

**PrP»CЦ PjPμPrPëC‡PSPsPiPs P•P°CíC,PsCíCíPPIP°PSPSCЦ P»C-PeP°CíCíCñPePsPiPs
P•P°CíPsP±Cí**

CP†P[´]P•P“P P_h

(SINEGRA)

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

$$PrC-C\bar{H}C\bar{\neq}P^{\circ} \ C\bar{H}P_{\mu}C\bar{\neq}PsPIP\bar{e}PSP^{\circ}: C\bar{I}P\bar{e}P\gg PrP_{\mu}PSP^{\circ}C_{,,}C-P\gg;$$
$$1 \text{ C}, \text{P}^\circ \text{P} \pm \text{P} \gg \text{P}_{\mu} \text{C}, \text{P} \epsilon \text{P}^\circ \text{PjC} - \text{C} \acute{\text{I}} \text{C}, \text{P} \ddot{\text{e}} \text{C}, \text{CH} \text{ C} \acute{\text{I}} \text{P} \ddot{\text{e}} \text{P} \gg \text{PrP}_{\mu} \text{PSP}^\circ \text{C}, \text{C} - \text{P} \gg \text{C} \acute{\text{r}} \text{ C} \dagger \text{P} \ddot{\text{e}} \text{C}, \text{C} \overline{\text{H}} \text{P}^\circ \text{C}, \\ \text{P}_{\mu} \text{P} \epsilon \text{PIC} - \text{PIP}^\circ \text{P} \gg \text{P}_{\mu} \text{PSC}, \text{PSPs} \text{ C} \acute{\text{I}} \text{P} \ddot{\text{e}} \text{P} \gg \text{PrP}_{\mu} \text{PSP}^\circ \text{C}, \text{C} - \text{P} \gg \text{C} \acute{\text{r}} \text{ 25 PjPi};$$
$$1 \text{ C}, \text{P}^\circ \text{P} \pm \text{P} \gg \text{P}_{\mu} \text{C}, \text{PeP}^\circ \text{ PjC} - \text{C}^{\dagger} \text{C}, \text{P}^{\dagger} \text{C}, \text{CH} \text{ C}^{\dagger} \text{P}^{\dagger} \text{P} \gg \text{PrP}_{\mu} \text{PSP}^\circ \text{C}, \text{C} - \text{P} \gg \text{Cr} \text{ C}^{\dagger} \text{P}^{\dagger} \text{C}, \text{C}^{\dagger} \text{P}^\circ \text{C}, \text{P}_{\mu} \text{PePIC} - \text{PIP}^\circ \text{P} \gg \text{P}_{\mu} \text{PSC}, \text{PSPs} \text{ C}^{\dagger} \text{P}^{\dagger} \text{P} \gg \text{PrP}_{\mu} \text{PSP}^\circ \text{C}, \text{C} - \text{P} \gg \text{Cr} \text{ 50 PjPi};$$

1 C,P°P±P»PμC,PεP° PjC-CΓC,PëC,CH CΓPëP»PrPμPSP°C,,C-P»Cr C†PëC,CΓP°C,
PμPePIC-PIP°P»PμPSC,PSPs CΓPëP»PrPμPSP°C,,C-P»Cr 100 PjPi;

PrPsPiPsPjC-P¶PSC- C¶PµC±PsPIPëPSPë: PeP°P»CHC†C-CH PiC-PrC¶PsC,,PsC†C,,P°C, P±PµP·PIPsPrPSPëPNë, C†PµP»CHP»PsP·P° PjC-PeC¶PsPeC¶PëC†C,P°P»C-C±PSP°, PSP°C,C¶C-CH PeC¶PsC†PeP°C¶PjPµP»PsP·P°, PeC¶PµPjPSC-CH PrC-PsPeC†PëPr PePsP»PsC—PrPSPëPNë P±PµP·PIPsPrPSPëPNë, PjP°PiPSC-CH C†C,PµP°C¶P°C,, PiPsPeC¶PëC,C,C¶ Opadry II Blue 85F20578: C†PiPëC¶C, PiPsP»C-PIC-PSC-P»PsPIPëPNë, PjP°PeC¶PsPiPsP» 4000, C,PëC,P°PSC† PrC-PsPeC†PëPr (E 171), C,P°P»CHPe, C-PSPrPëPiPsPeP°C¶PjC-PSC† P°P»CHPjC-PSC-C¶PIPëPNë P»P°Pe (E 132), P·P°P»C-P·P° PsPeC†PëPr P¶PsPIC,PëPNë (E 172).

