīPμCḡPsCḡP°P»CḡPSPsC— C-PSPiP°P»CḶCṛC-C— PsP±PëPrPIC- CṛPīPsP»CṛPePë
PrC-CḤC,Ḥ PjC-CṛCṛPμPIPs PSP° PrPëC...P°P»CḡPSC- CḡP»CḶC...Pë, PIPSP°CṛP»C-PrPsPe
C≠PsPiPs PIC-PrP±CṛPIP°C"ḤCḡCḶ CḡPsP·CḡPëCḡPμPSPSCḶ P±CḡPsPSC...C-PI Cṛ
CḡPμP·CṛP»CḡC,P°C,C- CḡC-P·PSPëC... PjPμC...P°PSC-P·PjC-PI PrC-C—.

PjPjPμPeP»C-PrPëPSC-CṛPj

PjPjPμPeP»C-PrPëPSC-CṛPj C" P°PSC,P°PiPsPSC-CṛC,PsPj PjCṛCṛPeP°CḡPëPSPsPiPëC...
CḡPμCṛPμPīC,PsCḡC-PI C,CḡPëPIP°P»PsC— PrC-C— (P°PSC,PëC...PsP»C-PSPμCḡPiC-C≠PSPëPNḡ
P·P°CṛC-P±). P'C-PS C" PīPsC...C-PrPSPëPj C...C-PSCṛPeP»C-PrPëPSCṛ P·
P°PeC,PëPIPSC-CṛC,Ḥ C%PsPrPs C≠PëCṛP»PμPSPSPëC... PīC-PrC,PëPīC-PI
PjCṛCṛPeP°CḡPëPSPsPiPëC... CḡPμCṛPμPīC,PsCḡC-PI. PjPjPμPeP»C-PrPëPSC-CṛPj
C≠PëPSPëC,Ḥ P±CḡPsPSC...PsCḡPsP·CḡPëCḡPīP°P»CḡPSCṛ PrC-CḤ CḡP»CḶC...PsPj
PePsPSPeCṛCḡPμPSC,PSPsPiPs PīCḡPëPiPSC-C≠PμPSPSCḶ P·Pīḡ™ CḶP·CṛPIP°PSPSCḶ
P°CṛPμC,PëP»C...PsP»C-PSCṛ P· PjCṛCṛPeP°CḡPëPSPsPiPëPjPë CḡPμCṛPμPīC,PsCḡP°PjPë
PīP»P°PrPePëC... Pjḡ™ CḶP·C-PI PrPëC...P°P»CḡPSPëC... CḡP»CḶC...C-PI. P'PëCḶPIP»PμPSP°
PīPsPIC-P»CḡPSP° PsP±PsCḡPsC,PSC-CṛC,Ḥ PSP° PīC-PrC,PëPīC- Pḡ3
PjCṛCṛPeP°CḡPëPSPsPiPëC... CḡPμCṛPμPīC,PsCḡC-PI P»CḡPrPëPSPë *in vitro* C- C,CḡPëPIP°P»P°
PrC-CḶ *in vivo* PīCḡPë P±PμP·PīPsCṛPμCḡPμPrPSCḡPsPjCṛ PīPIPμPrPμPSPSC- PI P»PμPiPμPSC-
PI PrPsPeP»C-PSC-C≠PSPëC... PjPsPrPμP»CḶC....

P'C-P»P°PSC,PμCḡPsP»

P'C-P»P°PSC,PμCḡPsP» C" CṛPμP»PμPeC,PëPIPSPëPj P°PiPsPSC-CṛC,PsPj P±PμC,P°₂-
P°PrCḡPμPSPμCḡPiC-C≠PSPëC... CḡPμCṛPμPīC,PsCḡC-PI C,CḡPëPIP°P»PsC— PrC-C—
(P±PμC,P°₂-P°PrCḡPμPSPμCḡPiC-C≠PSPëPNḡ P°PiPsPSC-CṛC,).
PαP°CḡPjP°PePsP»PsPiC-C≠PSPëPNḡ PIPīP»PëPI P±PμC,P°₂-P°PrCḡPμPSPμCḡPiC-C≠PSPëC...
P°PiPsPSC-CṛC,C-PI, PIPeP»CḡCṛP°CḡCṛPë PIC-P»P°PSC,PμCḡPsP», PīCḡPëPSP°PNḡPjPSC-
C≠P°CṛC,PePsPiPs PīPsCḶCṛPSCḤ"C,ḤCḡCḶ CṛC,PëPjCṛP»CḶCṛC-C"Ḥ
PIPSCṛC,CḡC-CḡPSCḡPsPeP»C-C,PëPSPSPsC— P°PrPμPSC-P»P°C,CṛPëPeP»P°P·Pë,
C,,PμCḡPjPμPSC,Cṛ, CḶPePëPNḡ PeP°C,P°P»C-P·CṛC" PīPμCḡPμC,PIPsCḡPμPSPSCḶ
P°PrPμPSPsP·PëPSC,CḡPëC,,PsCṛC,,P°C,Cṛ (PḡPṽPα) PSP° CṛPëPeP»C-C≠PSPëPNḡ-3',5'-
P°PrPμPSPsP·PëPSPjPsPSPsC,,PsCṛC,,P°C, (CṛAMPα). PḡC-PrPIPëC%PμPSC- CḡC-PIPSC-
CṛPḡPḡPα PIPëPeP»PëPeP°CḡC,Ḥ CḡPsP·CṛP»P°P±P»PμPSPSCḶ PīP»P°PrPePëC...
Pjḡ™ CḶP·C-PI P±CḡPsPSC...C-PI C- PīCḡPëPiPSC-CṛCḡC,Ḥ PIPëPIC-P»CḡPSPμPSPSCḶ
PjPμPrC-P°C,PsCḡC-PI PīC-PīPμCḡCṛC,P»PëPIPsCṛC,C- PSPμPiP°PNḡPSPsPiPs C,PëPīCṛ P·

PeP»C-C,PëPS, PsCÍPsP±P»PëPIPs P· PsPíP°CÍPëCÍC,PëC... PeP»C-C,PëPS.

PαP°CḡPjP°PePsPrPëPSP°PjC-C±PSC- PμC,,PμPeC,Pë

Pj 6-PjC-CÍCḡC±PSPëC... PrPsCÍP»C-PrPḡPμPSPSCḡC... P†P†P† C,,P°P·Pë
PePsPjP±C-PSP°C†C-Cḡ C†PjPμPeP»C-PrPëPSC-C†Pj/PIC-P»P°PSC,PμCḡPsP»
PíCḡPsPrPμPjPsPSCÍC,CḡC†PíP°P»P° PeP»C-PSC-C±PSPs P·PSP°C±C†C%Pμ
PíPsP»C-PíC€PμPSPSCḡ C,,C†PSPeC†C-C— P»PμPíPμPSCḡ PíPsCḡC-PIPSCḡPSPs P·
PíP»P°C†PμP±Ps (PIPëPjC-CḡCḡPíP°PíCÍCḡ PsP±Bḡ™ C”Pj C,,PsCḡCÍPsPíP°PSPsPíPs
PíPëPrPëC...C† P·P° PíPμCḡC€C† C†PμPeC†PSP†C† [PhPαP’₁]) PíCḡPsC,CḡPíPsPj 24 PíPsPrPëPS
PíC-CÍP»Cḡ PíCḡPëPNëPsPjC† PsPrPëPS CḡP°P· PSP° PrPμPSCḡ, C%Ps P±C†P»Ps
P·P°C,,C-PeCÍPsPíP°PSPs C±PμCḡPμP· 15 C...PIPëP»PëPS PíC-CÍP»Cḡ
P·P°CÍC,PscÍC†PíP°PSPSCḡ PíPμCḡC€PsC— PrPsP·Pë (PíPsP»C-PíC€PμPSPSCḡ
PíPsCḡC-PIPSCḡPSPs P· PíP»P°C†PμP±Ps CÍC,P°PSPsPíPëP»Ps 112 PjP» (Cḡ < 0,001*).
PḡP°PeCÍPëPjP°P»CḡPSPμ PíPsPeCḡP°C%PμPSPSCḡ PhPαP’₁ PíCḡPsC,CḡPíPsPj
PíPμCḡC€PëC... 6 PíPsPrPëPS PíC-CÍP»Cḡ PíCḡPëPNëPsPjC† PíPsCḡC-PIPSCḡPSPs P·
PíP»P°C†PμP±Ps Pí C†PμCḡPμPrPSCḡPsPjC† CÍC,P°PSPsPíPëP»Ps 224 PjP» (p < 0,001*)
PSP° 24 C,PëPḡPSC- PrPsCÍP»C-PrPḡPμPSPSCḡ. PkPμ PsC,CḡPëPjP°PSPs PrPsPeP°P·C-PI
CḡPsP·PíPëC,PeC† C,P°C...C-C,,C-P»P°PeCÍC-C— PIPSP°CÍP»C-PrPsPe PíCḡPëPNëPsPjC†
P»C-PeP°CḡCÍCḡPePsPíPs P·P°CÍPsP±C† PhPSPsCḡPs P·P»P»C-PíC,P° P·P° C±P°CÍ
PrPsCÍP»C-PrPḡPμPSCḡ.

PḡP°CḡPrC-PsPμP»PμPeC,CḡPsC,,C-P·C-PsP»PsPíC-Cḡ. P’PíP»PëPI
C†PjPμPeP»C-PrPëPSC-C†PjC†/PIC-P»P°PSC,PμCḡPsP»C† PSP° QT-C-PSC,PμCḡPíP°P»
PsC†C-PSCḡPíP°P»Pë C† PíP»P°C†PμP±Ps- C,P° PjPsPeCÍPëC,,P»PsPeCÍP°C†PëPS-
PePsPSC,CḡPsP»CḡPsPíP°PSPsPjC† PrPsCÍP»C-PrPḡPμPSPSC- P·P° C†C±P°CÍC,Cḡ 103
P·PrPsCḡPsPíPëC... PrPsP±CḡPsPíPsP»CḡC†C-PI, CḡPeC- PíCḡPsC,CḡPíPsPj 10 PrPSC-PI
PíCḡPëPNëPjP°P»Pë C†PjPμPeP»C-PrPëPSC-C†Pj/PIC-P»P°PSC,PμCḡPsP» PsPrPëPS CḡP°P· PSP°
PrPsP±C† C† PrPsP·C- 113/22 PjPePí P°P±Ps 500/100 PjPePí (PíPsPíPμCḡPμPrPSCḡPs
PIC-PrPjC-CḡCḡPSP° PrPsP·P° C†PjPμPeP»C-PrPëPSC-C†PjC† C† PIC-CÍC-Pj CḡP°P·C-PI
PíPëC%P° P·P° CḡPμPePsPjPμPSP†PsPíP°PSC† PrPsP·C†, P° PrPsP·P° PIC-P»P°PSC,PμCḡPsP»C†
C† C±PsC,PëCḡPë CḡP°P·Pë PíPëC%P° P·P° CḡPμPePsPjPμPSP†PsPíP°PSC† PrPsP·C†).
PḡP°PeCÍPëPjP°P»CḡPSP° C†PμCḡPμPrPSCḡ CḡC-P·PSPëC†Cḡ C† PíCḡPsP»PsPSPíP°C†C-C—
C-PSC,PμCḡPíP°P»C† QT (C†PePsCḡPëPíPsPíP°PSP° P·P° PjPμC,PsPrPsPj PαCḡPμPrPμCḡC-PeP°,
QT_cF) PíPsCḡC-PIPSCḡPSPs P· PíP»P°C†PμP±Ps PíC-CÍP»Cḡ PíPsPíCḡP°PíPëPë PSP°
PíPëC...C-PrPSPëPNë CḡC-PIPμPSCḡ CÍC,P°PSPsPíPëP»P° 4,3 (90 % P”P† [PrPsPIC-CḡC±PëPNë
C-PSC,PμCḡPíP°P»] 2,2Bḡ”6,4) PjC-P»C-CÍPμPeC†PSP†Pë C±PμCḡPμP· 10 C...PIPëP»PëPS
PíC-CÍP»Cḡ C-PSPíP°P»CḡC†C-C—
C†PjPμPeP»C-PrPëPSC-C†PjC†/PIC-P»P°PSC,PμCḡPsP»C† C† PrPsP·C- 113/22 PjPePí C,P°
8,2 (90 % P”P† 6,2Bḡ”10,2) PjC-P»C-CÍPμPeC†PSP†Pë C±PμCḡPμP· 30 C...PIPëP»PëPS
PíC-CÍP»Cḡ C-PSPíP°P»CḡC†C-C— C†PjPμPeP»C-PrPëPSC-C†PjC†/PIC-P»P°PSC,PμCḡPsP»C† C†
PrPsP·C- 500/100 PjPePí. PhC,PḡPμ, PíCḡPë C-PSPíP°P»CḡC†C-C—
C†PjPμPeP»C-PrPëPSC-C†PjC†/PIC-P»P°PSC,PμCḡPsP»C† C† PrPsP·C- 113/22 PjPePí
PḡPsPrPSPsPíPs PeP»C-PSC-C±PSPs P·PSP°C±C†C%PsPíPs PíCḡPsP°CḡPëC,PjPsPíPμPSPSPsPíPs
PμC,,PμPeC,C†, PíPsPíBḡ™ CḡP·P°PSPsPíPs C-P· PíCḡPsP»PsPSPíP°C†C-C”Cḡ
C-PSC,PμCḡPíP°P»C† QT, PSPμ PíPëPSPëPeP°P»Ps.

PŸPíPsCÍC,PμCḡC-PíP°P»PsCÍCḡ C,P°PePsPḡ PrPsP·PsP·P°P»PμPḡPSPμ
P·P±C-P»CḡC€PμPSPSCḡ C±P°CÍC,Psc,Pë C†PμCḡC†PμPíPëC... C†PePsCḡPsC±PμPSCḡ.
PḡP°PeCÍPëPjP°P»CḡPSP° C†PμCḡPμPrPSCḡ CḡC-P·PSPëC†Cḡ C±P°CÍC,Psc,Pë

PJ PrPμPeC-P»CHPePsC... PrPsCfP»C-PrP¶PμPSPSCfC... P»C-PeP°CfCfCHPePePN₂ P·P°CfC-P± PhPSPsCfPs P·P»P»C-PiC,P° PiCfPsPrPμPjPsPSCfC,CfCfPIP°PI PiPsP»C-PiC¶PμPSPSCf C,,CrPSPeCfC-C— P»PμPiPμPSCf (C%Ps PsCfC-PSCfPIP°P»PsCfCf P·P° P·PjC-PPSPsCf PhPαP'₁ PIC-Pr PiPsCfP°C,PePsPiPsPiPs CfC-PIPSCf). P' PsPrPSPsPjCr 6-PjC-CfCfCfPSPsPjCr PrPsCfP»C-PrP¶PμPSPSC- P†P†P† C,,P°P·Pë Cr PiCfCfPiC- P·P°CfC,PsCfCfPIP°PSPSCf P»C-PeP°CfCfCHPePsPiPs P·P°CfPsP±Cr PhPSPsCfPs P·P»P»C-PiC,P° PiCfPsPrPμPjPsPSCfC,CfPsPIP°PSPs CfC,P°C,PëCfC,PëCfPSPs P·PSP°CfCrC%Pμ PiPsP»C-PiC¶PμPSPSCf PhPαP'₁ (PiPμCfPIPëPSPSP° PeC-PSC†PμPIP° C,PsCfPeP°) PSP° 24-PjCr C,PëP¶PSC- PiPsCfC-PIPSCfPSPs P· PiCfCfPiPsCf PiP»P°C†PμP±Ps C,P° PePsP¶PSPsCf PiCfCfPiPsCf PjPsPSPsC,PμCfP°PiC-C—. PñCfC-Pj C,PsPiPs, P»C-PeP°CfCfCHPePePN₂ P·P°CfC-P± PhPSPsCfPs P·P»P»C-PiC,P° PiCfPsPrPμPjPsPSCfC,CfCfPIP°PI PeP»C-PSC-CfPSPs P·PSP°CfCrC%PëPN₂ C,P° CfC,P°C,PëCfC,PëCfPSPs PrPsCfC,PsPIC-CfPSPëPN₂ PiCfPëCfC-CfC, PhPαP'₁ PiPsCfC-PIPSCfPSPs P· C,C-PsC,CfPsPiC-C" Pj Cr PrPiPsC... C-P·

PŠP°CÍC,PeP° PíP°C†C-C”PSC,C-PI, Cr CĬPePëC... PíPsPeP°P·PSPëPe SGRQ P·PSPëP·PëPICÍCĬ
 PíCĤPëPSP°P№PjPSC- PSP° 4 PsPrPëPSPëC†C- (4 PsPrPëPSPëC†C- бĤ” MPњP—P PrP»CĬ
 PsPíPëC,CrPIP°P»CHPSPëPeP° SGRQ) PSP° 24-PjCr C,PëP¶PSC-, P±CrP»P° P±C-P»CHC€P° Cr
 PíCĤCrPíC- P»C-PeP°CĤCÍCHPePsPíPs P·P°CÍPsP±Cr PhPSPsCĤPs P·P»P»C-PíC,P° (49 %)

PíPsCĤC-PIPSCĬPSPs P· PíP»P°C†PµP±Ps (34 %) C,P° PjPsPSPsC,PµCĤP°PíC-C”CĤ (44 % PrP»CĬ
 CrPjPµPeP»C-PrPëPSC-CrPjCr C,P° 48 % PrP»CĬ PIC-P»P°PSC,PµCĤPsP»Cr). P’ PsPrPSPsPjCr
 PrPsCÍP»C-PrP¶PµPSPSC- P· P°PeC,PëPIPSPëPj PíCĤPµPíP°CĤP°C,PsPj PíPsCĤC-PIPSCĬPSPSCĬ
 P±C-P»CHC€PëP№ PIC-PrCÍPsC,PsPe PíP°C†C-C”PSC,C-PI, CĬPeC- PsC,CĤPëPjCrPIP°P»Pë
 P»C-PeP°CĤCÍCHPePëP№ P·P°CÍC-P± PhPSPsCĤPs P·P»P»C-PíC,P°, PIC-PrPíPsPIC-PI
 PeP»C-PSC-C‡PSPs P·PSP°C‡CrC‰PëPj PíPsP»C-PíC€PµPSPSCĬPj PíPsPeP°P·PSPëPeP° SGRQ
 PSP° 24-PjCr C,PëP¶PSC- (53 %) PíPsCĤC-PIPSCĬPSPs P· PíP°C†C-C”PSC,P°PjPë, CĬPeC-
 PsC,CĤPëPjCrPIP°P»Pë C,C-PsC,CĤPsPíC-P№ (46 %). P’ C-PSC€PëC... PrPIPsC...

PrPsCfP»C-PrP¶PµPSPSC¶C... P· P°PeC, PēPIPSpēPj PīCḡPµPīP°CḡP°C, PsPj
PīPsCḡC-PIPSC¶PSPSC¶ P°PSP°P»PsPiC-C‡PSP° C‡P°CfC, PeP° PīP°C†C-C°PSC, C-PI
PrPsCfC¶PīP»P° PīCḡPēPSP°PnēPjPSC- PḡPḡP—P P· P»C-PeP°CḡCfCḡPePēPj P·P°CfPsP±PsPj
PḡPSPsCḡPs P·P»P»C-PīC, P° C, P° C, C-PsC, CḡPsPīC-C°Pj: 49 % C- 54 % Bḡ” P·
P»C-PeP°CḡCfCḡPePēPj P·P°CfPsP±PsPj PḡPSPsCḡPs P·P»P»C-PīC, P° Cḡ PrPsP·C- 55/22 PjPePi,
52 % C- 55 % Bḡ” P· C, C-PsC, CḡPsPīC-C°Pj.

P’PēPePsCḡPēCfC, P°PSPSC¶ PīCḡPµPīP°CḡP°C, C-PI PrP»C¶ PīPsP»PµPiCēPµPSPSC¶
CfPēPjPīC, PsPjC-PI

PµCḡPē P·P°CfC, PsCfCḡPīP°PSPSC- P»C-PeP°CḡCfCḡPePsPiPs P·P°CfPsP±Cḡ PḡPSPsCḡPs
P·P»P»C-PīC, P° P·PjPµPSCēPēP»PsCfC¶ PīPēPePsCḡPēCfC, P°PSPSC¶ PīCḡPµPīP°CḡP°C, C-PI
PrP»C¶ PīPsP»PµPiCēPµPSPSC¶ CfPēPjPīC, PsPjC-PI (CfP°P»CḡP±CḡC, P°PjPsP»)
PīCḡPsC, C¶PīPsPj 1Bḡ”24 C, PēP¶PSC-PI PrPsCfP»C-PrP¶PµPSPSC¶ PīPsCḡC-PIPSC¶PSPs P·
PīP»P°C†PµP±Ps C, P° CḡPjPµPeP»C-PrPēPSC-CḡPjPsPj (PrPēPI. C, P°P±P»PēC†Cḡ 1) C, P°
P±CḡP»Ps PīCḡPsPrPµPjPsPSCfC, CḡPsPīP°PSPs P·P±C-P»CḡCēPµPSPSC¶
PeC-P»CḡPePsCfC, C- PrPSC-PI, PePsP»Pē PSPµ PīPsC, CḡC-P±PSPs P±CḡP»Ps
P·P°CfC, PsCfCḡPīCḡPīP°C, Pē PīCḡPµPīP°CḡP°C, Pē PrP»C¶ PīPsP»PµPiCēPµPSPSC¶
CfPēPjPīC, PsPjC-PI (PI CfPµCḡPµPrPSCḡPsPjCḡ PSP° 11,1 %) PīPsCḡC-PIPSC¶PSPs P·
PīCḡCḡPīPsCḡ PīP»P°C†PµP±Ps, Cḡ C¶PeC-Pnē P·PjPµPSCēPēP»P°CfC¶ PeC-P»CḡPeC-CfC, Cḡ
C, P°PePēC... PrPSC-PI PI CfPµCḡPµPrPSCḡPsPjCḡ PSP° 0,9 %.

Pj C, CḡCḡPsC... 6-PjC-CfC¶C‡PSPēC... PrPsCfP»C-PrP¶PµPSPSC¶C...,
PePsPSC, CḡPsP»CḡPsPīP°PSPēC... P·P° P°PeC, PēPIPSpēPj PīCḡPµPīP°CḡP°C, PsPj
PīPsCḡC-PIPSC¶PSPSC¶, P·P°CfC, PsCfCḡPīP°PSPSC¶ P»C-PeP°CḡCfCḡPePsPiPs
P·P°CfPsP±Cḡ PḡPSPsCḡPs P·P»P»C-PīC, P° P·PjPµPSCēPēP»Ps PīPēPePsCḡPēCfC, P°PSPSC¶
PīCḡPµPīP°CḡP°C, C-PI PrP»C¶ PīPsP»PµPiCēPµPSPSC¶ CfPēPjPīC, PsPjC-PI
PīPsCḡC-PIPSC¶PSPs P· C, C-PsC, CḡPsPīC-C°Pj P·C- CfC, P°C, PēCfC, PēC‡PSPs
P·PSP°C‡CḡC%PēPjPē P·PSPēP¶PµPSPSC¶PjPē Cḡ PrPīPsC... PrPsCfP»C-PrP¶PµPSPSC¶C...
(PrPēPI. C, P°P±P»PēC†Cḡ 1). P·C-PeP°CḡCfCḡPePēPnē P·P°CfC-P± PḡPSPsCḡPs
P·P»P»C-PīC, P° C, P°PePsP¶ PīCḡPsPrPµPjPsPSCfC, CḡCḡPīP°PI P·P±C-P»CḡCēPµPSPSC¶
C‡P°CfC, PePē PrPSC-PI, PePsP»Pē PSPµ PīPsC, CḡC-P±PSPs P±CḡP»Ps
PīPēPePsCḡPēCfC, PsPīCḡPīP°C, Pē PīCḡPµPīP°CḡP°C, Pē PrP»C¶ PīPsP»PµPiCēPµPSPSC¶
CfPēPjPīC, PsPjC-PI, Cḡ PīCfC-C... C, CḡCḡPsC... PrPsCfP»C-PrP¶PµPSPSC¶C...
(CfPµCḡPµPrPSC-Pnē PīPsPeP°P·PSPēPe PI PjPµP¶P°C... PIC-Pr 17,6 % PrPs 21,5 %)
PīPsCḡC-PIPSC¶PSPs P· C, C-PsC, CḡPsPīC-C°Pj (CfPµCḡPµPrPSC-Pnē PīPsPeP°P·PSPēPe PI
PjPµP¶P°C... PIC-Pr 11,7 % PrPs 13,4 %).

PḡP°P±P»PēC†C¶ 1. P·PµPīPµPSPµPīP° C, CḡPSPeC†C-C¶, CfPēPjPīC, PsPjPē C, P° C¶PeC-CfC, Cḡ
P¶PēC, C, C¶, PīPsPīBḡ™ C¶P·P°PSC- P·C- P·PrPsCḡPsPīBḡ™ C¶Pj, PSP° 24-PjCḡ C, PēP¶PSC-

PµPsCḡC-PIPSC¶PSPSC¶ P»C-PeCḡPīP°PSPSC¶ P»C-PeP°CḡCfCḡPePēPj P·P°CfPsP±PsPj PḡPSPsCḡPs P·P»P»C-PīC, P° Cḡ PrPsP·C- 55/22 PjPePi	P C-P·PSPēC†C¶ PjC-P¶ PīPsPeP°P·PSPēPeP°PjPē P»C-PeCḡPīP°PSPSC¶ ¹ (95 % PrPsPIC-CḡC‡PēPnē C-PSC, PµCḡPīP°P», P·PSP°C‡PµPSPSC¶ p)			
	P’PēC...C-PrPSPµ P·PSP°C‡PµPSPSC¶ PḡPḡP’ ₁ (PjP»)	PµPsPeP°P·PSPēPe TDI	PµPsPeP°P·PSPēPe SGRQ	P’PēPePsCḡPēCfC, P°PSPSC¶ PīCḡPµPīP°CḡP°C, C-PI PrP»C¶ PīPsP»PµPiCēPµPSPSC¶ CfPēPjPīC, PsPjC-PI ³
PḡPSPsCḡPs P·P»P»C-PīC, P° (N = 413) PīPsCḡC-PIPSC¶PSPs P· PīP»P°C†PµP±Ps (N = 280)	167 (128, 207) < 0,001	1,2 (0,7, 1,7) < 0,001	Bḡ”5,51 (Bḡ”7,88, Bḡ”3,13) < 0,001*	Bḡ”0,8 (Bḡ”1,3, Bḡ”0,3) 0,001*

PhPSPsC ₇ Ps P•P»P»C-PiC,P° (N = 413) PiPsC ₇ C-PIPSC ₇ IPSPs P· CrPjPμPeP»C-PrPēPSC-CrPjPsPj Cr PrPsP·C- 55 PjPePi (N = 418)	52 (17, 87) 0,004	0,3 (B ₇ “0,2, 0,7) 0,244	B ₇ “0,82 (B ₇ “2,90, 1,27) 0,441	B ₇ “0,6 (B ₇ “1,0, B ₇ “0,1) 0,014*
PhPSPsC ₇ Ps P•P»P»C-PiC,P° (N = 413) PiPsC ₇ C-PIPSC ₇ IPSPs P· PIC-P»P°PSC,PμC ₇ PsP»PsPj Cr PrPsP·C- 22 PjPePi (N = 421)	95 (60, 130) < 0,001	0,4 (B ₇ “0,1, 0,8) 0,117	B ₇ “0,32 (B ₇ “2,41, 1,78) 0,767	0,1 (B ₇ “0,3, 0,5) 0,675
PhPSPsC ₇ Ps P•P»P»C-PiC,P° (N = 454) PiPsC ₇ C-PIPSC ₇ IPSPs P· C,C-PsC,C ₇ PsPiC-C”Pj Cr PrPsP·C- 18 PjPePi (N = 451) (PrPsC ₇ P»C-PrP¶PμPSPSC ₇ ZEP117115)	112 (81, 144) < 0,001	PS/Ps	B ₇ “2,10 (B ₇ “3,61, B ₇ “0,59) 0,006	B ₇ “0,5 (B ₇ “0,7, B ₇ “0,2) < 0,001
PhPSPsC ₇ Ps P•P»P»C-PiC,P° B (N = 207) PiPsC ₇ C-PIPSC ₇ IPSPs P· C,C-PsC,C ₇ PsPiC-C”Pj Cr PrPsP·C- 18 PjPePi (N = 203) (PrPsC ₇ P»C-PrP¶PμPSPSC ₇ DB2113360)	90 (39, 141) < 0,001	0,1 ² (B ₇ “0,4, 0,5) 0,817	0,75 (B ₇ “2,12, 3,63) 0,607	B ₇ “0,7 (B ₇ “1,2, B ₇ “0,1) 0,022
PhPSPsC ₇ Ps P•P»P»C-PiC,P° (N = 217) PiPsC ₇ C-PIPSC ₇ IPSPs P· C,C-PsC,C ₇ PsPiC-C”Pj Cr PrPsP·C- 18 PjPePi (N = 215) (PrPsC ₇ P»C-PrP¶PμPSPSC ₇ DB2113374)	60 (10, 109) 0,018*		B ₇ “0,17 (B ₇ “2,85, 2,52) 0,904	B ₇ “0,6 (B ₇ “1,2, 0,0) 0,069

N B₇” C±PēC₇PμP»CH₇PSC-C₇C,CH₇ PiPsPiC₇P»C₇C±C-C— P±P₇P₇ (P±P₇P₇ B₇”
PiPsPiC₇P»C₇C±C-C₇ PIC₇C-C... PiP°C±C-C”PSC,C-PI, C₇P°PSPPrPsPjC-P·PsPIP°PSPēC... PrP»C₇
P»C-PēC₇PIP°PSPSC₇)

PjPePi B₇” PjC-PēC₇PsPiC₇P°Pj

PS/Ps B₇” PSPμ PsC±C-PSCH₇PIP°PIC₇C₇

¹ P₇P₇μC₇PμPrPSC₇PsPePIP°PrC₇P°C,PēC±PSPμ C₇PμC₇PμPrPSC”.

² PhP±B₇™ C”PrPSP°PSC- PrP°PSC- PrPsC₇P»C-PrP¶PμPSPSC₇ DB2113360 C,P°
PrPsC₇P»C-PrP¶PμPSPSC₇ DB2113374.

³ P C-P·PSPēC±C₇ C₇PμC₇PμPrPSC₇PsC— PeC-P»CH₇PePsC₇C, C- P·P°C₇C,PsC₇C₇PIP°PSPSC₇
PiC₇PμPiP°C₇P°C,C₇ PSP° PrPμPSC₇ PiC₇PsC,C₇PiPsPj C,PēP¶PSC-PI 1B₇“24.

P’C-P»CH₇Cē PIPēC₇PsPeCr PrPsP·Cr CrPjPμPeP»C-PrPēPSC-CrPjCr/PIC-P»P°PSC,PμC₇PsP»Cr
(113/22 PjPePi) C,P°PePsP¶ PIPēPIC±P°P»Pē Cr 24-C,PēP¶PSPμPIPjCr PiP»P°C±PμP±Ps-
PePsPSC,C₇PsP»CH₇PsPIP°PSPjCr PeP»C-PSC-C±PSPsPjCr PrPsC₇P»C-PrP¶PμPSPSC- C,P° Cr
PrPIPsC... C-P· C,C₇CH₇PsC... 24-C,PēP¶PSPμPIPēC... PrPsC₇P»C-PrP¶PμPSPSC₇C...,
PePsPSC,C₇PsP»CH₇PsPIP°PSPēC... P·P° P°PeC,PēPIPS₇Pj PiC₇PμPiP°C₇P°C,PsPj.
P PμP·CrP»CH₇C,P°C,Pē P±C₇P»Pē P°PSP°P»PsPiC-C±PSC- C₇PμP·CrP»CH₇C,P°C,P°Pj PiC₇Pē
P·P°C₇C,PsC₇C₇PIP°PSPSC- P»C-PēP°C₇C₇CH₇PePsPiPs P·P°C₇PsP±Cr PhPSPsC₇Ps
P·P»P»C-PiC,P° Cr PrPsP·C- 55/22 PjPePi C,P° PrPsPrP°C,PePsPIPs PiC-PrC,PIPμC₇PrPēP»Pē

$P_{\mu C}, P_{\mu PeC}, P_{\mu PIPSC-ClC}, CHB$ $P \gg C-PeP^{\circ}CH_3ClCHPePsPiPs$ $P \cdot P^{\circ}ClPsP \pm Cr$ $PhPSPsCHPs$
 $P \cdot P \gg P \gg C-PiC, P^{\circ}$.

$P-P^{\circ}PiPsClC, CH_3P_{\mu PSPSCl} PIPhP-P$

PJ 24-C, $P_{\mu P}PSP_{\mu PIPsPjCr}$ $PiP \gg P^{\circ}C \dagger P_{\mu P \pm Ps-PePsPSC}, CH_3PsP \gg CH_3PsPIP^{\circ}PSPsPjCr$
 $PrPsClP \gg C-PrP_{\mu P}PSPSC- PiP^{\circ}C \dagger C-C''PSC, C-PI C-P$ $ClP_{\mu}PjPiC, PsPjPSP_{\mu}Pj PiP_{\mu}CH_3P_{\mu P \pm C-PiPsPj}$
 $PIPhP-P \gg P \cdot P^{\circ}ClC, PsClC \dagger PIP^{\circ}PSPSCl P \gg C-PeP^{\circ}CH_3ClCHPePsPiPs$ $P \cdot P^{\circ}ClPsP \pm Cr$ $PhPSPsCHPs$
 $P \cdot P \gg P \gg C-PiC, P^{\circ}$ $P \cdot PSP_{\mu}P_{\mu}ClPIP^{\circ}P \gg Ps$ $CH_3P_{\mu}P_{\mu}P_{\mu}PiPs/C, ClIP_{\mu}PePsPiPs$
 $P \cdot P^{\circ}PiPsClC, CH_3P_{\mu}PSPSCl PIPhP-P \gg PSP^{\circ} 50 \%$ $PiPsCH_3C-PIPSClPSPs$ $P \cdot PiP \gg P^{\circ}C \dagger P_{\mu P \pm Ps}$ (PSP°
 $PsClPSPsPIC- P^{\circ}PSP^{\circ}P \gg C-P \cdot Cr$ $C \dagger P^{\circ}ClCr$ $PrPs$ $CH_3PsP \cdot PIP_{\mu}C, PeCr$ $PiP_{\mu}CH_3C_{\mu}PsPiPs$
 $P \cdot P^{\circ}PiPsClC, CH_3P_{\mu}PSPSCl: PIC-PrPSPsC_{\mu}P_{\mu}PSPSCl$ $CH_3P_{\mu}P_{\mu}P_{\mu}C-PI$ ($P'P$) 0,5; 95 % $P''P \dagger$ 0,3,
0,8; $p = 0,004^*$), $PSP^{\circ} 20 \%$ $PiPsCH_3C-PIPSClPSPs$ $P \cdot CrPjP_{\mu}PeP \gg C-PrP_{\mu}PSC-CrPjPsPj$ ($P'P$ 0,8; 95
% $P''P \dagger$ 0,5, 1,3; $p = 0,391$) C, P° $PSP^{\circ} 30 \%$ $PiPsCH_3C-PIPSClPSPs$ $P \cdot PIC-P \gg P^{\circ}PSC, P_{\mu}CH_3PsP \gg PsPj$
($P'P$ 0,7; 95 % $P''P \dagger$ 0,4, 1,1; $p = 0,121$).

$P' PsPrPSPsPjCr$ $P \cdot C, CH_3CHPsC... PrPsClP \gg C-PrP_{\mu}P_{\mu}PSCH$ Cr $PiP^{\circ}C \dagger C-C''PSC, C-PI C-P$
 $ClP_{\mu}PjPiC, PsPjPSP_{\mu}Pj PiP_{\mu}CH_3P_{\mu P \pm C-PiPsPj}$ $PIPhP-P \gg P \cdot P^{\circ}ClC, PsClC \dagger PIP^{\circ}PSPSCl$
 $P \gg C-PeP^{\circ}CH_3ClCHPePsPiPs$ $P \cdot P^{\circ}ClPsP \pm Cr$ $PhPSPsCHPs$ $P \cdot P \gg P \gg C-PiC, P^{\circ}$ Cr
 $PiPsCH_3C-PIPSClPSPSC- P \cdot C, C-PsC, CH_3PsPiC-C''Pj$ $P \cdot PSP_{\mu}P_{\mu}ClPIP^{\circ}P \gg Ps$ $CH_3P_{\mu}P_{\mu}P_{\mu}Pe$
 $PiPsPjC-CH_3PSPsPiPs/C, ClIP_{\mu}PePsPiPs$ $P \cdot P^{\circ}PiPsClC, CH_3P_{\mu}PSPSCl PIPhP-P \gg PSP^{\circ} 50 \%$ ($P'P$ 0,5;
95 % $P''P \dagger$ 0,3, 1,0; $p = 0,044$). PJ $PrPIPsC... C-PSC_{\mu}P_{\mu}C... PrPsClP \gg C-PrP_{\mu}P_{\mu}PSPSClC...$
 $CH_3P_{\mu}P_{\mu}P_{\mu}Pe PiPsPjC-CH_3PSPsPiPs/C, ClIP_{\mu}PePsPiPs$ $P \cdot P^{\circ}PiPsClC, CH_3P_{\mu}PSPSCl PIPhP-P \gg Cr$
 $PiP^{\circ}C \dagger C-C''PSC, C-PI C-P$ $ClP_{\mu}PjPiC, PsPjPSP_{\mu}Pj PiP_{\mu}CH_3P_{\mu P \pm C-PiPsPj}$ $PIPhP-P \gg P \pm CrPI$
 $PIC-PrPiPsPIC-PrPSPs$ $PSP^{\circ} 20 \% C, P^{\circ} 90 \% PIP_{\mu}C_{\mu}P_{\mu}Pj$ ($P'P$ 1,2; 95 % $P''P \dagger$ 0,5, 2,6; $p = 0,709$
 C, P° $P'P$ 1,9; 95 % $P''P \dagger$ 1,0, 3,6; $p = 0,062$ $PIC-PrPiPsPIC-PrPSPs$). $P''P_{\mu}P \cdot P^{\circ}P_{\mu}PS$ $C \dagger P_{\mu}C...$
 $PrPsClP \gg C-PrP_{\mu}P_{\mu}PSCH$ $PSP_{\mu} P \pm CrPI$ $CH_3PsP \cdot CH_3PsP \pm P \gg P_{\mu}PSP_{\mu}P_{\mu}P_{\mu} PrP \gg Cl PsC \dagger C-PSPeP_{\mu}$
 $PIPiP \gg P_{\mu}PICr P \gg C-PeCrPIP^{\circ}PSPSCl PSP^{\circ}$ $P \cdot P^{\circ}PiPsClC, CH_3P_{\mu}PSPSCl PIPhP-P \gg, C-$
 $PiP^{\circ}C \dagger C-C''PSC, P_{\mu}PiC-ClP \gg Cl$ $CH_3PsP \cdot PIP_{\mu}C, PeCr$ $P \cdot P^{\circ}PiPsClC, CH_3P_{\mu}PSPSCl$
 $PIP_{\mu}PeP \gg CHC \dagger P^{\circ}P \gg P_{\mu}ClC$ $P \cdot PrPsClP \gg C-PrP_{\mu}P_{\mu}PSPSCl$.

$P''PsClP \gg C-PrP_{\mu}P_{\mu}PSPSCl, C_{\mu}Ps PiC-PrC, PIP_{\mu}CH_3PrP_{\mu}C \dagger CHC, CH$
 $P_{\mu C}, P_{\mu PeC}, P_{\mu PIPSC-ClC}, CHB$

PJ $CH_3P^{\circ}PSP_{\mu}PsPjC-P \cdot PsPIP^{\circ}PSPsPjCr$ $PiPsPrPIC-PN_{\mu}PSPs$ $ClP \gg C-PiPsPjCr$ 52-C, $P_{\mu}P_{\mu}PSP_{\mu}PIP_{\mu}PjCr$
 $PrPsClP \gg C-PrP_{\mu}P_{\mu}PSPSC- (CTT116855, IMPACT) PI 10 355 PrPsCH_3PsClP \gg P_{\mu}C...$
 $PiP^{\circ}C \dagger C-C''PSC, C-PI C-P$ $ClP_{\mu}PjPiC, PsPjPSP_{\mu}Pj PiP_{\mu}CH_3P_{\mu P \pm C-PiPsPj}$ $PIPhP-P \gg, ClIP_{\mu}C-$
 $PiP_{\mu}CH_3P_{\mu}PSP_{\mu}ClP \gg P_{\mu}PsPrPSP_{\mu} C \dagger P_{\mu}PeC-P \gg CHPeP^{\circ}$ $PiPsPjC-CH_3PSP_{\mu}C... P^{\circ}P \pm Ps$
 $C, ClIP_{\mu}PeP_{\mu}C... P \cdot P^{\circ}PiPsClC, CH_3P_{\mu}PSCH PiCH_3PsC, ClIPiPsPj PiPsPiP_{\mu}CH_3P_{\mu}PrPSC-C... 12$
 $PjC-ClC \dagger C-PI, PiPsCH_3C-PIPSC_{\mu}PIP^{\circ}P \gg PsClCH$ $P \gg C-PeCrPIP^{\circ}PSPSCl$
 $CrPjP_{\mu}PeP \gg C-PrP_{\mu}PSC-CrPjPsPj/PIC-P \gg P^{\circ}PSC, P_{\mu}CH_3PsP \gg PsPj$ ($PjP_{\mu}P \cdot P_{\mu}/P'P \dagger 55/22$ $PjPePi$),
 $C_{\mu}P \gg CHC, P_{\mu}PeP^{\circ}P \cdot PsPSCr$ $C_{\mu}CrCH_3PsP^{\circ}C, PsPj / CrPjP_{\mu}PeP \gg C-PrP_{\mu}PSC-CrPjPsPj /$
 $PIC-P \gg P^{\circ}PSC, P_{\mu}CH_3PsP \gg PsPj$ ($P_{\mu}P_{\mu}/PjP_{\mu}P \cdot P_{\mu}/P'P \dagger 99/55/22$ $PjPePi$) $C \dagger P_{\mu}$
 $C_{\mu}P \gg CHC, P_{\mu}PeP^{\circ}P \cdot PsPSCr$ $C_{\mu}CrCH_3PsP^{\circ}C, PsPj / PIC-P \gg P^{\circ}PSC, P_{\mu}CH_3PsP \gg PsPj$ ($P_{\mu}P_{\mu}/P'P \dagger 99/22$
 $PjPePi$) $PsPrP_{\mu}PS$ $CH_3P^{\circ}P \cdot PSP^{\circ}$ $PrPsP \pm Cr$ $PI PsPrPSPsPjCr$ $C-PSPiP^{\circ}P \gg ClC, PsCH_3C-$
 $P_{\mu}P_{\mu}CH_3PIP_{\mu}PSPSPsCH$ $PeC-PSC \dagger P_{\mu}PIP_{\mu}CH$ $C, PsC \dagger PePsCH$ $PI PrPsClP \gg C-PrP_{\mu}P_{\mu}PSPSC-$
 $P \pm CrP \gg P^{\circ} C_{\mu}PsCH_3C-C \dagger PSP^{\circ} C \dagger P^{\circ}ClC, PsC, P^{\circ}$ $PiPsPjC-CH_3PSP_{\mu}C... C \dagger P_{\mu}C, ClIP_{\mu}PeP_{\mu}C...$
 $P \cdot P^{\circ}PiPsClC, CH_3P_{\mu}PSCH$ Cr $C, P_{\mu}C..., C...C, Ps$ $PiCH_3P_{\mu}P_{\mu}PjP^{\circ}PI$ $P_{\mu}P_{\mu}/PjP_{\mu}P \cdot P_{\mu}/P'P \dagger$,
 $PiPsCH_3C-PIPSClPSPs$ $P \cdot P_{\mu}P_{\mu}/P'P \dagger P^{\circ}P \pm Ps$ $PjP_{\mu}P \cdot P_{\mu}/P'P \dagger$. $P_{\mu}P_{\mu}CH_3P_{\mu}PrPSCl$ $CH_3C-C \dagger PSP^{\circ}$
 $C \dagger P^{\circ}ClC, PsC, P^{\circ}$ $P \cdot P^{\circ}PiPsClC, CH_3P_{\mu}PSCH$ $ClC, P^{\circ}PSPsPIP_{\mu}P \gg P^{\circ} 0,91, 1,07$ $C, P^{\circ} 1,21$ $PrP \gg Cl$
 $P_{\mu}P_{\mu}/PjP_{\mu}P \cdot P_{\mu}/P'P \dagger, P_{\mu}P_{\mu}/P'P \dagger C, P^{\circ}$ $PjP_{\mu}P \cdot P_{\mu}/P'P \dagger$ $PIC-PrPiPsPIC-PrPSPs$.