P>C-PeP°CTĜĆĤĤPeP° C,,PsCTĜPjP°. PŷP°P±P»PμC,PePě, PIPeCTĜPěC,C-PiP»C-PIPePsPIPsCTĤ PsP±PsP»PsPSPePsCTĤ.

$$Ph\dot{C}I\dot{P}SPsPIPSC-C_{\text{„}}C-P\cdot P\ddot{e}PePs-C...C-PjC-C\ddot{z}PSC-PIP\gg P^{\circ}CI\dot{C},P\ddot{e}PIPsCI\dot{C},C-:$$

$C, P^{\circ}P \pm P \gg P_{\mu}C, PeP\ddot{e} \text{ } \overline{P}iPs \text{ } 25 \text{ } PjPi: C\acute{I}P\ddot{e}PSCHbPsPiPs \text{ } PePsP \gg CHbPsC\overline{H}C\acute{r} \text{ } C, P^{\circ}P \pm P \gg P_{\mu}C, PeP\ddot{e},$
 $PIPeC\overline{H}P\ddot{e}C, C- \overline{P}iP \gg C-PIPePsPIPsC\overline{H} \text{ } PsP \pm PsP \gg PsPSPePsC\overline{H}, PsPeC\overline{H}C\acute{r}PiP \gg Ps-$
 $PePIP^{\circ}P\overline{r}C\overline{H}P^{\circ}C, PSPsC- C_{,,}PsC\overline{H}PiP\ddot{e};$

$C, P^{\circ}P \pm P \gg P_{\mu}C, PeP\ddot{e} \text{ } PiPs \text{ } 50 \text{ } PjPi$: $C\acute{I}P\ddot{e}PSC\grave{H}PsPiPs \text{ } PePsP \gg C\grave{H}PsC\grave{H}C\acute{r} \text{ } C, P^{\circ}P \pm P \gg P_{\mu}C, PeP\ddot{e},$
 $PIPeC\grave{H}P\ddot{e}C, C - PiP \gg C - PIPePsPIPsC\grave{H} \text{ } PsP \pm PsP \gg PsPSPePsC\grave{H}, \text{ } PsPeC\grave{H}C\acute{r}PiP \gg Ps -$
 $PePIP^{\circ}PrC\grave{H}P^{\circ}C, PSPsC - C_{,,}PsC\grave{H}PjP\ddot{e} \text{ } P \cdot P \gg C - PSC - C''C\grave{H} \text{ } C\grave{H}PsP \cdot P \gg PsPjC\acute{r} \text{ } P \cdot \text{ } PsP \pm PsC \dots$
 $P \pm PsPeC - PI$;

$C, P^{\circ}P \pm P \gg P_{\mu}C, P_{\epsilon}P_{\delta} \quad P\ddot{I}Ps \quad 100 \quad PjPi: \quad C\ddot{I}P_{\delta}PSC\ddot{H}P_{\delta}P_{\delta}P_{\delta}P_{\delta} \quad P_{\epsilon}P_{\delta}P \gg C\ddot{H}P_{\delta}C\ddot{I}C\ddot{r} \quad C, P^{\circ}P \pm P \gg P_{\mu}C, P_{\epsilon}P_{\delta},$

PIPeCḡPëC,C- PİP»C-PIPePsPIPsCḡ PsP±PsP»PsPSPePsCḡ, PsPeCḡCfPiP»Ps-
PePIP°PrCḡP°C,PSPsC— C„PsCḡPjPë P· PİPsPrPIC-PNëPSPsCḡ P»C-PSC-C”Cḡ CḡPsP·P»PsPjCf P·
PsPrPSPsPiPs P±PsPeCf.