$P_{\mu}PsC\bar{H}C-PIPSC\bar{U}PSPSC\bar{U}$ $P_{\alpha}P_{\alpha}/PJP_{\bar{H}}P \cdot P_{\bar{L}}/P'P \dagger P \cdot P_{\alpha}P_{\alpha}/P'P \dagger C, P^{\circ} PJP_{\bar{H}}P \cdot P_{\bar{L}}/P'P \dagger$
 $P\bar{i}PsPeP^{\circ}P \cdot P^{\circ}P \gg Ps$ $C\bar{I}C, P^{\circ}C, P\bar{e}C\bar{I}C, P\bar{e}C\ddagger PSPs$ $P \cdot PSP^{\circ}C\ddagger CrC\%_{\circ}P_{\mu}$ $P \cdot PSP\bar{e}P\bar{I}P_{\mu}PSPSC\bar{U}$
 $C\bar{H}P\bar{e}P \cdot P\bar{e}PeCr$ $P\bar{i}PsPjC-C\bar{H}PSPsPiPs/C, C\bar{U}P\bar{I}PePsPiPs$ $P \cdot P^{\circ}PiPsC\bar{I}C, C\bar{H}P_{\mu}PSPSC\bar{U}$ $PSP^{\circ} 14,8 \%$
 $(PSP^{\circ} PsC\bar{I}PSPsPIC- P^{\circ}PSP^{\circ}P \gg C-P \cdot Cr$ $C\ddagger P^{\circ}C\bar{I}Cr$ $PrPs$ $C\bar{H}PsP \cdot PIP\bar{e}C, PeCr$ $P\bar{i}P_{\mu}C\bar{H}C\bar{e}PsPiPs$
 $P \cdot P^{\circ}PiPsC\bar{I}C, C\bar{H}P_{\mu}PSPSC\bar{U})$ $(P'P \ 0,85; 95 \%$ $P''P \dagger 0,80, 0,91; p < 0,001)$ $C, P^{\circ} PSP^{\circ} 16 \%$
 $PIC-PrP\bar{i}PsPIC-PrPSPs$ $(PSP^{\circ} PsC\bar{I}PSPsPIC- P^{\circ}PSP^{\circ}P \gg C-P \cdot Cr$ $C\ddagger P^{\circ}C\bar{I}Cr$ $PrPs$ $C\bar{H}PsP \cdot PIP\bar{e}C, PeCr$
 $P\bar{i}P_{\mu}C\bar{H}C\bar{e}PsPiPs$ $P \cdot P^{\circ}PiPsC\bar{I}C, C\bar{H}P_{\mu}PSPSC\bar{U})$ $(P'P \ 0,84; 95 \%$ $P''P \dagger 0,78, 0,91; p < 0,001)$.

$P'P\bar{e}C, C\bar{H}P\bar{e}PIP^{\circ}P \gg C-C\bar{I}C, CH$ $C, P^{\circ} PsP \pm B\bar{H}^{\text{TM}} C''Pj$ $P \gg P_{\mu}PiP_{\mu}PSC\bar{H}$

$P' PsPrPSPsPjCr$ $P \cdot PrPIPSC \dots PrPsC\bar{I}P \gg C-PrP\bar{I}P_{\mu}PSC\bar{H}$ $P \gg C-PeP^{\circ}C\bar{H}C\bar{I}CHPePsPiPs$
 $P \cdot P^{\circ}C\bar{I}PsP \pm Cr$ $P\bar{H}PSPsC\bar{H}Ps$ $P \cdot P \gg P \gg C-P\bar{i}C, P^{\circ} Cr$ $PrPsP \cdot C- 55/22 PjPePi$
 $P\bar{i}PsPeC\bar{H}P^{\circ}C\%_{\circ}P\bar{e}PIC\bar{I}C\bar{U}$ $C\ddagger P^{\circ}C\bar{I} PIP\bar{e}C, C\bar{H}P\bar{e}PIP^{\circ}P \gg PsC\bar{I}C, C-$ $P\bar{i}PsC\bar{H}C-PIPSC\bar{U}PSPs$ $P \cdot$
 $C, P^{\circ}PeP\bar{e}Pj$ $P\bar{i}C\bar{H}P\bar{e}$ $P \cdot P^{\circ}C\bar{I}C, PsC\bar{I}CrPIP^{\circ}PSPSC-$ $P\bar{i}P \gg P^{\circ}C\ddagger P_{\mu}P \pm Ps, C\%_{\circ}Ps$
 $PsC\ddagger C-PSC\bar{H}PIP^{\circ}P \gg PsC\bar{I}C\bar{U}$ $P \cdot P^{\circ} PrPsP\bar{i}PsPjPsPiPsC\bar{H}$ $C\bar{I}C, CrP\bar{i}C-PSC\ddagger P^{\circ}C\bar{I}C, PsPiPs$ $C, P_{\mu}C\bar{I}C, Cr$
 $C\ddagger PsPIPSP\bar{e}PePsPiPsC-$ $C \dots PsPrCHP \pm P\bar{e}$ $(C\bar{e}P^{\circ}C, P \gg C, P_{\mu}C\bar{I}C, Cr)$. $P' PsP \pm PsC \dots$
 $PrPsC\bar{I}P \gg C-PrP\bar{I}P_{\mu}PSPSC\bar{U}C \dots C\bar{I}P\bar{i}PsC\bar{I}C, P_{\mu}C\bar{H}C-PiP^{\circ}P \gg PsC\bar{I}C\bar{U}$ $P \cdot P \pm C-P \gg CHC\bar{e}P_{\mu}PSPSC\bar{U}$
 $P\bar{i}PsPeP^{\circ}P \cdot PSP\bar{e}PeC-Pi$ $PsP \pm B\bar{H}^{\text{TM}} C''PjCr$ $P \gg P_{\mu}PiP_{\mu}PSC\bar{H}$ $P\bar{i}PsC\bar{H}C-PIPSC\bar{U}PSPs$ $P \cdot$
 $P\bar{i}P \gg P^{\circ}C\ddagger P_{\mu}P \pm Ps$ Cr $PrPsC\bar{H}PsC\bar{I}P \gg P\bar{e}C \dots P\bar{i}P^{\circ}C\ddagger C-C''PSC, C-Pi$ $C-P$ $P\bar{I}P\bar{H}P-P \gg P \cdot$
 $PiC-PiP_{\mu}C\bar{H}C-PSC, P \gg C\bar{U}C\ddagger C-C''C\bar{H}$ $(C, CrPSPeC\ddagger C-PsPSP^{\circ}P \gg CHPSP^{\circ} P \cdot P^{\circ}P \gg P\bar{e}C\bar{e}PePsPIP^{\circ}$
 $C''PjPSC-C\bar{I}C, CH$ $(P_{\alpha}P-P,) > 120 \%)$. Pj $P\bar{i}P_{\mu}C\bar{H}C\bar{e}PsPjCr$ $PrPsC\bar{I}P \gg C-PrP\bar{I}P_{\mu}PSPSC-$
 $P \gg C-PeP^{\circ}C\bar{H}C\bar{I}CHPeP\bar{e}PN_{\bar{e}}$ $P \cdot P^{\circ}C\bar{I}C-P \pm P\bar{H}PSPsC\bar{H}Ps$ $P \cdot P \gg P \gg C-P\bar{i}C, P^{\circ} Cr$ $PrPsP \cdot C- 55/22 PjPePi$
 $P\bar{i}C\bar{H}PsPrP_{\mu}PjPsPSC\bar{I}C, C\bar{H}CrPIP^{\circ}Pi$ $C\bar{I}C, P^{\circ}C, P\bar{e}C\bar{I}C, P\bar{e}C\ddagger PSPs$ $PrPsC\bar{I}C, PsPIC-C\bar{H}PSP_{\mu}$ C, P°
 $PeP \gg C-PSC-C\ddagger PSPs$ $P \cdot PSP^{\circ}C\ddagger CrC\%_{\circ}P_{\mu}$ $P\bar{i}PsPeC\bar{H}P^{\circ}C\%_{\circ}P_{\mu}PSPSC\bar{U}$ $C\ddagger P^{\circ}C\bar{I}Cr$
 $PIP\bar{e}C, C\bar{H}P\bar{e}PIP^{\circ}P \gg PsC\bar{I}C, C-$ $P\bar{i}C\bar{H}P\bar{e}$ $C, C-P \cdot P\bar{e}C\ddagger PSPsPjCr$ $PSP^{\circ}PIP^{\circ}PSC, P^{\circ}P\bar{I}P_{\mu}PSPSC-$ $(69,4$
 $C\bar{I}P_{\mu}PeCrPSP\bar{r}P\bar{e}, p = 0,003)$ $P\bar{i}PsC\bar{H}C-PIPSC\bar{U}PSPs$ $P \cdot P\bar{i}P \gg P^{\circ}C\ddagger P_{\mu}P \pm Ps, C\%_{\circ}Ps$
 $PsC\ddagger C-PSC\bar{H}PIP^{\circ}P \gg PsC\bar{I}CH$ $PSP^{\circ} 12$ $C, P\bar{e}P\bar{I}PSC-$ $PrPsC\bar{I}P \gg C-PrP\bar{I}P_{\mu}PSPSC\bar{U}$ $C\ddagger P_{\mu}C\bar{H}P_{\mu}P \cdot 3$
 $PiPsPrP\bar{e}PSP\bar{e}$ $P\bar{i}C-C\bar{I}P \gg C\bar{U}$ $P \cdot P^{\circ}C\bar{I}C, PsC\bar{I}CrPIP^{\circ}PSPSC\bar{U}$ $P\bar{i}C\bar{H}P_{\mu}P\bar{i}P^{\circ}C\bar{H}P^{\circ}C, Cr$
 $(PjC-PSC-PjP^{\circ}P \gg CHPSP^{\circ} PeP \gg C-PSC-C\ddagger PSPs$ $P \cdot PSP^{\circ}C\ddagger P\bar{e}PjP^{\circ}$ $C\bar{H}C-P \cdot PSP\bar{e}C\ddagger C\bar{U}$ $(P_{\bar{H}}P_{\bar{L}}P-P)$
 $PrP \gg C\bar{U}$ $C\bar{e}P^{\circ}C, P \gg C, P_{\mu}C\bar{I}C, Cr$ $C\bar{I}C, P^{\circ}PSPsPIP\bar{e}C, CH$ $45B\bar{H}^{\text{TM}} 85$ $C\bar{I}$). $P_{\mu}PsPeC\bar{H}P^{\circ}C\%_{\circ}P_{\mu}PSPSC\bar{U}$
 $C\ddagger P^{\circ}C\bar{I}Cr$ $PIP\bar{e}C, C\bar{H}P\bar{e}PIP^{\circ}P \gg PsC\bar{I}C, C-$ $P\bar{i}C\bar{H}P\bar{e}$ $C, C-P \cdot P\bar{e}C\ddagger PSPsPjCr$ $PSP^{\circ}PIP^{\circ}PSC, P^{\circ}P\bar{I}P_{\mu}PSPSC-$
 $P\bar{i}PsC\bar{H}C-PIPSC\bar{U}PSPs$ $P \cdot C, P^{\circ}PeP\bar{e}Pj$ $P\bar{i}C\bar{H}P\bar{e}$ $P \cdot P^{\circ}C\bar{I}C, PsC\bar{I}CrPIP^{\circ}PSPSC-$ $P\bar{i}P \gg P^{\circ}C\ddagger P_{\mu}P \pm Ps$
 $C\bar{I}P\bar{i}PsC\bar{I}C, P_{\mu}C\bar{H}C-PiP^{\circ}P \gg PsC\bar{I}CH$ $PSP^{\circ} 2$ $PrP_{\mu}PSC\bar{H}$ $C, P^{\circ} P \cdot P \pm P_{\mu}C\bar{H}C-PiP^{\circ}P \gg PsC\bar{I}C\bar{U}$ $PSP^{\circ} 6-$
 $PjCr$ $C, P^{\circ} 12-PjCr$ $C, P\bar{e}P\bar{I}PSC\bar{U}C \dots PrPsC\bar{I}P \gg C-PrP\bar{I}P_{\mu}PSPSC\bar{U}$. Pj $PrC\bar{H}CrPiPsPjCr$
 $PrPsC\bar{I}P \gg C-PrP\bar{I}P_{\mu}PSPSC-$ $C\bar{H}C-P \cdot PSP\bar{e}C\ddagger C\bar{U}$ $PjC-P\bar{I}$ $P\bar{i}PsPeP^{\circ}P \cdot PSP\bar{e}PeP^{\circ}PjP\bar{e}$ $C\ddagger P^{\circ}C\bar{I}Cr$
 $PIP\bar{e}C, C\bar{H}P\bar{e}PIP^{\circ}P \gg PsC\bar{I}C, C-$ $P\bar{i}C\bar{H}P\bar{e}$ $C, C-P \cdot P\bar{e}C\ddagger PSPsPjCr$ $PSP^{\circ}PIP^{\circ}PSC, P^{\circ}P\bar{I}P_{\mu}PSPSC-$ Cr
 $PiC\bar{H}CrPiC-$ $P \cdot P^{\circ}C\bar{I}C, PsC\bar{I}CrPIP^{\circ}PSPSC\bar{U}$ $P \gg C-PeP^{\circ}C\bar{H}C\bar{I}CHPePsPiPs$ $P \cdot P^{\circ}C\bar{I}PsP \pm Cr$ $P\bar{H}PSPsC\bar{H}Ps$
 $P \cdot P \gg P \gg C-P\bar{i}C, P^{\circ} Cr$ $PrPsP \cdot C- 55/22 PjPePi$ $C, P^{\circ} P\bar{i}P \gg P^{\circ}C\ddagger P_{\mu}P \pm Ps$ $PSP^{\circ} 12$ $C, P\bar{e}P\bar{I}PSC-$
 $C\bar{I}C, P^{\circ}PSPsPIP\bar{e}P \gg P^{\circ} 21,9$ $C\bar{I}P_{\mu}PeCrPSP\bar{r}P\bar{e}$ $(p = 0,234)$.

$P \gg C-PeP^{\circ}C\bar{H}C\bar{I}CHPeP\bar{e}PN_{\bar{e}}$ $P \cdot P^{\circ}C\bar{I}C-P \pm P\bar{H}PSPsC\bar{H}Ps$ $P \cdot P \gg P \gg C-P\bar{i}C, P^{\circ} Cr$ $PrPsP \cdot C- 55/22 PjPePi$
 $C, P^{\circ}PePsP\bar{I}$ $P\bar{i}PsPeP^{\circ}P \cdot P^{\circ}Pi$ $C\bar{I}C, P^{\circ}C, P\bar{e}C\bar{I}C, P\bar{e}C\ddagger PSPs$ $P \cdot PSP^{\circ}C\ddagger CrC\%_{\circ}P_{\mu}$ $P\bar{i}PsP \gg C-P\bar{i}C\bar{e}P_{\mu}PSPSC\bar{U}$
 $PIP\bar{e}C \dots C-PrPSP\bar{e}C \dots P\bar{i}PsPeP^{\circ}P \cdot PSP\bar{e}PeC-Pi$ $PsP \pm B\bar{H}^{\text{TM}} C''PjCr$ $P \gg P_{\mu}PiP_{\mu}PSC\bar{H}$
 $P\bar{i}PsC\bar{H}C-PIPSC\bar{U}PSPs$ $P \cdot P\bar{i}P \gg P^{\circ}C\ddagger P_{\mu}P \pm Ps$. $P' P\bar{i}P_{\mu}C\bar{H}C\bar{e}PsPjCr$ $PrPsC\bar{I}P \gg C-PrP\bar{I}P_{\mu}PSPSC-$
 $PIP\bar{e}PjC-C\bar{H}C\bar{H}PIP^{\circ}PSPSC\bar{U}$ $P\bar{i}C\bar{H}PsPiPsPrP\bar{e}P \gg P\bar{e}$ $PSP^{\circ} 12-PjCr$ $C, P\bar{e}P\bar{I}PSC-$ $P\bar{i}P_{\mu}C\bar{H}P_{\mu}Pr$
 $P \cdot P^{\circ}C\bar{I}C, PsC\bar{I}CrPIP^{\circ}PSPSC\bar{U}Pj$ $P\bar{i}C\bar{H}P_{\mu}P\bar{i}P^{\circ}C\bar{H}P^{\circ}C, Cr$ $C, P^{\circ} C\ddagger P_{\mu}C\bar{H}P_{\mu}P \cdot 3$ $PiPsPrP\bar{e}PSP\bar{e}$
 $P\bar{i}C-C\bar{I}P \gg C\bar{U}$ $C-PSPiP^{\circ}P \gg C\bar{U}C\ddagger C-C-$ $C''PjPSC-C\bar{I}C, CH$ $PIP\bar{r}P\bar{e}C \dots Cr$ $C\bar{I}C, P^{\circ}PSPsPIP\bar{e}P \gg P^{\circ}$
 $237 PjP \gg C- 316 PjP \gg PIC-PrP\bar{i}PsPIC-PrPSPs$, $P \cdot P^{\circ}P \gg P\bar{e}C\bar{e}PePsPIP\bar{e}PN_{\bar{e}}$ $PsP \pm B\bar{H}^{\text{TM}} C''Pj$
 $B\bar{H}^{\text{TM}} 466 PjP \gg C, P^{\circ} B\bar{H}^{\text{TM}} 643 PjP \gg PIC-PrP\bar{i}PsPIC-PrPSPs$ $C, P^{\circ} C, CrPSPeC\ddagger C-PsPSP^{\circ}P \gg CHPSP^{\circ}$
 $P \cdot P^{\circ}P \gg P\bar{e}C\bar{e}PePsPIP^{\circ} C''PjPSC-C\bar{I}C, CH$ $B\bar{H}^{\text{TM}} 351 PjP \gg C- B\bar{H}^{\text{TM}} 522 PjP \gg PIC-PrP\bar{i}PsPIC-PrPSPs$
 $(PIC\bar{I}C- C\bar{H} < 0,001)$. Pj $PrC\bar{H}CrPiPsPjCr$ $PrPsC\bar{I}P \gg C-PrP\bar{I}P_{\mu}PSPSC-$ $PIP\bar{e}PjC-C\bar{H}C\bar{H}PIP^{\circ}PSPSC\bar{U}$

C,P°PePsP¶ PīCḡPsPIPsPrPēP»Pē PSP° 12-PjCṛ C,PēP¶PSC- PrPs C-PSPIP°P»CḶC†C-C—
PīCḡPμPīP°CḡP°C,Cṛ C,P° C‡PμCḡPμP· 3 PīPsPrPēPSPē PīC-CḶP»CḶ C-PSPIP°P»CḶC†C-C—:
C”PjPSC-CḶC,Cḡ PIPrPēC...Cṛ CḶC,P°PSPsPIPēP»P° 198 PjP» C- 238 PjP» PIC-PrPīPsPIC-PrPSPs,
P·P°P»PēC€PePsPIPēP№ PsP±Bḡ™ C”Pj Bḡ”295 PjP» C- Bḡ”351 PjP» PIC-PrPīPsPIC-PrPSPs
C,P° C,,CṛPSPeC†C-PsPSP°P»CḡPSP° P·P°P»PēC€PePsPIP° C”PjPSC-CḶC,Cḡ Bḡ”238 PjP»
C- Bḡ”302 PjP» PIC-PrPīPsPIC-PrPSPs (PICḶC- p < 0,001*).

PṛP°CḡPjP°PePsPeC-PSPμC,PēPeP°

PḡPsP»Pē PePsPjP±C-PSP°C†C-Cḡ CṛPjPμPeP»C-PrPēPSC-CṛPjCṛ C,P° PIC-P»P°PSC,PμCḡPsP»Cṛ
PIPIPsPrPēP»Pē C-PSPIP°P»CḶC†C-P№PSPs, C,,P°CḡPjP°PePsPeC-PSPμC,PēPeP° PePsP¶PSPsPīPs
PePsPjPīPsPSPμPSC,P° P±CṛP»P° PīPsPrC-P±PSP° C,,P°CḡPjP°PePsPeC-PSPμC,PēC†C-, C%oPs
CḶPīPsCḶC,PμCḡC-PiP°P»P°CḶCḶ PīCḡPē P·P°CḶC,PsCḶCṛPIP°PSPSC- PePsP¶PSPsC—
P°PeC,PēIPSPsC— CḡPμC‡PsPIPēPSPē PSP°CḡC-P·PSPs. PḡC,P¶Pμ, P·
C,,P°CḡPjP°PePsPeC-PSPμC,PēC‡PSPsPīPs PīPsPīP»CḶPrCṛ PePsP¶PμPS PePsPjPīPsPSPμPSC,
PjPsP¶PSP° CḡPsP·PiP»CḶPrP°C,Pē PsPeCḡPμPjPs.

PḡP±CḶPsCḡP±C†C-CḶ

PjPjPμPeP»C-PrPēPSC-CṛPj

PḡC-CḶP»CḶ PIPiPμPrPμPSPSCḶ CṛPjPμPeP»C-PrPēPSC-CṛPjCṛ Cṛ PIPēPiP»CḶPrC-
C-PSPIP°P»CḶC†C-P№ P·PrPsCḡPsPIPēPj PrPsP±CḡPsPIPsP»CḡC†CḶPj C_{max}
PrPsCḶCḶPīP°P»P°CḶCḶ C‡PμCḡPμP· 5Bḡ”15 C...PIPēP»PēPS. PḡP±CḶPsP»CḡC,PSP°
P±C-PsPrPsCḶC,CṛPīPSC-CḶC,Cḡ CṛPjPμPeP»C-PrPēPSC-CṛPjCṛ, PIPiPμPrPμPSPsPīPs Cṛ
PIPēPiP»CḶPrC- C-PSPIP°P»CḶC†C-P№, CḶC,P°PSPsPIPēP»P° Pī CḶPμCḡPμPrPSCḡPsPjCṛ 13 %
PIC-Pr PrPsP·Pē, PrPμ C‡P°CḶC,PeP°, PīPsPīBḡ™ CḶP·P°PSP° P· PīPμCḡPsCḡP°P»CḡPSPsCḡ
P°P±CḶPsCḡP±C†C-C”Cḡ, PSPμP·PSP°C‡PSP°. PḡC-CḶP»CḶ PīPsPIC,PsCḡPSPsC— PrPsP·Pē
CṛPjPμPeP»C-PrPēPSC-CṛPjCṛ, PIPiPμPrPμPSPsPīPs Cṛ PIPēPiP»CḶPrC- C-PSPIP°P»CḶC†C-P№,
CḡC-PIPSPsPIP°P¶PSPēP№ CḶC,P°PS PrPsCḶCḶPīP°PICḶCḶ PīCḡPsC,CḶPīPsPj 7Bḡ”10 PrPSC-PI
P· 1,5Bḡ”1,8-PeCḡP°C,PSPēPj PSP°PePsPīPēC‡PμPSPSCḶPj.

P’C-P»P°PSC,PμCḡPsP»

PḡC-CḶP»CḶ PIPiPμPrPμPSPSCḶ PIC-P»P°PSC,PμCḡPsP»Cṛ Cṛ PIPēPiP»CḶPrC-
C-PSPIP°P»CḶC†C-P№ P·PrPsCḡPsPIPēPj PrPsP±CḡPsPIPsP»CḡC†CḶPj C_{max}
PrPsCḶCḶPīP°P»P°CḶCḶ C‡PμCḡPμP· 5Bḡ”15 C...PIPēP»PēPS. PḡP±CḶPsP»CḡC,PSP°
P±C-PsPrPsCḶC,CṛPīPSC-CḶC,Cḡ PIC-P»P°PSC,PμCḡPsP»Cṛ, PIPiPμPrPμPSPsPīPs Cṛ
PIPēPiP»CḶPrC- C-PSPIP°P»CḶC†C-P№, CḶC,P°PSPsPIPēP»P° 27 %, PrPμ C‡P°CḶC,PeP°,
PīPsPīBḡ™ CḶP·P°PSP° P· PīPμCḡPsCḡP°P»CḡPSPsCḡ P°P±CḶPsCḡP±C†C-C”Cḡ,
PSPμP·PSP°C‡PSP°. PḡC-CḶP»CḶ PīPsPIC,PsCḡPSPsC— PrPsP·Pē PIC-P»P°PSC,PμCḡPsP»Cṛ,
PIPIPμPrPμPSPsPīPs Cṛ PIPēPiP»CḶPrC- C-PSPIP°P»CḶC†C-P№, CḡC-PIPSPsPIP°P¶PSPēP№
CḶC,P°PS PrPsCḶCḶPīP°PICḶCḶ PīCḡPsC,CḶPīPsPj 6 PrPSC-PI P· 2,4-PeCḡP°C,PSPēPj
PSP°PePsPīPēC‡PμPSPSCḶPj.

P PsP·PīPsPrC-P»

PjPjPμPeP»C-PrPēPSC-CṛPj

PḡC-CḶP»CḶ PIPSCṛC,CḡC-C€PSCḡPsPIPμPSPSPsPīPs PIPiPμPrPμPSPSCḶ P·PrPsCḡPsPIPēPj

PrPsP±C₇PsPIPsP»CHC†C₁Pj C₁P_μC₇P_μPrPSC-PN₂ PsP±B₇TMC^{''}Pj C₇PsP·P_iPsPrC-P»C₁
C₁C, P°PSPsPIPēPI 86 P»C-C, C₇C-PI. *In vitro* P·PIB₇TMC₁P·C₁PIP°PSPSC₁ P· P±C-P»PeP°PjPē
P_iP»P°P·PjPē PeC₇PsPIC- P»C₇PrPēPSPē C₁C, P°PSPsPIPēP»Ps PI C₁P_μC₇P_μPrPSC₇PsPjC₁ 89 %.

P'C-P»P°PSC, P_μC₇PsP»

P_μC-C₁P»C₁ PIPSC₁C, C₇C-C€PSC₇PsPIP_μPSPSPsPiPs PIP_μPrP_μPSPSC₁ P·PrPsC₇PsPIPēPj
PrPsP±C₇PsPIPsP»CHC†C₁Pj C₁P_μC₇P_μPrPSC-PN₂ PsP±B₇TMC^{''}Pj C₇PsP·P_iPsPrC-P»C₁ PI
C₇C-PIPSsPIP°P¶PSPsPjC₁ C₁C, P°PSC- C₁C, P°PSPsPIPēPI 165 P»C-C, C₇C-PI. *In vitro*
P·PIB₇TMC₁P·C₁PIP°PSPSC₁ P· P±C-P»PeP°PjPē P_iP»P°P·PjPē PeC₇PsPIC- P»C₇PrPēPSPē
C₁C, P°PSPsPIPēP»Ps PI C₁P_μC₇P_μPrPSC₇PsPjC₁ 94 %.

P'C-PsC, C₇P°PSC₁C, PsC₇PjP°C†C-C₁

PjPjP_μPeP»C-PrPēPSC-C₁Pj

P^{''}PsC₁P»C-PrP¶P_μPSPSC₁ C-*n vitro* P_iPsPeP°P·P°P»Pē, C%₀Ps C₁PjP_μPeP»C-PrPēPSC-C₁Pj
PiPsP»PsPIPSpēPj C†PēPSPsPj PjP_μC, P°P±PsP»C-P·C₁C^{''}C, CHC₁C₁ C†PēC, PsC...C₇PsPjPsPj P450
2D6 (CYP2D6) C- C^{''} C₁C₁P±C₁C, C₇P°C, PsPj PrP»C₁ P_iP_μC₇P_μPSPsC₁PSPēPeP° P -
PiP»C-PePsPiC₇PsC, P_μC—PSC₁ (P-gp). P_μP_μC₇PIPēPSPSC- PjP_μC, P°P±PsP»C-C†PSC-
C€P»C₁C...Pē PrP»C₁ C₁PjP_μPeP»C-PrPēPSC-C₁PjC₁ C^{''} PsPePēC₁PSPēPjPē
(PiC-PrC₇PsPeC₁PēP»C₇PIP°PSPSC₁, Ph-PrP_μP°P»PeC-P»C₁PIP°PSPSC₁) P·
P_iPsPrP°P»CHC€PsC₇ PePsPSB₇TMC₇PiP°C†C-C^{''}C₇ (PiP»C₇PeC₁C₇PsPSC-P·P°C†C-C₁
C, PsC%₀Ps), C%₀Ps C₁PiC₇PēC†PēPSC₁C^{''} C₁C, PIPsC₇P_μPSPSC₁ PSPēP·PePē
PjP_μC, P°P±PsP»C-C, C-PI P°P±Ps P·C- P·PSPēP¶P_μPSPsC₇ C, P°C₇PjP°PePsP»PsPiC-C†PSPsC₇
P°PeC, PēPIPS-C₁C, C₇, P°P±Ps P· PSP_μPIC₁C, P°PSPsPIP»P_μPSPsC₇
C, P°C₇PjP°PePsP»PsPiC-C†PSPsC₇ P°PeC, PēPIPS-C₁C, C₇. P₇PēC₁C, P_μPjPSPēPN₂ PIP_iP»PēPI
PjP_μC, P°P±PsP»C-C, C-PI C^{''} PSPēP·CHPePēPj.

P'C-P»P°PSC, P_μC₇PsP»

P^{''}PsC₁P»C-PrP¶P_μPSPSC₁ C-*n vitro* P_iPsPeP°P·P°P»Pē, C%₀Ps PIC-P»P°PSC, P_μC₇PsP»
PiPsP»PsPIPSpēPj C†PēPSPsPj PjP_μC, P°P±PsP»C-P·C₁C^{''}C, CHC₁C₁ C†PēC, PsC...C₇PsPjPsPj P 450
3P₄ (CYP3A4) C- C^{''} C₁C₁P±C₁C, C₇P°C, PsPj PrP»C₁ P_iP_μC₇P_μPSPsC₁PSPēPeP° P-gp.
PhC₁PSPsPIPSpēPjPē PjP_μC, P°P±PsP»C-C†PSPēPjPē C€P»C₁C...P°PjPē PrP»C₁
PIC-P»P°PSC, P_μC₇PsP»C₁ C^{''} Ph-PrP_μP°P»PeC-P»C₁PIP°PSPSC₁ P· P_iPsPrP°P»CHC€PēPj
C₁C, PIPsC₇P_μPSPSC₁Pj C₇C₁PrC₁ PjP_μC, P°P±PsP»C-C, C-PI P·C- P·PSP°C†PSPs
P·PjP_μPSC€P_μPSPsC₇ P°PeC, PēPIPS-C₁C, C₇ C%₀PsPrPs P±P_μC, P°₁- C- P±P_μC, P°₂-
P°PrC₇P_μPSPsC₇P_μC†P_μPiC, PsC₇C-PI. P_μP»P°P·PjPsPIPēPN₂ PjP_μC, P°P±PsP»C-C†PSPēPN₂
PiC₇PsC, C-P»CH PiC-C₁P»C₁ P_iP_μC₇PsC₇P°P»CHPSPsPiPs P·P°C₁C, PsC₁C₁PIP°PSPSC₁
PIC-P»P°PSC, P_μC₇PsP»C₁ PI C₇P°PrC-PsC-P·PsC, PsP_iPSPsPjC₁ PrPsC₁P»C-PrP¶P_μPSPSC- C₁
P»C₇PrPēPSPē PIC-PrP_iPsPIC-PrP°C^{''} PIPēC₁PsPePsPjC₁ PjP_μC, P°P±PsP»C-P·PjC₁
P_iP_μC₇C€PsPiPs PiC₇PsC...PsPrP¶P_μPSPSC₁. P₇PēC₁C, P_μPjPSPēPN₂ PIP_iP»PēPI
PjP_μC, P°P±PsP»C-C, C-PI C^{''} PSPēP·CHPePēPj.

P'PēPIP_μPrP_μPSPSC₁

PjPjP_μPeP»C-PrPēPSC-C₁Pj

P_μP»P°P·PjPsPIPēPN₂ PeP»C-C₇P_μPSC₁ PiC-C₁P»C₁ PIPSC₁C, C₇C-C€PSC₇PsPIP_μPSPSPsPiPs
PIP_μPrP_μPSPSC₁ C₁C, P°PSPsPIPēPI 151 P»C-C, C₇/PiPsPrPēPSC₁. P_μC-C₁P»C₁
PIPSC₁C, C₇C-C€PSC₇PsPIP_μPSPSPsPiPs PIP_μPrP_μPSPSC₁ PiC₇PēP±P»PēP·PSPs 58 %

PIPIPμPrPμPSPsC— PjC-C†PμPSPsC— PrPsP·Pë (P°P±Ps 73 % PIPëCΠIP»PμPSPsC—
 CḡP°PrC-PsP°PeC,PëPIPSsC†C,C-) PIPëPIPsPrPëP»PëC†CΠ P· PsCḡPiP°PSC-P·PjC† P·
 C„PμPeP°P»C-CΠPjPë C†PμCḡPμP· 192 PiPsPrPëPSPë PïC-C†P»CΠ PIPμPrPμPSPSCΠ PrPsP·Pë.
 P†P· C†PμC†PμCḡ PIPëPIPsPrPëC,CḡC†CΠ 22 % PIPμPrPμPSPsC— PjC-C†PμPSPsC— PrPsP·Pë
 C†PμCḡPμP· 168 PiPsPrPëPS (27 % PIPëCΠIP»PμPSPsC— CḡP°PrC-PsP°PeC,PëPIPSsC†C,C-).
 P°PëPIPμPrPμPSPSCΠ CḡPμC†PsPIPëPSPë, PïPsPIBḡ™ CΠP·P°PSPsC— P· P»C-PeP°CḡC†CḡPePëPj
 P·P°C†PsP±PsPj, P· C„PμPeP°P»C-CΠPjPë PïC-C†P»CΠ PIPSC†C,CḡC-CëPSCḡPsPIPμPSPSPsPiPs
 PIPμPrPμPSPSCΠ C†PIC-PrC†PëC,Cḡ PïCḡPs C†PμPeCḡPμC†C-Cḡ C† P¶PsPIC†. Pj CḡP°P·C-
 PïPμCḡPsCḡP°P»CḡPSPsPiPs P·P°C†C,PsC†C†PIP°PSPSCΠ C† P·PrPsCḡPsPIPëC...
 C†PsP»PsPIC-PeC-PI-PrPsP±CḡPsPIPsP»CḡC†C-PI P·P°PiP°P»CḡPSP°
 CḡP°PrC-PsP°PeC,PëPIPSsC-C†C,Cḡ PIPëPIPsPrPëP»P°C†CΠ P· PsCḡPiP°PSC-P·PjC†
 PïPμCḡPμPIP°P¶PSPs P· C„PμPeP°P»C-CΠPjPë (92 % PIPμPrPμPSPsC— PjC-C†PμPSPsC—
 PrPsP·Pë P°P±Ps 99 % PIPëCΠIP»PμPSPsC— CḡP°PrC-PsP°PeC,PëPIPSsC†C,C-) C†PμCḡPμP·
 168 PiPsPrPëPS PïC-C†P»CΠ PïCḡPëPμPsPjC† PrPsP·Pë. PḡPμPSCëPμ 1 %
 PïPμCḡPsCḡP°P»CḡPSPs PIPμPrPμPSPsC— PrPsP·Pë (1 % PIPëCΠIP»PμPSPsC—
 CḡP°PrC-PsP°PeC,PëPIPSsC†C,C-) PIPëPIPsPrPëP»PsC†CΠ P· PsCḡPiP°PSC-P·PjC† C-P·
 C†PμC†PμCḡ, C°Ps C†PIC-PrC†PëC,Cḡ PïCḡPs PSPμP·PSP°C†PSC† P°P±C†PsCḡP±C†C-Cḡ
 PïC-C†P»CΠ PïPμCḡPsCḡP°P»CḡPSPsPiPs P·P°C†C,PsC†C†PIP°PSPSCΠ. PμPμCḡC-PsPr
 PSP°PïC-PIPIPëPIPμPrPμPSPSCΠ C†PjPμPeP»C-PrPëPSC-C†PjC† PïC-C†P»CΠ PIPμPrPμPSPSCΠ
 C† PIPëPiP»CΠPrC- C-PSPiP°P»CΠC†C-C— PïCḡPsC,CΠPiPsPj 10 PrPSC-PI C†C,P°PSPsPIPëPI C†
 C†PμCḡPμPrPSCḡPsPjC† 19 PiPsPrPëPS C† P·PrPsCḡPsPIPëC... PrPsP±CḡPsPIPsP»CḡC†C-PI,
 PïCḡPëC†PsPjC† 3Bḡ“4 % PIPëPIPsPrPëP»PsC†CΠ P· PsCḡPiP°PSC-P·PjC† C†
 PSPμP·PjC-PSPμPSPsPjC† PIPëPiP»CΠPrC- C-P· C†PμC†PμCḡ C† CḡC-PIPSsPIP°P¶PSPsPjC†
 C†C,P°PSC-.

P°C-P»P°PSC,PμCḡPsP»

PμP»P°P·PjPsPIPëPμ PeP»C-CḡPμPSC† PIC-P»P°PSC,PμCḡPsP»C† PïC-C†P»CΠ
 PIPSC†C,CḡC-CëPSCḡPsPIPμPSPSPsPiPs PIPμPrPμPSPSCΠ C†C,P°PSPsPIPëPI
 108 P»C-C,CḡC-PI/PiPsPrPëPSC†. PμC-C†P»CΠ PïPμCḡPsCḡP°P»CḡPSPsPiPs
 PIPμPrPμPSPSCΠ PjC-C†PμPSPsPiPs PIC-P»P°PSC,PμCḡPsP»C† P±C†P»Ps PIPëCΠIP»PμPSPs
 70 % PjC-C†PμPSPsC— PrPsP·Pë C† C†PμC†C- C- 30 % Bḡ” C† C„PμPeP°P»C-CΠC....
 PμPμCḡPIPëPSPSPëPj PIPëPIPμPrPμPSPSCΠPj PIC-P»P°PSC,PμCḡPsP»C† C”
 PjPμC,P°P±PsP»C-P·Pj C-P· PSP°C†C,C†PïPSPëPj PIPëPrC-P»PμPSPSCΠPj
 PjPμC,P°P±PsP»C-C,C-PI C-P· C†PμC†PμCḡ C- C„PμPeP°P»C-CΠPjPë. PμPμCḡC-PsPr
 PSP°PïC-PIPIPëPIPμPrPμPSPSCΠ PIC-P»P°PSC,PμCḡPsP»C† PïC-C†P»CΠ
 PIPμPrPμPSPSCΠ C† PIPëPiP»CΠPrC- C-PSPiP°P»CΠC†C-C— PïCḡPsC,CΠPiPsPj 10
 PrPSC-PI C†C,P°PSPsPIPëPI C† C†PμCḡPμPrPSCḡPsPjC† 11 PiPsPrPëPS.

PḡC†PsP±P»PëPIC- PïCḡC†PïPë PïP°C†C-C”PSC,C-PI B

PμP°C†C-C”PSC,Pë P»C-C,PSCḡPsPiPs PIC-PeCé (> 65 CḡPsPeC-PI).
 PμP°CḡPjP°PePsPeC-PSPμC,PëC†PSPëPμ P°PSP°P»C-P· PïPsPïC†P»CΠC†C-C— PïPsPeP°P·P°PI,
 C°Ps C„P°CḡPjP°PePsPeC-PSPμC,PëPeP° C†PjPμPeP»C-PrPëPSC-C†PjC† C-
 PIC-P»P°PSC,PμCḡPsP»C† P±C†P»P° C†C...PsP¶P° C† PïP°C†C-C”PSC,C-PI P· P¶PḡP—P»
 PIC-PePsPj PIC-Pr 65 CḡPsPeC-PI C- PïP°C†C-C”PSC,C-PI PIC-PePsPj PrPs 65 CḡPsPeC-PI.

PμPsCḡCéCëPμPSPSCΠ C„CéPSPeC†C-C— PSPëCḡPsPe. Pj PïP°C†C-C”PSC,C-PI P·
 C,CΠP¶PePsCḡ PSPëCḡPePsPIPsCḡ PSPμPrPsC†C,P°C,PSC-C†C,Cḡ PSPμ P±C†P»Ps
 PIPëCΠIP»PμPSPs P¶PsPrPSPëC... PsP·PSP°Pe P·P±C-P»CḡCëPμPSPSCΠ
 C†PëC†C,PμPjPSPsPiPs PIP°P»PëPIC† C†PjPμPeP»C-PrPëPSC-C†PjC† P°P±Ps

PIC-P»P°PSC,PµCḡPsP»Ć (C_{max} C- AUC) P̄C-ĆP»Cḡ PIPµPrPµPSPSCḡ
ĆPjPµPeP»C-PrP̄PSC-ĆPjĆ/PIC-P»P°PSC,PµCḡPsP»Ć, PrPµ PrPsP·P°
ĆPjPµPeP»C-PrP̄PSC-ĆPjĆ P̄PµCḡPµPIP̄C%oĆPIP°P»P° CḡPµPePsPjPµPSPPrPsPIP°PSĆ
PrPsP·Ć Ć PrPIP° CḡP°P·P̄, P° PrPsP·P° PIC-P»P°PSC,PµCḡPsP»Ć PIC-PrP̄PsPIC-PrP°P»P°
CḡPµPePsPjPµPSPPrPsPIP°PSC-PN̄ PrPsP·C-. P—PIbḡ™ CḡP·ĆPIP°PSPSCḡ P· P±C-P»PeP°PjP̄
P̄P»P°P·PjP̄ P·P°P»P̄C̄P°P»PsĆCḡ PSPµP·PjC-PSPSP̄Pj P̄PsCḡC-PIPSCḡPSPs P·C-
P·PrPsCḡPsPIP̄PjP̄ PrPsP±CḡPsPIP̄sP»CHC†CḡPjP̄.

PµPsCḡC̄C̄PµPSPSCḡ C,,C̄PSP̄C†C-C— P̄PµC†C-PSP̄P̄. Pj P̄P°C†C-C”PSC,C-PI P·
P̄PsPjC-CḡPSP̄PjP̄ P̄PsCḡC̄C̄PµPSPSCḡPjP̄ C,,C̄PSP̄C†C-C— P̄PµC†C-PSP̄P̄ (PeP»P°Ć B
P·P° PeP»P°ĆP̄C,,C-PeP°C†C-C”Cḡ P̄P°PN̄P»PrP° Bḡ” P̄Pbḡ™ Cḡ) PSPµ
ĆP̄PsĆC,PµCḡC-PiP°P»PsĆCḡ PḡPsPrPSP̄C... PsP·PSP°Pe P·P±C-P»CHC̄PµPSPSCḡ
ĆP̄C̄C,PµPjPSPsPiPs PIP̄P»P̄PIĆ ĆPjPµPeP»C-PrP̄PSC-ĆPjĆ P°P±Ps
PIC-P»P°PSC,PµCḡPsP»Ć (C_{max} C- AUC) P̄C-ĆP»Cḡ PIPµPrPµPSPSCḡ
ĆPjPµPeP»C-PrP̄PSC-ĆPjĆ/PIC-P»P°PSC,PµCḡPsP»Ć, PrPµ PrPsP·P°
ĆPjPµPeP»C-PrP̄PSC-ĆPjĆ P̄PµCḡPµPIP̄C%oĆPIP°P»P° CḡPµPePsPjPµPSPPrPsPIP°PSĆ
PrPsP·Ć Ć PrPIP° CḡP°P·P̄, P° PrPsP·P° PIC-P»P°PSC,PµCḡPsP»Ć PIC-PrP̄PsPIC-PrP°P»P°
CḡPµPePsPjPµPSPPrPsPIP°PSC-PN̄ PrPsP·C-. P—PIbḡ™ CḡP·ĆPIP°PSPSCḡ P· P±C-P»PeP°PjP̄
P̄P»P°P·PjP̄ P·P°P»P̄C̄P°P»PsĆCḡ PSPµP·PjC-PSPSP̄Pj P̄PsCḡC-PIPSCḡPSPs P·C-
P·PrPsCḡPsPIP̄PjP̄ PrPsP±CḡPsPIP̄sP»CHC†CḡPjP̄. P—P°ĆC,P̄ĆC̄C̄PIP°PSPSCḡ
ĆPjPµPeP»C-PrP̄PSC-ĆPjĆ/PIC-P»P°PSC,PµCḡPsP»Ć PSPµ PsC†C-PSCḡPIP°P»P̄ Ć
P̄P°C†C-C”PSC,C-PI P· C,CḡPḡPePsCḡ P̄PµC†C-PSP̄PsPIP̄sCḡ PSPµPrPsĆC,P°C,PSC-ĆC,Cḡ.

P†PSC̄C- PsĆPsP±P»P̄PIC- PiCḡC̄P̄P̄ B

P̄P°CḡPjP°PePsPeC-PSPµC,P̄C†PSP̄PN̄ P°PSP°P»C-P· P̄PsP̄C̄P»CḡC†C-C— P̄PsPeP°P·P°PI,
C%oPs PSPµ P̄PsC,CḡC-P±PSPs PḡPsPrPSPsPiPs PePsCḡP̄P̄C̄PIP°PSPSCḡ PrPsP·P̄
ĆPjPµPeP»C-PrP̄PSC-ĆPjĆ P°P±Ps PIC-P»P°PSC,PµCḡPsP»Ć P·P°P»PµPḡPSPs PIC-Pr
PIC-PeĆ, CḡP°ĆP̄, ĆC,P°C,C-, PIP̄PePsCḡP̄C̄C,P°PSPSCḡ C-PSPiP°P»CḡC†C-PN̄PSP̄C...
PePsCḡC,P̄PePsĆC,PµCḡPsC—PrC-PI P°P±Ps PjP°ĆP̄ C,C-P»P° P̄P°C†C-C”PSC,P°.
P”PsĆP»C-PrPḡPµPSPSCḡ, P̄CḡPsPIPµPrPµPSPµ Ć P̄PsPIC-P»CHPSP̄C...
PjPµC,P°P±PsP»C-P·P°C,P̄CḡC-PI CYP2D6, PSPµ PIP̄CḡPIP̄P̄P»Ps PḡPsPrPSP̄C... PsP·PSP°Pe
PeP»C-PSC-C†PSPs P·PSP°C†C̄C%oPsPiPs PµC,,PµPeC,Ć PiPµPSPµC,P̄C†PSPsPiPs
P̄PsP»C-PjPsCḡC,,C-P·PjĆ CYP2D6 C%oPsPrPs ĆP̄C̄C,PµPjPSPsPiPs PIP̄P»P̄PIĆ
ĆPjPµPeP»C-PrP̄PSC-ĆPjĆ. B

Р̄P»C-PSC-C†PSC- C...P°CḡP°PeC,PµCḡP̄C̄C,P̄PeP̄

P̄PsPeP°P·P°PSPSCḡ. P”P»Cḡ P̄C-PrC,CḡP̄PjC̄CḡC†PsC—
P±CḡPsPSC...PsP»C-C,P̄C†PSPsC— C,PµCḡP°P̄C-C— P· PjPµC,P̄Cḡ P̄PsP»PµPiC̄PµPSPSCḡ
ĆP̄PjP̄P̄C,P̄PjC-PI Ć PrPsCḡPsĆP»P̄C... P̄P°C†C-C”PSC,C-PI P· C...CḡPsPSC-C†PSP̄Pj
PsP±ĆC,CḡC̄PeC,P̄PIPSP̄Pj P·P°C...PIP̄CḡCḡPIP°PSPSCḡPj P»PµPiPµPSCḡ (P̄PhP—P°).

PµCḡPsC,P̄P̄PsPeP°P·P°PSPSCḡ. P̄C-PrPIP̄C%oPµPSP° C†C̄C,P»P̄PIC-ĆC,Cḡ PrPs
P°PeC,P̄PIPSP̄C... CḡPµC†PsPIP̄PS P°P±Ps PrPs P±C̄PrCḡ-CḡPePsC— P·
PrPsP̄PsPjC-PḡPSP̄C... CḡPµC†PsPIP̄PS.

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-PNë

PᄁP»C-PSC-C†PSPs P·PSP°C†CíC%oC- PIP·P°C" PjPsPrC-C— P· C-PSC€PëPjPë
P»C-PeP°CĤCÍCHPePëPjPë P·P°CÍPsP±P°PjPë, CIPeC- P·P°CÍC,PsCÍPsPICrCĤC,CH
PsPrPSPSc†P°CÍPSPs, PIPiP°P¶P°CĤC,CHCÍC CI PjP°P»PsPNëPjPsPIC-CĤPSPëPjPë P·
PsPiP»CIPIrC† PSP° PSPëP·CHPeC- PePsPSC†PµPSC,CĤP°C†C-C—
C†PjPµPeP»C-PrPëPSC-C†PjC†/PIC-P»P°PSC,PµCĤPsP»C† PI PiP»P°P·PjC- PeCĤPsPIC-
PiC-CÍP»CI C-PSPiP°P»CI†C-C-PNëPSPSc— PrPsP·Pë.

P'PµC,P°-P°PrCĤPµPSPsP±P»PsPeP°C,PsCĤPë

P>C-PeP°CĤCÍCHPeC- P·P°CÍPsP±Pë, C%oPs PjC-CÍC,CIC,CH P±PµC,P°-
P°PrCĤPµPSPsP±P»PsPeP°C,PsCĤPë, PjPsP¶C†C,CH PiPsCÍP»P°P±PëC,Pë P°P±Ps
P°PSC,P°PiPsPSC-P·C†PIP°C,Pë PµC,,PµPeC, P±PµC,P°₂-P°PrCĤPµPSPµCĤPiC-C†PSPëC...
P°PiPsPSC-CÍC,C-PI, C,P°PePëC... CIPIe PIC-P»P°PSC,PµCĤPsP». PŸP»C-Pr C†PSPëPeP°C,Pë
PsPrPSPSc†P°CÍPSPsPiPs P·P°CÍC,PsCÍC†PIP°PSPSC CIPIe PSPµCÍPµP»PµPeC,PëPIPSPëC...,
C,P°Pe C- CÍPµP»PµPeC,PëPIPSPëC... P±PµC,P°-P°PrCĤPµPSPsP±P»PsPeP°C,PsCĤC-PI,
CIPIeC%oPs C,C-P»CHPePë PSPµPjP°C" PIP°PiPsPjPëC... PiCĤPëC†PëPS PrP»CI C-C...
P·P°CÍC,PsCÍC†PIP°PSPSC.

*P'P·P°C" PjPsPrC-C— PSP° PsCÍPSPsPIC- PjPµC,P°P±PsP»C-P·PjCž C-
C,CĤP°PSCÍPiPsCĤC,CžPIP°PSPSC*

P'C-P»P°PSC,PµCĤPsP» C" CÍC†P±CÍC,CĤP°C,PsPj C†PëC,PsC...CĤPsPjC† P450 3A4 (CYP3A4).
PhPrPSPSc†P°CÍPSPµ P·P°CÍC,PsCÍC†PIP°PSPSC CI†PëP»CHPSPsPrC-CĤC†PëC...
C-PSPiC-P±C-C,PsCĤC-PI CYP3A4 (C,P°PePëC... CIPIe PePµC,PsPePsPSP°P·PsP»,
PeP»P°CĤPëC,CĤPsPjC-C†PëPS, C-C,CĤP°PePsPSP°P·PsP», CĤPëC,PsPSP°PIC-CĤ,
C,PµP»C-C,CĤPsPjC-C†PëPS) PjPsP¶Pµ PiCĤPëPiPSC-C†C†PIP°C,Pë PjPµC,P°P±PsP»C-P·Pj C-
P·P±C-P»CHC€C†PIP°C,Pë CÍPëCÍC,PµPjPSPëPNë PIPiP»PëPI PIC-P»P°PSC,PµCĤPsP»C†.
PhPrPSPSc†P°CÍPSPµ P·P°CÍC,PsCÍC†PIP°PSPSC P· PePµC,PsPePsPSP°P·PsP»PsPj (400 PjPi) C†
P·PrPsCĤPsPIPëC... PrPsP±CĤPsPIPsP»CHC†C-PI P·P±C-P»CHC€C†PIP°P»Ps CÍPµCĤPµPrPSC-
PiPsPeP°P·PSPëPePë AUC_(0-8h) C- C_{max} PIC-P»P°PSC,PµCĤPsP»C† PSP° 65 % C- 22 %
PIC-PrPiPsPIC-PrPSPs. PµPsCÍPëP»PµPSPSC PrC-C— PIC-P»P°PSC,PµCĤPsP»C† PSPµ
P°CÍPsC†C-CĤHIP°P»PsCÍC CI P·C- P·P±C-P»CHC€PµPSPSC IPj C†PëCÍC,PµPjPSPëC...
PIPiP»PëPIC-PI, PiPsPIbĤ™ CIPI·P°PSPëC... C-P· P±PµC,P°-
P°PrCĤPµPSPsP±P»PsPeP°C,PsCĤP°PjPë, PSPµ PIPiP»PëPIP°P»Ps PSP° C†P°CÍC,PsC,C†
CÍPµCĤC†PµPIPëC... C†PePsCĤPsC†PµPSCĤ, CĤC-PIPIPSCH PeP°P»C-CĤ C† PeCĤPsPIC-
P°P±Ps C-PSC,PµCĤHIP°P» QT (C†PePsCĤPëPiPsPIP°PSPëPNë P·P° C,,PsCĤPjC†P»PsCĤ
PαCĤPµPrPµCĤC-PeP°). PŸP»C-Pr P±C†C,Pë PsP±PµCĤPµP¶PSPëPj PiCĤPë
PsPrPSPSc†P°CÍPSPsPjC† P·P°CÍC,PsCÍC†PIP°PSPSC-
C†PjPµPeP»C-PrPëPSC-C†PjC†/PIC-P»P°PSC,PµCĤPsP»C† P· PePµC,PsPePsPSP°P·PsP»PsPj C,P°
C-PSC€PëPjPë PIC-PrPsPjPëPjPë C†PëP»CHPSPsPrC-CĤC†PëPjPë C-PSPiC-P±C-C,PsCĤP°PjPë
CYP3A4, PsCÍPeC-P»CHPePë C" PiPsC,PµPSC†C-P°P» PrP»CI P·P±C-P»CHC€PµPSPSC CI
C†PëCÍC,PµPjPSPsPiPs PIPiP»PëPIC† PIC-P»P°PSC,PµCĤPsP»C†, C%oPs PjPsP¶Pµ
PiC-PrPIPëC%oPëC,Pë CĤPëP·PëPe CĤPsP·PIPëC,PeC† PSPµP±P°P¶P°PSPëC... CĤPµP°PeC†C-PNë.
P'PµCĤP°PiP°PjC-P», PiPsPjC-CĤPSPëPNë C-PSPiC-P±C-C,PsCĤ CYP3A4, C-CÍC,PsC,PSPs PSPµ
PIPiP»PëPIP°C" PSP° C,,P°CĤPjP°PePsPeC-PSPµC,PëPeC† PIC-P»P°PSC,PµCĤPsP»C†.

PjPjPµPeP»C-PrPëPSC-C†Pj C" CÍC†P±CÍC,CĤP°C,PsPj C†PëC,PsC...CĤPsPjC† P450 2D6
(CYP2D6). PαP°CĤPjP°PePsPeC-PSPµC,PëPeC† C†PjPµPeP»C-PrPëPSC-C†PjC† C†
CĤC-PIPSPIPIP°P¶PSPsPjC† CÍC,P°PSC- PsC†C-PSCĤHIP°P»Pë C† P·PrPsCĤPsPIPëC...

PrPsP±CḡPsPIPsP»CHC†C-PI, CḡPeC- PjP°CHC,CH PrPμC,,C-C†PëC, CYP2D6
(PiPsPIC-P»CHPSC- PjPμC,P°P±PsP»C-P·P°C,PsCḡPë). PkPμ CḡPiPsCḡC,PμCḡC-PiP°P»PsCḡCḡ
P¶PsPrPSPsPiPs PiPiP»PëPICr PSP° AUC P°P±Ps C_{max} CrPjPμPeP»C-PrPëPSC-CrPjCr PiCḡPë
PrPsP·C-, C%Ps PiPμCḡPμPIPëC%CrC” PSPsCḡPjCr Cr 8 CḡP°P·C-PI. PμCḡPëP±P»PëP·PSPs 1,3-
PeCḡP°C,PSPμ P·P±C-P»CHC€PμPSPSCḡ AUC CrPjPμPeP»C-PrPëPSC-CrPjCr
CḡPiPsCḡC,PμCḡC-PiP°P»PsCḡCḡ PiCḡPë PrPsP·C-, C%Ps PiPμCḡPμPIPëC%CrC” PSPsCḡPjCr
Cr 16 CḡP°P·C-PI, P±PμP· PiPiP»PëPICr PSP° C_{max} CrPjPμPeP»C-PrPëPSC-CrPjCr. P—
PsPiP»CḡPrCr PSP° PiPμP»PëC†PëPSCr C†PëC... P·PjC-PS PSPμ PsC†C-PeCrC”C,CHCḡCḡ
PeP»C-PSC-C†PSPs P·PSP°C†CrC%PëC... P»C-PeP°CḡCḡCHPePëC... PiP·P°C”PjPsPrC-PNë PiCḡPë
PsPrPSPsC†P°CḡPSPsPjCr PiPiPμPrPμPSPSC-
CrPjPμPeP»C-PrPëPSC-CrPjCr/PIC-P»P°PSC,PμCḡPsP»Cr P· C-PSPiC-P±C-C,PsCḡP°PjPë CYP2D6
P°P±Ps PiCḡPë PiPiPμPrPμPSPSC- PiP°C†C-C”PSC,P°Pj P· PiPμPSPμC,PëC†PSPs
PrPμC,,C-C†PëC,PSPsCḡ P°PeC,PëPiPSC-CḡC,Cḡ CYP2D6 (PiPsPIC-P»CHPSC-
PjPμC,P°P±PsP»C-P·P°C,PsCḡPë).

PiPe CrPjPμPeP»C-PrPëPSC-CrPj, C,P°Pe C- PIC-P»P°PSC,PμCḡPsP» C”
CḡCrP±CḡC,CḡP°C,P°PjPë PiPμCḡPμPSPsCḡPSPëPeP° P -PiP»C-PePsPiCḡPsC,PμC—PSCr (P-gp).
P’PiP»PëPi PiPsPjC-CḡPSPsPiPs P-gp C-PSPiC-P±C-C,PsCḡCr PiPμCḡP°PiP°PjC-P»Cr (240 PjPi
PsPrPëPS CḡP°P· PSP° PrPsP±Cr) PSP° C,,P°CḡPjP°PePsPeC-PSPμC,PëPeCr
CrPjPμPeP»C-PrPëPSC-CrPjCr C- PIC-P»P°PSC,PμCḡPsP»Cr Cr CḡC-PIPSsPiP°P¶PSPsPjCr
CḡC,P°PSC- PsC†C-PSCḡPiP°P»Pë Cr P·PrPsCḡPsPiPëC... PrPsP±CḡPsPIPsP»CHC†C-PI. PkPμ
P±CrP»Ps PIC-PrPjC-C†PμPSPs P±CrPrCH-CḡPePsPiPs PiPiP»PëPICr PiPμCḡP°PiP°PjC-P»Cr
PSP° C_{max} CrPjPμPeP»C-PrPëPSC-CrPjCr P°P±Ps PIC-P»P°PSC,PμCḡPsP»Cr.
PŸPiPsCḡC,PμCḡC-PiP°P»PsCḡCḡ PiCḡPëP±P»PëP·PSPs 1,4-PeCḡP°C,PSPμ
P·P±C-P»CHC€PμPSPSCḡ AUC CrPjPμPeP»C-PrPëPSC-CrPjCr P·P° PIC-PrCḡCrC,PSPsCḡC,C-
PiPiP»PëPICr PSP° AUC PIC-P»P°PSC,PμCḡPsP»Cr. P— PsPiP»CḡPrCr PSP° PiPμP»PëC†PëPSCr
C†PëC... P·PjC-PS PSPμ PsC†C-PeCrC”C,CHCḡCḡ PeP»C-PSC-C†PSPs P·PSP°C†CrC%PëC...
P»C-PeP°CḡCḡCHPePëC... PiP·P°C”PjPsPrC-PNë PiCḡPë PsPrPSPsC†P°CḡPSPsPjCr
PiPiPμPrPμPSPSC- CrPjPμPeP»C-PrPëPSC-CrPjCr/PIC-P»P°PSC,PμCḡPsP»Cr P·
C-PSPiC-P±C-C,PsCḡP°PjPë P-gp.

P†PSC€C- P°PSC,PëPjC†CḡPeP°CḡPëPSPsPIC- P·P°CḡPsP±Pë C- CḡPëPjPiP°C,PsPjC-PjPμC,PëPePë

PkPrPSPsC†P°CḡPSPμ P·P°CḡC,PsCḡCrPiP°PSPSCḡ
CrPjPμPeP»C-PrPëPSC-CrPjCr/PIC-P»P°PSC,PμCḡPsP»Cr P· C-PSC€PëPjPë
PjCrCḡPeP°CḡPëPSPsPiPëPjPë P°PSC,P°PiPsPSC-CḡC,P°PjPë C,CḡPëPiP°P»PsC— PrC-C—,
P±PμC,P°₂-P°PrCḡPμPSPμCḡPiC-C†PSPëPjPë P°PiPsPSC-CḡC,P°PjPë C,CḡPëPiP°P»PsC— PrC-C—
P°P±Ps P»C-PeP°CḡCḡCHPePëPjPë P·P°CḡPsP±P°PjPë, C%Ps PjC-CḡC,CḡC,CH P±CrPrCH-
CḡPePëPNë P· C†PëC... P·P°CḡPsP±C-PI, PSPμ PrPsCḡP»C-PrP¶CrPiP°P»Pë. PŸP°PePμ
P·P°CḡC,PsCḡCrPiP°PSPSCḡ PSPμ CḡPμPePsPjPμPSPrCrC”C,CHCḡCḡ, PsCḡPeC-P»CHPePë C†Pμ
PjPsP¶Pμ PiPsCḡPëP»PëC,Pë PIC-PrPsPjC- PiPsP±C-C†PSC- CḡPμP°PeC†C-C—
C-PSPiP°P»CḡC†C-PNëPSPëC... PjCrCḡPeP°CḡPëPSPsPiPëC... P°PSC,P°PiPsPSC-CḡC,C-PI P°P±Ps
P±PμC,P°₂-P°PrCḡPμPSPμCḡPiC-C†PSPëC... P°PiPsPSC-CḡC,C-PI (PrPëPi. CḡPsP·PrC-P»Pë
B«PkCḡPsP±P»PëPiPsCḡC,C- P·P°CḡC,PsCḡCrPiP°PSPSCḡB» C,P°
B«PμPμCḡPμPrPsP·CrPiP°PSPSCḡB»).

P”C-PiPsPeP°P»C-C”PjC-Cḡ

PŸCrPiCrC,PSC” PiC-PiPsPeP°P»C-C”PjC-C†PSPμ P»C-PeCrPiP°PSPSCḡ C-P·
P·P°CḡC,PsCḡCrPiP°PSPSCḡPj PiPsC...C-PrPSPëC... PjPμC,PëP»PeCḡP°PSC,PëPSCr,
CḡC,PμCḡPsC—PrC-PI P°P±Ps PSPμPeP°P»C-PNëP·P±PμCḡC-PiP°CHC†PëC...
PrC-CrCḡPμC,PëPeC-PI PjPsP¶Pμ PiPsCḡPëP»CHPiP°C,Pë PiPsC,PμPSC†C-PNëPSPëPNë

PiC-PiPsPeP°P»C-C"PjC-C†PSPëPNë PµC,,PµPeC, P±PµC,P°₂-P°PrCḡPµPSPµCḡPiC-C†PSPëC...
P°PiPsPSC-CÍC,C-PI, C,PsPjCí CíPjPµPeP»C-PrPëPSC-CíPj/PIC-P»P°PSC,PµCḡPsP» CÍP»C-Pr
P·P°CÍC,PsCÍPsPICíPIP°C,Pë P· PsP±PµCḡPµP¶PSC-CÍC,Cḡ (PrPëPI. CḡPsP·PrC-P»
B«PŸPiPsCÍC-P± P·P°CÍC,PsCÍCíPIP°PSPSCÍ C,P° PrPsP·PëB»).

P†PSCëC- P»C-PeP°CḡCÍCHPeC- P·P°CÍPsP±Pë PrP»CÍ P»C-PeCáPIP°PSPSCÍ PÍPhP-P»

PÍPsC±P° P¶PsPrPSPëC... C,CḡP°PrPëC†C-PNëPSPëC... PrPsCÍP»C-PrP¶PµPSCḡ
PIP·P°C"PjPsPrC-C— P»C-PeP°CḡCÍCHPePëC... P·P°CÍPsP±C-PI *in vivo* PSPµ P±CíP»Ps
PiCḡPsPIPµPrPµPSPs, C-PSPiP°P»CÍC†C-PNëPSPëPNë P·P°CÍC-P±
CíPjPµPeP»C-PrPëPSC-CíPj/PIC-P»P°PSC,PµCḡPsP» P·P°CÍC,PsCÍPsPICíPIP°P»Pë
PsPrPSPsC±P°CÍPSPs P· C-PSCëPëPjPë P»C-PeP°CḡCÍCHPePëPjPë P·P°CÍPsP±P°PjPë
PiCḡPsC,Pë PÍPhP-P», PIPEP»CḡC±P°CḡC±Pë CÍPëPjPiP°C,PsPjC-PjPµC,PëC±PSC-
P±CḡPsPSC...PsP»C-C,PëPePë PePsCḡPsC,PePsC— PrC-C— C,P° C-PSPiP°P»CÍC†C-PNëPSC-
PePsCḡC,PëPePsCÍC,PµCḡPsC—PrPë, P±PµP· PeP»C-PSC-C†PSPëC... PiCḡPsCÍPIC-PI
P»C-PeP°CḡCÍCHPePsC— PIP·P°C"PjPsPrC-C—.

B

PhCÍPsP±P»PëPIPsCÍC,C- P·P°CÍC,PsCÍCáPIP°PSPSCÍ

PḡCÍC,PjP°

PjPjPµPeP»C-PrPëPSC-CíPj/PIC-P»P°PSC,PµCḡPsP» PSPµ CÍP»C-Pr
P·P°CÍC,PsCÍPsPICíPIP°C,Pë PiP°C†C-C"PSC,P°Pj P· P°CÍC,PjPsCḡ, PsCÍPeC-P»CHPePë C†PµPNë
P»C-PeP°CḡCÍCHPePëPNë P·P°CÍC-P± PSPµ PrPsCÍP»C-PrP¶CíPIP°PICÍCÍ PrP»CÍ C†C-C"C—
PiCḡCíPiPë PiP°C†C-C"PSC,C-PI.

PµP°CḡP°PrPsPeCÍP°P»CHPSPëPNë P±CḡPsPSC...PsCÍPiP°P·Pj

PiPe C- PiCḡPë P·P°CÍC,PsCÍCíPIP°PSPSC- C-PSCëPëC... PIPëPrC-PI
C-PSPiP°P»CÍC†C-PNëPSPsC— C,PµCḡP°PiC-C—, P·P°CÍC,PsCÍCíPIP°PSPSCÍ
CíPjPµPeP»C-PrPëPSC-CíPjCí/PIC-P»P°PSC,PµCḡPsP»Cí PjPsP¶Pµ PiCḡPëP·PIPµCÍC,Pë PrPs
PiP°CḡP°PrPsPeCÍP°P»CHPSPsPiPs P±CḡPsPSC...PsCÍPiP°P·PjCí, CÍPePëPNë PjPsP¶Pµ
P±CíC,Pë PSPµP±PµP·PiPµC±PSPëPj PrP»CÍ P¶PëC,C,CÍ. PŸPµCḡP°PiC-Cḡ
CíPjPµPeP»C-PrPëPSC-CíPjPsPj/PIC-P»P°PSC,PµCḡPsP»PsPj CÍP»C-Pr PSPµPiP°PNëPSPs
PiCḡPëPiPëPSPëC,Pë Cí CḡP°P·C- CḡPsP·PIPëC,PeCí PiP°CḡP°PrPsPeCÍP°P»CHPSPsPiPs
P±CḡPsPSC...PsCÍPiP°P·PjCí C,P° CḡPsP·PiPsC±P°C,Pë P°P»CHC,PµCḡPSP°C,PëPIPsCí
C,PµCḡP°PiC-Cḡ, CÍPeC%oPs C†Pµ PiPsC,CḡC-P±PSPs.

PkPµ PrP»CÍ PIPëPePsCḡPëCÍC,P°PSPSCÍ PiC-Pr C†P°CÍ PiPsCÍC,CḡPsPiPs CÍC,P°PSCá

PjPjPµPeP»C-PrPëPSC-CíPj/PIC-P»P°PSC,PµCḡPsP» PSPµ PiPsPeP°P·P°PSPëPNë PrP»CÍ
P»C-PeCíPIP°PSPSCÍ PiPsCÍC,CḡPëC... PµPiC-P·PsPrC-PI P±CḡPsPSC...PsCÍPiP°P·PjCí.

PµPsPiC-CḡCëPµPSPSCÍ PiPµCḡPµP±C-PiCá P·P°C...PIPsCḡCHPIP°PSPSCÍ

PµC-PrPIPëC%oPµPSPSCÍ C±P°CÍC,PsC,Pë PIPëPePsCḡPëCÍC,P°PSPSCÍ
PePsCḡPsC,PePsPrC-CḡC±PëC... P±CḡPsPSC...PsP»C-C,PëPeC-PI PrP»CÍ
PiPsP»PµPiCëPµPSPSCÍ CÍPëPjPiC,PsPjC-PI PIPEP°P·CíC" PSP° PiPsPiC-CḡCëPµPSPSCÍ
PePsPSC,CḡPsP»Cḡ P·P°C...PIPsCḡCHPIP°PSPSCÍ. Pj CḡP°P·C- PiPsPiC-CḡCëPµPSPSCÍ
PiPµCḡPµP±C-PiCí PÍPhP-P» PSP° C,P»C- P»C-PeCíPIP°PSPSCÍ
CíPjPµPeP»C-PrPëPSC-CíPjPsPj/PIC-P»P°PSC,PµCḡPsP»PsPj PSPµPsP±C...C-PrPSPs

PiPsPIC,PsCḡPSPs PsC†C-PSPëC,Pë C†C,P°PS PiP°C†C-C"PS,C- C,P°PeC,PëPeC†
P»C-PeC†PIP°PSPSCḡ.

P'PiP»PëPI PSP° C†PµCḡC†PµPIPs-C†C¿PrPëPSPSC¿ C†PëC†C,PµPjC¿

P'PiP»PëPI PSP° C†PµCḡC†PµPIPs-C†C†PrPëPSPSC† C†PëC†C,PµPjC†, C,P°PePëPNë CḡPe
PiPsCḡC†C†PµPSPSCḡ C†PµCḡC†PµPIPsPiPs CḡPëC,PjC†, PSP°PiCḡPëPeP»P°Pr
PjPëPiPsC,P»PëPIP° P°CḡPëC,PjC-Cḡ C,P° C,P°C...C-PeP°CḡPrC-Cḡ, PjPsPḡP»PëPIPëPNë
PiC-C†P»Cḡ PIPµPrPµPSPSCḡ P°PSC,P°PiPsPSC-C†C,C-PI PjC†C†PeP°CḡPëPSPsPIPëC...
CḡPµC†PµPiC,PsCḡC-PI C- C†PëPjPiP°C,PsPjC-PjPµC,PëPeC-PI, PIPeP»CḡC†P°CḡC†Pë
C†PjPµPeP»C-PrPëPSC-C†Pj/PIC-P»P°PSC,PµCḡPsP» PµP°C†C-C"PS,C- P· PeP»C-PSC-C†PSPs
P·PSP°C†C†C%PëPj PSPµPePsPSC,CḡPsP»CHPsPIP°PSPëPj C†PµCḡC†PµPIPs-C†C†PrPëPSPSPëPj
P·P°C...PIPsCḡCḡPIP°PSPSCḡPj P±C†P»Pë PIPëPeP»CḡC†PµPSC- P· PeP»C-PSC-C†PSPëC...
PrPsC†P»C-PrPḡPµPSCḡ. PýPsPjC† C†PjPµPeP»C-PrPëPSC-C†Pj/PIC-P»P°PSC,PµCḡPsP»
C†P»C-Pr PiCḡPëP·PSP°C†P°C,Pë P· PsP±PµCḡPµPḡPSC-C†C,Cḡ PiP°C†C-C"PS,C- P°Pj P·
C,CḡPḡPePëPj C†PµCḡC†PµPIPs-C†C†PrPëPSPSPëPj P·P°C...PIPsCḡCḡPIP°PSPSCḡPj.

PḡPSC,PëPjC¿C†PeP°CḡPëPSPsPIP° P°PeC,PëPIPSC-C†C,CH

P— PsPiP»CḡPrC† PSP° P°PSC,PëPjC†C†PeP°CḡPëPSPsPIC† P°PeC,PëPIPSC-C†C,CH
C†PjPµPeP»C-PrPëPSC-C†Pj/PIC-P»P°PSC,PµCḡPsP» C†P»C-Pr P·P°C†C,PsC†PsPIC†PIP°C,Pë P·
PsP±PµCḡPµPḡPSC-C†C,Cḡ PiP°C†C-C"PS,C- P°Pj C-P· P·P°C,CḡPëPjPePsCḡ C†PµC†C- P°P±Ps
P·P°PeCḡPëC,PsPeC†C,PsIPsCḡ PiP»P°C†PePsPjPsCḡ.

P"C-PiPsPeP°P»C-C" PjC-Cḡ

P'PµC,P°₂-P°PrCḡPµPSPµCḡPiC-C†PSC- P°PiPsPSC-C†C,Pë PjPsPḡC†C,CH
C†PiCḡPëC†PëPSCḡC,Pë P·PSP°C†PSC† PiC-PiPsPeP°P»C-C" PjC-Cḡ C† PrPµCḡPePëC...
PiP°C†C-C"PS,C-PI, C%Ps PjPsPḡPµ PsP±C†PjPsPIPëC,Pë CḡPsP·PIPëC,PsPe
PiPsP±C-C†PSPëC... CḡPµP°PeC†C-PNë P· P±PsPeC† C†PµCḡC†PµPIPs-C†C†PrPëPSPSPsC—
C†PëC†C,PµPjPë. P—PjPµPSC€PµPSPSCḡ PeP°P»C-Cḡ PI C†PëCḡPsPIP°C,C†C- PeCḡPsPIC-,
CḡPe PiCḡP°PIPëP»Ps, C" C,CḡP°PSP·PëC,PsCḡPSPëPj.

P-PsPrPSPëC... PeP»C-PSC-C†PSPs P·PSP°C†C†C%PëC... PµC,,PµPeC,C-PI
PiC-PiPsPeP°P»C-C" PjC-C— PSPµ C†PiPsC†C,PµCḡC-PiP°P»PsC†Cḡ C†
PeP»C-PSC-C†PSPëC... PrPsC†P»C-PrPḡPµPSPSCḡC... C-P·
P·P°C†C,PsC†C†PIP°PSPSCḡPj C†PjPµPeP»C-PrPëPSC-C†PjC†/PIC-P»P°PSC,PµCḡPsP»C†
C† CḡPµPePsPjPµPSPPrPsPIP°PSC-PNë C,PµCḡP°PiPµPIC,PëC†PSC-PNë PrPsP·C-. PÝP»C-Pr
PiCḡPsCḡPIP»CḡC,Pë PsP±PµCḡPµPḡPSC-C†C,CH, PePsP»Pë
C†PjPµPeP»C-PrPëPSC-C†Pj/PIC-P»P°PSC,PµCḡPsP» P·P°C†C,PsC†PsPIC†CḡC,CH P·
C-PSC€PëPjPë P»C-PeP°CḡC†CHPePëPjPë P·P°C†PsP±P°PjPë, CḡPeC- C,P°PePsPḡ
PjPsPḡC†C,CH C†PiCḡPëCḡC,Pë CḡPsP·PIPëC,PeC† PiC-PiPsPeP°P»C-C" PjC-C— (PrPëPI.
CḡPsP·PrC-P» B«P'P·P°C" PjPsPrC-Cḡ P· C-PSC€PëPjPë P»C-PeP°CḡC†CHPePëPjPë
P·P°C†PsP±P°PjPë C,P° C-PSC€C- PIPëPrPë PIP·P°C" PjPsPrC-PNëB»).

P"C-PiPµCḡPiP»C-PePµPjC-Cḡ

P'PµC,P°₂-P°PrCḡPµPSPµCḡPiC-C†PSC- P°PiPsPSC-C†C,Pë PjPsPḡC†C,CH
C†PiCḡPëC†PëPSCḡC,Pë C,CḡP°PSP·PëC,PsCḡPSC† PiC-PiPµCḡPiP»C-PePµPjC-Cḡ C†
PrPµCḡPePëC... PiP°C†C-C"PS,C-PI.

P-PsPrPSPëC... PeP»C-PSC-C†PSPs P·PSP°C†CfC%PëC... PIPiP»PëPIC-PI PSP° CĤC-PIPμPSCĤ
PiP»CĤPePsP·Pë PI PïP»P°P·PjC- PSPμ CĤPïPsCĤC,PμCĤC-PiP°P»PsCĤCĤ Cř
PeP»C-PSC-C†PSPëC... PrPsCĤP»C-PrP¶PμPSPSCĤC... C-P· P·P°CĤC,PsCĤCřPIP°PSPSCĤPj
CřPjPμPeP»C-PrPëPSC-CřPjCř/PIC-P»P°PSC,PμCĤPsP»Cř Cř CĤPμPePsPjPμPSPPrPsPIP°PSC-PNë
C,PμCĤP°PiPμPIC,PëC†PSC-PNë PrPsP·C-. PμC-CĤP»CĤ PïPsC†P°C,PeCř P»C-PeCřPIP°PSPSCĤ
CřPjPμPeP»C-PrPëPSC-CřPjPsPj/PIC-P»P°PSC,PμCĤPsP»PsPj PSPμPsP±C...C-PrPSPs
P±C-P»CHC€ CĤPμC,PμP»CHPSPs PePsPSC,CĤPsP»CĤPIP°C,Pë CĤC-PIPμPSCĤ
PiP»CĤPePsP·Pë PI PïP»P°P·PjC- PeCĤPsPIC- Cř C...PIPsCĤPëC... PSP° C†CřPeCĤPsPIPëPNë
PrC-P°P±PμC,.

PŸCžPìCžC,PSC- P·P°C...PIPsCĤCĤPIP°PSPSCĤ

PjPjPμPeP»C-PrPëPSC-CřPj/PIC-P»P°PSC,PμCĤPsP» CĤP»C-Pr P·P°CĤC,PsCĤPsPICřPIP°C,Pë P·
PsP±PμCĤPμP¶PSC-CĤC,CĤ PïP°C†C-C°PSC,P°Pj C-P· CĤCřPrPsPjPSPëPjPë CĤPsP·P»P°PrP°PjPë
P°P±Ps P· C,PëCĤPμPsC,PsPeCĤPëPePsP·PsPj, P° C,P°PePsP¶ PïP°C†C-C°PSC,P°Pj, CĤPeC-
PSPμP·PIPëC†PSPs CĤPμP°PiCřCĤC,CH PSP° P±PμC,P°₂-P°PrCĤPμPSPμCĤPiC-C†PSC-
P°PiPsPSC-CĤC,Pë.

P°PsPïPsPjC-P¶PSC- CĤPμC†PsPIPëPSPë

PiPμPNë P»C-PeP°CĤCĤCĤPePëPNë P·P°CĤC-P± PjC-CĤC,PëC,CH P»P°PeC,PsP·Cř.
PμP°C†C-C°PSC,P°Pj C-P· CĤC-PrPeC-CĤPSPëPjPë CĤPïP°PrPePsPIPëPjPë C,,PsCĤPjP°PjPë
PSPμPiPμCĤPμPSPsCĤPëPjPsCĤC,C- PiP°P»P°PeC,PsP·Pë, PSPμPrPsCĤC,P°C,PSC-CĤC,CĤ
P»P°PeC,P°P·Pë P°P±Ps CĤPëPSPrCĤPsPjPsPj PiP»CĤPePsP·Ps-PiP°P»P°PeC,PsP·PSPsC—
PjP°P»CHP°P±CĤPsCĤP±C†C-C— PSPμ CĤP»C-Pr P·P°CĤC,PsCĤPsPICřPIP°C,Pë C†PμPNë
P»C-PeP°CĤCĤCĤPePëPNë P·P°CĤC-P±.

***P—P°CĤC,PsCĤCžPIP°PSPSCĤ Cž PïPμCĤC-PsPr PIP°PiC-C,PSPsCĤC,C- P°P±Ps
PiPsPrCžPIP°PSPSCĤ PiCĤCžPrPrCĤ***

P°P°PiC-C,PSC-CĤC,CH. PkP°CĤP°P·C- PIC-PrCĤCřC,PSC- PrP°PSC- PïCĤPs
P·P°CĤC,PsCĤCřPIP°PSPSCĤ CřPjPμPeP»C-PrPëPSC-CřPjCř/PIC-P»P°PSC,PμCĤPsP»Cř
PIP°PiC-C,PSPëPj P¶C-PSPeP°Pj.

PjPjPμPeP»C-PrPëPSC-CřPj/PIC-P»P°PSC,PμCĤPsP» CĤP»C-Pr P·P°CĤC,PsCĤPsPICřPIP°C,Pë
PïC-Pr C†P°CĤ PIP°PiC-C,PSPsCĤC,C-, C,C-P»CHPePë CĤPeC%Ps PsC†C-PeCřPIP°PSP°
PePsCĤPëCĤC,CH PrP»CĤ PIP°PiC-C,PSPsC— PïPμCĤPμPIPëC%CřC” PïPsC,PμPSC†C-PNëPSPëPNë
CĤPëP·PëPe PrP»CĤ PïP»PsPrP°.

PμPμCĤC-PsPr PiPsPrCřPIP°PSPSCĤ PiCĤCřPrPrCĤ. PkPμPIC-PrPsPjPs, C†Pë
PïPsC,CĤP°PiP»CĤC” CřPjPμPeP»C-PrPëPSC-CřPj P°P±Ps PIC-P»P°PSC,PμCĤPsP» Cř
PiCĤCřPrPSPμ PjPsP»PsPePs. PμCĤPsC,Pμ C-PSC€C- P±PμC,P°₂-P°PrCĤPμPSPμCĤPiC-C†PSC-
P°PiPsPSC-CĤC,Pë PIPëCĤPIP»CĤCĤC,CHCĤCĤ Cř PiCĤCřPrPSPsPjCř PjPsP»PsC†C-. P PëP·PëPe
PrP»CĤ PSPsPiPsPSP°CĤPsPrP¶PμPSPëC.../PSPμPjPsPIP»CĤC, PSPμ PjPsP¶PSP°
PIPëPeP»CĤC†P°C,Pë. PμPsC,CĤC-P±PSPs PiCĤPëPNëPSCĤC,Pë CĤC-C€PμPSPSCĤ PïCĤPs
PïCĤPëPiPëPSPμPSPSCĤ PiPsPrCřPIP°PSPSCĤ PiCĤCřPrPrCĤ P°P±Ps PïCĤPëPiPëPSPμPSPSCĤ
C,PμCĤP°PiC-C— CřPjPμPeP»C-PrPëPSC-CřPjPsPj/PIC-P»P°PSC,PμCĤPsP»PsPj P·
CřCĤP°C...CřPIP°PSPSCĤPj PePsCĤPëCĤC,C- PiCĤCřPrPSPsPiPs PIPëPiPsPrPsPICřPIP°PSPSCĤ
PrP»CĤ PrPëC,PëPSPë C- PePsCĤPëCĤC,C- C,PμCĤP°PiC-C— PrP»CĤ P¶C-PSPePë.

$\text{P}_{\text{L}}\text{PsC}\overline{\text{Tj}}\text{P}\ddot{\text{e}}\text{PiCrPIP}^{\circ}\text{PSPSC}\sqcup \text{PrPsP}\cdot\text{P}\ddot{\text{e}} \text{ PSP}_{\mu} \text{ P}\ddot{\text{i}}\text{PsC},\overline{\text{Tj}}\text{C}-\text{P}\pm\text{PSPs PrP}\gg\sqcup \text{ P}\ddot{\text{i}}\text{P}^{\circ}\text{C}\dagger\text{C}-\text{C}''\text{PSC},\text{C}-\text{PI P}\cdot$
 $\text{PiPsC}\overline{\text{Tj}}\text{CrC}\epsilon\text{P}_{\mu}\text{PSPSC}\sqcup\text{pj C,,CrPSPeC}\dagger\text{C}-\text{— P}\ddot{\text{i}}\text{P}_{\mu}\text{C}\dagger\text{C}-\text{PSPeP}\ddot{\text{e}} \text{ P}\gg\text{P}_{\mu}\text{PiPePsPiPs C,P}^{\circ}$
 $\text{PiPsPjC}-\overline{\text{Tj}}\text{PSPsPiPs C}\acute{\text{I}}\text{C,CrP}\ddot{\text{i}}\text{P}_{\mu}\text{PSC}\sqcup . \text{P}\acute{\text{k}}\text{P}_{\mu} \text{ PrPsC}\acute{\text{I}}\text{P}\gg\text{C-PrP}\P\text{CrPIP}^{\circ}\text{P}\gg\text{PsC}\acute{\text{I}}\sqcup$

$P \cdot P^{\circ} C \dot{I} C, P s C \dot{I} P s P I C \acute{r} P I P^{\circ} P S P S C \dot{I} P \gg C - P e P^{\circ} C \dot{I} C \dot{I} C H P e P s P i P s P \cdot P^{\circ} C \dot{I} P s P \pm C \acute{r} P \dot{h} P S P s C \dot{I} P s$
 $P \cdot P \gg P \gg C - P \dot{i} C, P^{\circ} P \dot{i} P^{\circ} C \dot{I} C - C^{\prime \prime} P S C, P^{\circ} P j P \cdot C, C \dot{I} P \dot{I} P e P s C \dot{I} P \dot{i} P \mu C \dot{I} C - P S P e P s P I P s C \dot{I} H$
 $P S P \mu P r P s C \dot{I} C, P^{\circ} C, P S C - C \dot{I} C, C \dot{I} H, C, P s P j C \acute{r} P N \dot{e} P s P i P s C \dot{I} P \gg C - P r P \cdot P^{\circ} C \dot{I} C, P s C \dot{I} P s P I C \acute{r} P I P^{\circ} C, P \ddot{e} P \cdot$
 $P s P \pm P \mu C \dot{I} P \mu P \dot{I} P S C - C \dot{I} C, C \dot{I} H.$

$P^{\prime \prime} C - C, P \ddot{e}$

$P \gg C - P e P^{\circ} C \dot{I} C \dot{I} C H P e P \ddot{e} P N \dot{e} P \cdot P^{\circ} C \dot{I} C - P \pm P \dot{h} P S P s C \dot{I} P s P \cdot P \gg P \gg C - P \dot{i} C, P^{\circ} P S P \mu$
 $P \cdot P^{\circ} C \dot{I} C, P s C \dot{I} P s P I C \acute{r} P I P^{\circ} P \gg P \ddot{e} P r C - C, C \dot{I} P j (P I C - P e P s P j P r P s 18 C \dot{I} P s P e C - P I) P r P \gg C \dot{I} P$
 $P \gg C - P e C \acute{r} P I P^{\circ} P S P S C \dot{I} P \dot{I} P \dot{h} P - P \gg.$

$P \ddot{Y} P i P s C \dot{I} C - P \pm P \cdot P^{\circ} C \dot{I} C, P s C \dot{I} C \acute{r} P I P^{\circ} P S P S C \dot{I} P$

$P \gg C - P e P^{\circ} C \dot{I} C \dot{I} C H P e P \ddot{e} P N \dot{e} P \cdot P^{\circ} C \dot{I} C - P \pm P \dot{h} P S P s C \dot{I} P s P \cdot P \gg P \gg C - P \dot{i} C, P^{\circ} P \dot{i} C \dot{I} P \ddot{e} P \cdot P S P^{\circ} C \dot{I} P \mu P S P \ddot{e} P N \dot{e}$
 $C, C - P \gg C \dot{I} H P e P \ddot{e} P r P \gg C \dot{I} P C - P S P i P^{\circ} P \gg C \dot{I} C \dot{I} C - P N \dot{e}.$

$P^{\prime} P e P^{\circ} P \cdot C - P I P e P \ddot{e} C \% P s P r P s P \cdot P^{\circ} C \dot{I} C, P s C \dot{I} C \acute{r} P I P^{\circ} P S P S C \dot{I} P$

$P \dot{I} P S P i P^{\circ} P \gg C \dot{I} C, P s C \dot{I} P \cdot P \gg P \gg C - P \dot{i} C, P^{\circ} P j C - C \dot{I} C, P \ddot{e} C, C \dot{I} H P \dot{i} P s P \dot{i} P \mu C \dot{I} P \mu P r P S C \dot{I} H P s$
 $C \dot{I} P s P \cdot P \dot{i} P s P r C - P \gg P \mu P S C - P r P s P \cdot P \ddot{e} C - P i P s C, P s P I P \ddot{e} P N \dot{e} P r P s P I P \ddot{e} P e P s C \dot{I} P \ddot{e} C \dot{I} C, P^{\circ} P S P S C \dot{I} P.$

$P \dot{I} P S P i P^{\circ} P \gg C \dot{I} C, P s C \dot{I} C \acute{r} P \dot{i} P^{\circ} P e P s P I P^{\circ} P S P \ddot{e} P N \dot{e} C \acute{r} P \gg P s C, P s P e, C \% P s P j C - C \dot{I} C, P \ddot{e} C, C \dot{I} H$
 $P I P s P \gg P s P i P s P \dot{i} P s P i P \gg P \ddot{e} P S P^{\circ} C \dot{I} C \dot{I} P \ddot{e} P N \dot{e} P \dot{i} P^{\circ} P e P \mu C, P \ddot{e} P e. P^{\prime} P s P \gg P s P i P s P \dot{i} P s P i P \gg P \ddot{e} P S P^{\circ} C \dot{I} C \dot{I} P \ddot{e} P N \dot{e}$
 $P \dot{i} P^{\circ} P e P \mu C, P \ddot{e} P e C \dot{I} P \gg C - P r P I P \ddot{e} P e P \ddot{e} P S C \acute{r} C, P \ddot{e}, P N \dot{e} P s P i P s P I P j C - C \dot{I} C, P S P \mu P j P s P \dot{I} P S P^{\circ}$
 $P e P s P I C, P^{\circ} C, P \ddot{e} P^{\circ} P \pm P s P I P r P \ddot{e} C \dots P^{\circ} C, P \ddot{e}. P \mu P^{\circ} C \dot{I} C - C^{\prime \prime} P S C, P^{\circ} P I P^{\circ} C \dot{I} C, P s P \dot{i} P s P \dot{i} P \mu C \dot{I} P \mu P r P \ddot{e} C, P \ddot{e},$
 $C \% P s P \gg P s C, P s P e P \cdot C - P S P i P^{\circ} P \gg C \dot{I} C, P s C \dot{I} P s P j C \dot{I} P \gg C - P r P I C - P r P e C \dot{I} P \ddot{e} P I P^{\circ} C, P \ddot{e} C, P s P r C -,$
 $P e P s P \gg P \ddot{e} P I C - P S P \pm C \acute{r} P r P \mu P i P s C, P s P I P \ddot{e} P N \dot{e} P r P s C - P S P i P^{\circ} P \gg C \dot{I} C \dot{I} C - C - P r P s P \cdot P \ddot{e}.$

$P \dot{I} P S P i P^{\circ} P \gg C \dot{I} C, P s C \dot{I} P I P \gg P s C, P e C \acute{r} P \dot{i} P \mu C \dot{I} P \mu P \pm C \acute{r} P I P^{\circ} C^{\prime \prime} P I P \dot{i} P s P \gg P s P \dot{I} P \mu P S P S C -$
 $B \ll P \cdot P^{\circ} P e C \dot{I} P \ddot{e} C, P s B \gg. P \mu C - C \dot{I} P \gg C \dot{I} C, P s P i P s, C \dot{I} P e P \gg P s C, P s P e P \pm C \acute{r} P r P \mu P I C - P r P e C \dot{I} P \ddot{e} C, P s,$
 $P N \dot{e} P s P i P s P j P s P \dot{I} P S P^{\circ} P I P \ddot{e} P e P \ddot{e} P S C \acute{r} C, P \ddot{e}. P^{\prime \prime} P^{\circ} C, P^{\circ} P e C - P S C \dot{I} P \mu P I P s P i P s$
 $P \cdot P^{\circ} C \dot{I} C, P s C \dot{I} C \acute{r} P I P^{\circ} P S P S C \dot{I} P C - P S P i P^{\circ} P \gg C \dot{I} C, P s C \dot{I} P^{\circ} C \dot{I} C, P^{\circ} P S P s P I P \ddot{e} C, C \dot{I} H 6 C, P \ddot{e} P \dot{I} P S C - P I P \cdot$
 $P r P^{\circ} C, P \ddot{e} P I C - P r P e C \dot{I} P \ddot{e} C, C, C \dot{I} P \gg P s C, P e P^{\circ}. P^{\prime \prime} P^{\circ} C, C \acute{r} P e C - P S C \dot{I} P \mu P I P s P i P s$
 $P \cdot P^{\circ} C \dot{I} C, P s C \dot{I} C \acute{r} P I P^{\circ} P S P S C \dot{I} P C \dot{I} P \gg C - P r P I P \dot{i} P \ddot{e} C \dot{I} P^{\circ} C, P \ddot{e} C \acute{r} C \dot{I} P \dot{i} P \mu C \dot{I} C - P^{\circ} P \gg C \dot{I} H P S P s$
 $P I C - P r P I P \mu P r P \mu P S P s P j C \acute{r} P j C - C \dot{I} C \dot{I} C -, P \dot{i} P s P \cdot P S P^{\circ} C \dot{I} P \mu P S P s P j C \acute{r} B \ll P \acute{K} P \mu$
 $P I P \ddot{e} P e P s C \dot{I} P \ddot{e} C \dot{I} C, P s P I C \acute{r} P I P^{\circ} C, P \ddot{e} P \dot{i} C - C \dot{I} P \gg C \dot{I} P \cdot B \gg. P \acute{K} P \mu C \dot{I} P \gg C - P r P \cdot P^{\circ} C \dot{I} C, P s C \dot{I} P s P I C \acute{r} P I P^{\circ} C, P \ddot{e}$
 $C - P S P i P^{\circ} P \gg C \dot{I} C, P s C \dot{I} P \dot{i} C - C \dot{I} P \gg C \dot{I} P r P^{\circ} C, P \ddot{e} P e C - P S C \dot{I} P \mu P I P s P i P s P \cdot P^{\circ} C \dot{I} C, P s C \dot{I} C \acute{r} P I P^{\circ} P S P S C \dot{I} P.$

$P j P \dot{i} P^{\circ} P e P s P I P e P^{\circ} P \gg C - P e P^{\circ} C \dot{I} C \dot{I} C H P e P s P i P s P \cdot P^{\circ} C \dot{I} P s P \pm C \acute{r} P \dot{h} P S P s C \dot{I} P s P \cdot P \gg P \gg C - P \dot{i} C, P^{\circ} C, P^{\circ}$
 $C - C - P I P j C - C \dot{I} C,$

$P \mu P s C \dot{I} C \dot{I} P r P s P e P I P \ddot{e} P e P s C \dot{I} P \ddot{e} C \dot{I} C, P^{\circ} P S P S C \dot{I} P C - P S P i P^{\circ} P \gg C \dot{I} C, P s C \dot{I} P^{\circ} (C \dot{I} P \ddot{e} C \dot{I} . 1)$

$1. P \mu C \dot{I} P s C \dot{I} P \ddot{e} C, P^{\circ} C, P \ddot{e} P \dot{i} P \mu C \dot{I} P \mu P r P \dot{i} P s C \dot{I} P^{\circ} C, P e P s P j P \cdot P^{\circ} C \dot{I} C, P s C \dot{I} C \acute{r} P I P^{\circ} P S P S C \dot{I} P$

$P \dot{i} P e C \% P s P e C \dot{I} P \ddot{e} C \acute{e} P e C \acute{r} C - P S P i P^{\circ} P \gg C \dot{I} C, P s C \dot{I} P^{\circ} P \pm C \acute{r} P r P \mu P I C - P r P e C \dot{I} P \ddot{e} C, P s C -$
 $P \cdot P^{\circ} P e C \dot{I} P \ddot{e} C, P s P \pm P \mu P \cdot C - P S P i P^{\circ} P \gg C \dot{I} C \dot{I} C - C - P \dot{i} C \dot{I} P \mu P \dot{i} P^{\circ} C \dot{I} P^{\circ} C, C \acute{r}, C, P s P r P s P \cdot C \acute{r} P \pm C \acute{r} P r P \mu$
 $P I C, C \dot{I} P^{\circ} C \dot{I} P \mu P S P s. P^{\prime} C, C \dot{I} P^{\circ} C \dot{I} P \mu P S P^{\circ} P r P s P \cdot P^{\circ} P S P^{\circ} P r C - P N \dot{e} P S P s P \cdot P \pm P \mu C \dot{I} C - P i P^{\circ} C^{\prime \prime} C, C \dot{I} H C \dot{I} C \dot{I} P$

PI C-PSIP°P»CЦC,PsCЦC-, P°P»Pμ PIP¶Pμ PSPμ C” PrPsCГC,CrPіPSPsCЦ PrP»CЦ
C-PSIP°P»CЦC†C-C—.

РќРμРјРѕР¶Р»РөРІРѕ РІРөРІ°РrРeРѕРІРѕ РіCЦРөР№PSCЦC,Рө PSP°РrР»РөCөРeРѕРІCГ P°P±Ps
РіРѕРrРІC-R№PSCГ РrРѕР·CГ РіCЦРμРІР°CЦР°C,CГ P·P° PsPrPSCГ C-PSIP°P»CЦC†C-CЦ.

P>C-C†PөP»CЦPSPөPө PrPsP·

P’C-PS РіРѕРeР°P·CГC”, CГPөC-P»CЦPөPө РrРѕР· РіCЦРμРІР°CЦР°C,CГ P·P°P»РөCөРөP»PsCГCЦ
PI C-PSIP°P»CЦC,PsCЦC-.

P”Ps P·P°CГC,PsCГCГPIP°PSPSCЦ РіРμCЦCөPөC— PrPsP·Pө P· C-PSIP°P»CЦC,PsCЦP° P»C-C†PөP»CЦPSPөPө РіРѕРeР°P·CГC” C,PsC†PSPs 30 PrPsP·.

P’C-PS PIC-PrCЦP°C...PsPICГC” РіРѕ 1 РrРѕР·C- PI P·РІРѕCЦРѕC,PSPsРјCГ РіРѕCЦCЦPPrPөCГ
C%PsCЦP°P·CГ, CЦPө PIC-PrPөCЦРөРІР°C”C,CЦCГCЦ PөCЦPөCөPөP°.

РљРѕР»Рө P·P°P»РөCөP°C”C,CЦCГCЦ РјРμPSCөPμ 10 PrPsP·, РіРѕР»P»PsPIPөPSP° P»C-C†PөP»CЦPSPөPөP° CГC,P°C” C†PμCЦPIPsPSPsРіРѕ РөРѕР»CЦPөP°.

РμC-CГP»CЦ РІРөРeРѕCЦРөCГC,P°PSPSCЦ PsCГC,P°PSPSCЦPөC— РrРѕР·Pө РіРѕР»P»PsPIPөPSP°
**P»C-C†PөP»CЦPSPөPөP° РrРѕР· P·P°P±P°CЦPIP»PμPSP° C†PμCЦPIPsPSPөPј C-
PIC-PrPsP±CЦP°P¶P°C”C,CЦCГCЦ C†PөCГP»Ps 0.** РђРμРіРμCЦ C-PSIP°P»CЦC,PsCЦ
РіРѕCЦPѕP¶PSC-R№.

РІРөC%Ps PIC-PrPөCЦPөC,Рө PөCЦPөCөPөCГ РіC-CГP»CЦ C†CЦPѕРіРѕ, C,Ps

P»C-C†PөP»CЦPSPөPө РrРѕР· B CГC,P°PSPμ РіРѕРІPSC-CГC,CЦ C†PμCЦPIPsPSPөPј.B

РљCЦPөCөPөP°

**P©PsCЦP°P·CГ РіCЦРө PIC-PrPөCЦPөC,C,C- PIPө РіРѕC,CГC”C,Рμ PsPrPSCГ РrРѕР·CГ
РіCЦРμРІР°CЦР°C,CГ**

P РөCГ. 1

B 2. РІРө РіC-PrРіРѕC,CГPIP°C,Рө РrРѕР·CГ

РќРμ PIC-PrPөCЦРөРІР°P№C,Рμ PөCЦPөCөPөCГ, РrРѕРeРө PSPμ P±CГPrPμC,Рμ РіРѕC,PsPIC-
РіCЦРөР№PSCЦC,Рө РrРѕР·CГ. РќРμ CГC,CЦCГCөCГP№C,Рμ C-PSIP°P»CЦC,PsCЦ.

РμРѕC,CЦPіPSC-C,CЦ PөCЦPөCөPөCГ PIPSPөP·, РrРѕРeРө PSPμ РіРѕC†CГC”C,Рμ
B«PөP»P°C†B» (CЦPөCГ. 2).

РђРμРіРμCЦ РіCЦРμРІР°CЦР°C, РіРѕC,PsPIPөP№ РrРѕ C-PSIP°P»CЦC†C-C—.

P>C-C†PөP»CЦPSPөPө РrРѕР· PIC-PrCЦP°C...PsPICГC” PI P·РІРѕCЦРѕC,PSPsРјCГ
РіРѕCЦCЦPPrPөCГ PSP° 1, C%Ps РіC-PrC,РІРμCЦPPrP¶CГC” РіРѕC,PsPіPSC-CГC,CЦ.

РІРөC%Ps РіC-CГP»CЦ C,PsРіРѕ, CЦPө PIPө РіРѕC†CГP»Pө B«PөP»P°C†B»,

P»C-C†PөP»CЦPSPөPө РrРѕР· PSPμ PIC-PrCЦP°C...CГPIP°PI РrРѕР·CГ PI

P·РІРѕCЦРѕC,PSPsРјCГ РіРѕCЦCЦPPrPөCГ, C,Ps C-PSIP°P»CЦC,PsCЦ PSPμ

РіCЦP°C†CЦC”. РќРμРѕР±C...C-PrPSPs P·РІРμCЦPSCГC,РөCГCЦ P·

C-PSIP°P»CЦC,PsCЦPѕРј PI P°PіC,РμPөCГ P·P° РіРѕCЦР°P°РrРѕCЦ. Рј P±CГPrCЦ-

CЦPөPѕРјCГ РІРөРІ°РrРөCГ CГC,CЦCГCөCГPIP°C,Рө C-PSIP°P»CЦC,PsCЦ PSPμ
CГP»C-Pr.

P РөCГ. 2

3. РІРө РІРөРeРѕPSP°C,Рө C-PSIP°P»CЦC†C-CЦ РіCЦРμРІР°CЦР°C,CГ

PJC, C₇H₆P₆Jc₇P₆N₆C, P_μ C-PSPIP°P»C₇IC, PsC₇H PSP° PIC-PrC₇IC, P°PSC- PIC-Pr C₇H₆PsC, P° C-
P·C₇H₆PsP±C-C, CH₆ PIP₆PrP₆PeC... PSP°C₇IC, C-P»CH₆PeP₆ PiP»P₆PeP±PsPeP₆PN₆,
PSP°C₇IC-P»CH₆PeP₆ C†P_μ PrP»C₇IC PIP°C₇IC PePsPjC,, PsC₇H₆C, PSPs.

P₆KP_μ PIP₆PrP₆PeC...P°PN₆C, P_μ PI C-PSPIP°P»C₇IC, PsC₇H.

P PsP·PjC-C₇IC, C-C, CH₆ PjC₇PrP₆PeC, C₇IC PjC-P₇ PiC₇P±P°PjP₆ C- C%₆C-P»CH₆PSPs
P₆C₇H₆PeC, P₆C₇IC-PSC-C, CH₆ C—C... PSP°PIPePsP»Ps PjC₇PrP₆PeC, C₇IC P₆° (C₇H₆PeC₇IC. 3).

P₆KP_μ PiP_μC₇H₆P_μPeC₇H₆PIP°PN₆C, P_μ PiP°P»CH₆C†C₇IC PjP₆ PiPsPIC-C, C₇H₆IC-PSC- PeP»P°PiP°PSP₆.

P—C₇H₆PsP±C-C, CH₆ PsPrP₆PePS PrPsPIPiP₆PN₆, C₇H₆C-PIPSP₆PN₆, PiP»P₆PeP±PsPeP₆PN₆ PIP₆PrP₆PeC...,
PiC-C₇IC»C₇IC C†PsPiPs P·P°C, C₇H₆PePjP°PN₆C, P_μ PrP₆PeC...P°PSPSC₇IC C₇ICPePsPjPsPiP° PrPsPIC₆P_μ
(C%₆PsPSP°PN₆PjP_μPSC₆P_μ 3в₇“4 C₇ICP_μPeC₇PrP₆PeC).

P°P₆PN₆PjC-C, CH₆ C-PSPIP°P»C₇IC, PsC₇H P· C₇H₆PsC, P°.

P_μPsPIC-P»CH₆PSPs C- PsP±P_μC₇H₆P_μP₇PSPs PIP₆PrP₆PeC...PSC-C, CH₆.

P P₆C₇IC. 3

P₆K°PIC-C, CH₆ PiC₇H₆Pe PiC₇H₆P°PIP₆Pe»CH₆PSPsPjC₇ PIP₆PePsC₇H₆PeC₇IC, P°PSPSC-
C-PSPIP°P»C₇IC, PsC₇H₆P° P°P₆ PjPsP₇P_μC, P_μ PSP_μ PIC-PrC†C₇IC, P₆ PiC₇H₆P_μPiP°C₇H₆P°C,,
P·PsPeC₇H₆P_μPjP° PN₆PsPiPs C₇ICPjP°Pe.

4. P—P°PeC₇H₆PePN₆C, P_μ C-PSPIP°P»C₇IC, PsC₇H

P₆PeC%₆Ps C” PSP_μPsP±C...C-PrPSC-C₇IC, CH₆ PI PsC†P₆C₇IC, C†C- PjC₇PrP₆PeC, C₇IC P₆°,
PIP₆PePsC₇H₆PeC₇IC, PsPIC₇PN₆C, P_μ C₇IC₇C...C₇IC, PeP°PSP₆PeSC₇IC PrPs P·P°PeC₇H₆PeC, C, C₇IC
PeC₇H₆PeC₆PeP₆.

P©PsP± PSP°PeC₇H₆PeC, P₆ PjC₇PrP₆PeC, C₇IC, PiPsPIP_μC₇H₆PSC-C, CH₆ PeC₇H₆PeC₆PeC₇IC PSP°
PjC-C₇IC†P_μ, C₇IC₇C...P°C₇H₆PeC—C— PIP_μC₇H₆C... PrPs C₇ICPiPsC₇IC (C₇H₆PeC₇IC. 4).

P P₆C₇IC. 4

P°C-C, P₆. P>C-PeP°C₇IC₇CH₆PeP₆PN₆ P·P°C₇IC-P± P₆H₆PSPsC₇H₆Ps P·P»P»C-PiC, P° PSP_μ
P·P°C₇IC, PsC₇IC₇PsPIC₇PIP°P»P₆ PrC-C, C₇IC Pj (PIC-PePsPj PrPs 18 C₇H₆PsPeC-PI) PrP»C₇IC
P»C-PeC₇ICPIP°PSPSC₇IC P₇ICP—P>.

P_μP_μC₇H₆P_μPrPsP·C₇PIP°PSPSC₇IC

P_μP_μC₇H₆P_μPrPsP·C₇PIP°PSPSC₇IC C₇ICPjP_μPeP»C-PrP₆PeSC-C₇IC₇IC/PIC-P»P°PSC, P_μC₇H₆PsP»C₇IC,
PN₆PjPsPIC-C₇H₆PSPs, PjP°C, P₆PePjP_μ PsP·PSP°PeP₆ C- C₇IC₇ICPjPiC, PsPjP₆, PiPsPIB₇™ C₇ICP·P°PSC- P·
PrC-C”C₇H₆PsPeC₇H₆PjP₆PeC... PePsPjPiPsPSP_μPSC, C-PI, C%₆Ps C₇IC·PiPsPrP₇IC₇IC”C, CH₆C₇IC₇IC P·
PIC-PrPsPjP₆PePjP₆ PiPsP±C-C†PSP₆PePjP₆ C₇H₆P_μP°PeC†C-C₇IC PjP₆ C-PSPIP°P»C₇IC†C-PN₆PSP₆PeC...

PjCfCfPeP°CfPePSPsPIPeC... P°PSC,P°PiPsPSC-CfC,C-PI (PSP°PiCfPePeP»P°Pr,
CfCfC...C-CfC,Ch Cf CfPsC,C-, PiPsCfCfC€PµPSPSCf P·PsCfPsPIPsC—
P°PePsPjPsPrP°C†C-C— C,P° C,P°C...C-PeP°CfPrC-Cf) P°P±Ps CfPµP°PeC†C-CfPjPe, C%Ps
CfPiPsCfC,PµCfC-PiP°P»PëCfCf PiCfPe PiPµCfPµPrPsP·CfPIP°PSPSC- C-PSC€PeC...
P±PµC,P°₂-P°PrCfPµPSPµCfPiC-C†PSPëC... P°PiPsPSC-CfC,C-PI (PSP°PiCfPePeP»P°Pr,
P°CfPeC,PjC-Cf, C,CfPµPjPsCf, PiPsP»PsPIPsPëPµ P±C-P»Ch, PiPsCfPeP»PµPSPµ
CfPµCfC†PµP±PeC,C,Cf, PSCfPrPsC,P°, PiC-PiPµCfPiP»C-PePµPjC-Cf C,P°
PiC-PiPsPeP°P»C-C”PjC-Cf).

PJ CfP°P·C- PiPµCfPµPrPsP·CfPIP°PSPSCf P»C-PeCfPIP°PSPSCf PiP°C†C-C”PSC,P° PjP°C”
P±CfC,Pë CfPePjPiC,PsPjP°C,PëC†PSPëPj P· PIC-PrPiPsPIC-PrPSPëPj PjPsPSC-C,PsCfPePSPiPsPj
Cf CfP°P·C- PiPsC,CfPµP±Pe.

PµPsP±C-C†PSC- CfPµP°PeC†C-C—

PhPiP»CfIPr PiCfPsC,,C-P»Cf P±PµP·PiPµPePe

PkP°PµP±C-P»ChC€ C†P°CfC,PsCf P·P°CfPµC”CfC,CfPsPIP°PSPsCf PiPsP±C-C†PSPsCf
CfPµP°PeC†C-C”Cf PiCfPe P·P°CfC,PsCfCfPIP°PSPSC-
CfPjPµPeP»C-PrPëPSC-CfPjCf/PIC-P»P°PSC,PµCfPsP»Cf P±CfPI PSP°P·PsC,,P°CfPePSPiC-C, (9
%).

PµPµCfPµP»C-Pe PiPsP±C-C†PSPëC... CfPµP°PeC†C-Pµ

PµCfPsC,,C-P»Ch P±PµP·PiPµPePe P»C-PeP°CfCfChPePsPiPs P·P°CfPsP±Cf PhPSPsCfPs
P·P»P»C-PiC,P° T”CfCfPSC,CfC”C,ChCfCf PSP° PrP°PSPëC... C%PsPrPs
P·P°CfC,PsCfCfPIP°PSPSCf PePsPjP±C-PSP°C†C-C—
CfPjPµPeP»C-PrPëPSC-CfPjCf/PIC-P»P°PSC,PµCfPsP»Cf C,P° PsPeCfPµPjPeC... C—C—
PePsPjPiPsPSPµPSC,C-PI, PsC,CfPePjP°PSPëC... Cf PjPµP¶P°C... PiCfPsPiCfP°PjPe
PeP»C-PSC-C†PSPëC... PrPsCfP»C-PrP¶PµPSCh, C%Ps PIPeP»CfC†P°P»P° 6 855
PiP°C†C-C”PSC,C-PI C-P· PfPhP—P», C,P° PiC-Pr C†P°Cf PiPsCfC,PjP°CfPePµC,PëPSPiPsPIPsPiPs
CfPiPsCfC,PµCfPµP¶PµPSPSCf. P”Ps PiCfPsPiCfP°PjPe PeP»C-PSC-C†PSPëC...
PrPsCfP»C-PrP¶PµPSCh P±CfP»Ps P·P°P»CfC†PµPSPs 2 354 PiP°C†C-C”PSC,Pë, C%Ps
PsC,CfPePjCfPIP°P»Pë CfPjPµPeP»C-PrPëPSC-CfPj/PIC-P»P°PSC,PµCfPsP» PsPrPëPS CfP°P·
PSP° PrPsP±Cf Cf PeP»C-PSC-C†PSPëC... PrPsCfP»C-PrP¶PµPSPSCfC... PSP° PµC,P°PiC-
C,,P°P·Pë 3 C,CfPePIP°P»C-CfC,Cf 24 C,PëP¶PSC- C- P±C-P»ChC€Pµ, P· CfPePeC... 1 296
PiP°C†C-C”PSC,C-PI PsC,CfPePjCfPIP°P»Pë CfPµPePsPjPµPSPPrPsPIP°PSCf PrPsP·Cf 55/22 PjPePi
Cf 24-C,PëP¶PSPµPIPsPjCf PrPsCfP»C-PrP¶PµPSPSC-, 832 PiP°C†C-C”PSC,Pë
PsC,CfPePjCfPIP°P»Pë PIPëCfPsPeCf PrPsP·Cf 113/22 PjPePi Cf 24-C,PëP¶PSPµPIPsPjCf
PrPsCfP»C-PrP¶PµPSPSC- C- 226 PiP°C†C-C”PSC,C-PI PsC,CfPePjCfPIP°P»Pë 113/22 PjPePi
PiCfPsC,CfPiPsPj 12-PjC-CfCfC†PSPsPiPs PrPsCfP»C-PrP¶PµPSPSCf.

PSP°CfC,PsC,P° PiPsP±C-C†PSPëC... CfPµP°PeC†C-Pµ, PIPëP·PSP°C†PµPSPëC... Cf
C,P°P±P»PëC†C- 2, PIPeP»CfC†P°C” PiPsPeP°P·PSPëPePe P·P°PiP°P»ChPSPsPiPs
PePsPµC,,C-C†C-C”PSC,P° P·P°C...PIPsCfChPIP°PSPsCfC,C-, PIPëP·PSP°C†PµPSC- PiCfPe
PsP±Bf™C”PrPSP°PSPSC- Pibf™CfC,Pë 24-C,PëP¶PSPµPIPëC... PrPsCfP»C-PrP¶PµPSCh C- 12-
PjC-CfCfC†PSPsPiPs PrPsCfP»C-PrP¶PµPSPSCf P±PµP·PiPµPePe.

P—P°CfC,PsCfPsPICfC”C,ChCfCf C,P°PeP° PeP»P°CfPeC,,C-PeP°C†C-Cf C†P°CfC,PsC,Pë
PIPëPSPëPePSPµPSPSCf PiPsP±C-C†PSPëC... PµC,,PµPeC,C-PI: PrCfP¶Pµ C†P°CfC,Ps Bf™³ 1/10;

$C\ddot{P}^{\circ}C\dot{I}C,Ps\text{ в}\ddot{H}''^3 1/100\text{ }C,P^{\circ} < 1/10$; $PSP_{\mu}C\ddot{P}^{\circ}C\dot{I}C,Ps\text{ в}\ddot{H}''^3 1/1000\text{ }C,P^{\circ} < 1/100$; $C\ddot{H}C-PrPePs\text{ в}\ddot{H}''^3 1/10000\text{ }C,P^{\circ} < 1/1000$; $PrCrP\Pi P_{\mu}C\ddot{H}C-PrPePs\text{ в}\ddot{H}'' < 1/10000$; $PSP_{\mu}PIC-PrPsPjPs\text{ в}\ddot{H}'' PSP_{\mu}PjPsP\Pi PSP^{\circ}PIP\ddot{e}P\cdot PSP^{\circ}C\ddot{P}eC,P\ddot{e}P\cdot PSP^{\circ}C\Pi PIPSP\ddot{e}C... PrP^{\circ}PSP\ddot{e}C....$

PyP°P±P»PëC†CΠ 2

PyPëC†C,PμPjPë C,P° PsCḡPiP°PSPë	PkPμP±P°P\P°PSP° CḡPμP°PeC†C-CΠ	P§P°C†C,PsC,P°
P†PSC,,PμPeC†C-C— C,P° C-PSPIP°P·C-C—	P†PSC,,PμPeC†C-CΠ C†PμC†PsPIPëPIC-PrPSPëC... CëP»CΠC...C-PI PyPëPSCrC†PëC, PkP°P·PsC,,P°CḡPëPSPiC-C, PαP°CḡPëPSPiC-C, P†PSC,,PμPeC†C-CΠ PIPμCḡC...PSC-C... PrPëC...P°P»CHPSPëC... CëP»CΠC...C-PI	P§P°C†C,Ps
PμPsCḡCrCëPμPSPSCΠ P· P±PsPeCr C-PjCrPSPSPsC— C†PëC†C,PμPjPë	P PμP°PeC†C-C— PiC-PiPμCḡC†C†C,P»PëPIPsC†C,C-, PIPeP»CḡC†P°CḡC†Pë PIPëC†PëPi, P°PSP°C,,C-P»P°PeC†C-Cḡ, P°PSPiC-PsPSPμPICḡPsC,PëC†PSPëPNë PSP°P±CḡCΠPe, PeCḡPsPiPëPiḡ™ CΠPSPeCr	PkPμC†P°C†C,Ps P C-PrPePs P C-PrPePs P C-PrPePs
PμPsCḡCrCëPμPSPSCΠ P· P±PsPeCr PSPμCḡPIPsIPsC— C†PëC†C,PμPjPë	P°PsP»PsPIPSPëPNë P±C-P»CH PyCḡPμPjPsCḡ P°PëC†PiPμPIP·C-CΠ P—P°PiP°PjPsCḡPsC†PμPSPSCΠ	P§P°C†C,Ps PkPμC†P°C†C,Ps PkPμC†P°C†C,Ps PkPμPIC-PrPsPjPs
PμPsCḡCrCëPμPSPSCΠ P· P±PsPeCr PiC†PëC...C-PePë	PyCḡPëPIPsPiP°	PkPμC†P°C†C,Ps
PμPsCḡCrCëPμPSPSCΠ P· P±PsPeCr PsCḡPiP°PSC-PI P·PsCḡCr	PμPsCḡCrCëPμPSPSCΠ C†C-C,PePsC†C,C- P·PsCḡCr P°P»P°C†PePsPjP° PμC-PrPIPëC%PμPSPSCΠ PIPSC†C,CḡC-CëPSCḡPsPsC†PSPsPiPs C,PëC†PeCr P°C-P»CH PI PsC†P°C...	P C-PrPePs P C-PrPePs P C-PrPePs P C-PrPePs
PμPsCḡCrCëPμPSPSCΠ P· P±PsPeCr C†PμCḡC†PμPIPs-C†CrPrPëPSPSPsC— C†PëC†C,PμPjPë	PḡPëPiPsC,P»PëPIP° P°CḡPëC,PjC-CΠ PyCrPiCḡP°PIPμPSC,CḡPëPeCrP»CΠCḡPSP° C,P°C...C-PeP°CḡPrC-CΠ P†PrC-PsPIPμPSC,CḡPëPeCrP»CΠCḡPSPëPNë CḡPëC,Pj PyP°C...C-PeP°CḡPrC-CΠ PyCrPiCḡP°PIPμPSC,CḡPëPeCrP»CΠCḡPSC- PμPeC†C,CḡP°C†PëC†C,PsP»Pë PμPsC†PëP»PμPSPμ C†PμCḡC†PμP±PëC,C,CΠ	PkPμC†P°C†C,Ps
PμPsCḡCrCëPμPSPSCΠ P· P±PsPeCr PsCḡPiP°PSC-PI PrPëC...P°P»CHPSPsC— C†PëC†C,PμPjPë, PiCḡCrPrPSPsC— PeP»C-C,PePë C,P° C†PμCḡPμPrPsC†C,C-PSPSCΠ	PḡP°CëPμP»CH P PsC,PsPiP»PsC,PePsPIPëPNë P±C-P»CH P°PëC†C,,PsPSC-CΠ PμP°CḡP°PrPsPeC†P°P»CHPSPëPNë P±CḡPsPSC...PsC†PiP°P·Pj	P§P°C†C,Ps P§P°C†C,Ps PkPμC†P°C†C,Ps P C-PrPePs
PμPsCḡCrCëPμPSPSCΠ P· P±PsPeCr CëP»C†PSPePsIPs-PePëCëPePsPIPsPiPs C,CḡP°PeC,Cr	P—P°PiPsCḡ PyCrC...C-C†C,CH Cr CḡPsC,C-	P§P°C†C,Ps
PμPsCḡCrCëPμPSPSCΠ P· P±PsPeCr CëPeC-CḡPë C,P° PiC-PrCëPeC-CḡPSPëC... C,PeP°PSPëPS	P°PëC†PëPi	PkPμC†P°C†C,Ps

PsPrCḡP°P·Cṛ P¶ PīC-CṛP»Cḷ PrC-CṛC,P°PIP°PSPSCḷ C-PSPiP°P»CḷC,PSCḡP° P· P»PsC,PeP°.

PJṖiP°PePsPIPeP°. PḷPsCḡPsC€PsPe PrP»Cḷ C-PSPiP°P»CḷC†C-PNḷ, PrPsP·PsPIP°PSPëPNḷ, PīPs 55 PjPePi / 22 PjPePi / PrPsP·Cṛ, PīPs 30 PrPsP· Cṛ PīPsCḡPsC€PePsPIPsPjCṛ C-PSPiP°P»CḷC,PSCḡC- B,, - 1.

P†PSPiP°P»CḷC,PSCḡ PjC-CṛC,PeC,CH PrPIC- P±P»C-CṛC,PḷCḡPSC- CṛC,CḡC-C†PePë PīPs 30 CḡC-PIPPSPsPjC-CḡPSPs CḡPsP·PīPsPrC-P»PḷPSPëC... C†P°CḡCṛPSPsPe, PePsP¶PSP° P· CḷPePëC... PjC-CṛC,PeC,CH PīPs 1 PrPsP·C- PePsP¶PSPsC— PrC-CḡC†PsC— CḡPḷC†PsPIPëPSPë, C%Ps PrPsCṛC,P°PIP»CḷCḡC,CHCṛCḷ PsPrPSPsC†P°CṛPSPs.

PḷP»P°CṛC,PëPePsPIPëPNḷ C-PSPiP°P»CḷC,PSCḡ P·C- CṛPIC-C,P»Ps-CṛC-CḡPëPj PePsCḡPīCṛCṛPsPj, C†PḷCḡPIPsPSPëPj PePsPIPīP°C†PePsPj CḡPsP·PīPëP»CḡPIP°C†P° C,P° P»C-C†PëP»CHPSPëPePsPj PrPsP· CṛPīP°PePsPIP°PSPs PI P»PsC,PsPe P· C,,PsP»CHPiPë, C%Ps PjC-CṛC,PeC,CH PīP°PePḷC,PëPe P· PIPsP»PsPiPsPīPsPiP»PëPSP°C†PḷPj C-P· CṛPëP»C-PeP°PiPḷP»Cḡ. P>PsC,PsPe, P·P°PīP°CḷPSPëPNḷ PIC-PrCḡPëPIPPSPsCḡ PeCḡPëC€PePsCḡ, PīPsPjC-C%PḷPSPs Cṛ PeP°CḡC,PSPSPSCṛ PePsCḡPsP±PeCṛ.

PḷP°C,PḷPiPsCḡC-Cḷ PIC-PrPīCṛCṛPeCṛ. P—P° CḡPḷC†PḷPīC,PsPj. B

P°PëCḡPsP±PSPëPe. P°P»P°PeCṛPs PhPiPḷCḡPḷPNḷC€PSCṛ P®Pḷ P>C-PjC-C,PḷPr / Glaxo Operations UK Ltd. B

PḡC-CṛC†PḷP·PSP°C...PsPrP¶PḷPSPSCḷ PIPëCḡPsP±PSPëPeP° C,P° PNḷPsPiPs P°PrCḡPḷCṛP° PjC-CṛC†Cḷ PīCḡPsPIP°PrP¶PḷPSPSCḷ PrC-CḷP»CHPSPsCṛC,C-.
P°P»P°PeCṛPs PhPiPḷCḡPḷPNḷC€PSCṛ P®Pḷ P>C-PjC-C,PḷPr, PḷCḡP°PNḷPsCḡC- PṽC,CḡC-C,, P°PḷP°, SG12 0DJ, P°PḷP»PëPeP° P°CḡPëC,P°PSC-Cḷ / Glaxo Operations UK Ltd, Priory Street, Ware, SG12 0DJ, United Kingdom.

PḷCḡPḷPrCṛC,P°PIPSPëPe P·P°CḷPIPSPëPeP°. PṽPhP'
B<P°P»P°PeCṛPsPṽPjC-C,PḷP»CḷPNḷPs PḡP°CḡPjP°CṛCHCḡC,C-PeP°P»Cṛ PJPeCḡP°C—PSP°B».

PḡC-CṛC†PḷP·PSP°C...PsPrP¶PḷPSPSCḷ PīCḡPḷPrCṛC,P°PIPSPëPeP° P·P°CḷPIPSPëPeP°.
PJPeCḡP°C—PSP°, Pj. PḷPëC—PI, 02152, PīCḡPsCṛPīPḷPeC, PḷP°PIP»P° PṽPëC†PëPSPë, P±CṛPr. 1-P', C,PḷP»: (044) 585-51-85, C,,P°PeCṛ: (044) 585-51-92.

PṽP°PePsP¶ PīCḡPs CṛCṛC- PIPëPīP°PrPePë PīC-PrPsP·CḡCḡPIP°PSPëC... PīPsP±C-C†PSPëC... CḡPḷP°PeC†C-PNḷ C,P° PIC-PrCṛCṛC,PSPsCṛC,C- PḷC,,PḷPeC,PëPIPPSPsCṛC,C- P»C-PeP°CḡCṛCHPePsPiPs P·P°CṛPsP±Cṛ PjPsP¶PSP° PīPsPIC-PrPsPjP»CḷC,Pë PṽPhP'

* Pj C†CHbPsPjC† PrPsC†P»C-PrP¶PµPSPSC- PIPëPePsC¶PëC†C,PSPIC†PIP°P»P°C†C¶
PiC¶PsC†PµPrC†C¶P° C†C,P°C,PëC†C,PëC†PSPsPiPs C,PµC†C,C†PIP°PSPSC¶ P·
PiPsC†C,C†PiPsPIPëPj P·PSPëP¶PµPSPSC¶Pj, C- C†Pµ PiPsC¶C-PIPSC¶PSPSC¶ P±C†P»Ps
PSPëP¶C†Pµ C¶C-PIPSC¶ PrPsC†C¶PiPSPµPSPSC¶ C†C,P°C,PëC†C,PëC†PSPsC—
P·PSP°C†C†C%PsC†C,C- PhC,P¶Pµ, PSPµ PjPsP¶PSP° P·C¶PsP±PëC,Pë PIPëC†PSPsPIPeC†
PiC¶Ps C†C,P°C,PëC†C,PëC†PSC† P·PSP°C†C†C%C-C†C,CH C†CHbPsPiPs
PiPsC¶C-PIPSC¶PSPSC¶.

* PJ C†CHbPsPjCr PrPsCfP»C-PrP¶PµPSPSC- PIPëPePsCḡPëCfC,PsPICfPIP°P»P°CfC¶
PïCḡPsC†PµPrCrCḡP° CfC,P°C,PëCfC,PëC†PSPsPiPs C,PµCfC,CrPIP°PSPSC¶ P·
PïPsCfC,CrPïPsPIPëPj P·PSPëP¶PµPSPSC¶Pj, C- C†Pµ PïPsCḡC-PIPSCL¶PSPSC¶ P±CrP»Ps
PSPëP¶C†Pµ CḡC-PIPSCL PrPsCfC¶PiPSPµPSPSC¶ CfC,P°C,PëCfC,PëC†PSPsC—
P·PSP°C†CrC‰PsCfC,C-. PhC,P¶Pµ, PSPµ PjPsP¶PSP° P·CḡPsP±PëC,Pë PIPëCfPSPsPIPeCr
PïCḡPs CfC,P°C,PëCfC,PëC†PSCr P·PSP°C†CrC‰C-CfC,CH C†CHbPsPiPs
PïPsCḡC-PIPSCL¶PSPSC¶.