P»P°CḡPjP°PePsC,PμCḡP°PİPμPIC,PëC±PSP° PiCḡCfPİP°. P—P°CfPsP±Pë, C%Ps
P·P°CfC,PsCfPsPICfCḡC,CHCfCḡ PİCḡPë PμCḡPμPeC,PëP»CHPSC-PNë
PrPëCfC,,CfPSPeC†C-C—. PŸPëP»PrPμPSP°C,,C-P». PḡPsPr PḡPŸPİ G04B E03.

P»P°CḡPjP°PePsP»PsPiC-C±PSC- PIP»P°CfC,PëPIPsCfC,C-.

P»P°CḡPjP°PePsPrPëPSP°PjC-PeP°.

PḡPμC...P°PSC-P·Pj PrC-C—. PŸPëP»PrPμPSP°C,,C-P» C” PİCḡPμPİP°CḡP°C,PsPj PrP»Cḡ
PİPμCḡPsCḡP°P»CHPSPsPiPs P·P°CfC,PsCfCfPIP°PSPSCḡ, PİCḡPëP·PSP°C±PμPSPëPj
PrP»Cḡ P»C-PeCfPIP°PSPSCḡ PμCḡPμPeC,PëP»CHPSPsC— PrPëCfC,,CfPSPeC†C-C—.
PμCḡPë CfC,P°C,PμPIPsPjCf P·P±CfPrPḡPμPSPSC- PİCḡPμPİP°CḡP°C,
PIC-PrPSPsPIP»CḡC” P·PSPëPḡPμPSCf PμCḡPμPeC,PëP»CHPSCf C,,CfPSPeC†C-Cḡ
C€P»CḡC...PsPj PİPsCfPëP»PμPSPSCḡ PİCḡPëC,PsPeCf PeCḡPsPIC- PrPs PİPμPSC-CfP°.

P»C-P·C-PsP»PsPiC-C±PSPëPNë PjPμC...P°PSC-P·Pj, C%Ps PIC-PrPİPsPIC-PrP°C” P·P°
PμCḡPμPeC†C-Cḡ, PİPeP»CḡC±P°C” PİPëPIC-P»CHPSPμPSPSCḡ PsPeCfPëPrCf P°P·PsC,Cf (NO)
Cf PeP°PIPμCḡPSPsP·PSPëC... C,C-P»P°C... PİC-Pr C±P°Cf CfPμPeCfCfP°P»CHPSPsPiPs
P·P±CfPrPḡPμPSPSCḡ. PḡPeCfPëPr P°P·PsC,Cf P°PeC,PëPICfC” C,,PμCḡPjPμPSC,
PiCfP°PSC-P»P°C,C†PëPeP»P°P·Cf, C%Ps CfC,PëPjCfP»CḡC” PİC-PrPIPëC%PμPSPSCḡ
CḡC-PIPSCḡ C†PëPeP»C-C±PSPsPiPs PiCfP°PSPsP·PëPSPjPsPSPsC,,PsCfC,,P°C,Cf (C†P”PḡPḡ),
C%Ps CfPİCḡPëC±PëPSCḡC” CḡPsP·CfP»P°P±P»PμPSPSCḡ PiP»P°PrPePsC—
PjCfCfPeCfP»P°C,CfCḡPë PİPμC±PμCḡPëCfC,PëC... C,C-P», CfPİCḡPëCḡCḡC±Pë
PİCḡPëC,PsPeCf PeCḡPsPIC-.

PŸPëP»PrPμPSP°C,,C-P» C” PİPsC,CfPḡPSPëPj CfPμP»PμPeC,PëPIPSPëPj
C-PSPiC-P±C-C,PsCḡPsPj C†P”PḡPḡ-CfPİPμC†PëC,,C-C±PSPsC—
C,,PsCfC,,PsPrC-PμCfC,PμCḡP°P·Pë 5 (P»P”P·5) Cf PeP°PIPμCḡPSPsP·PSPëC... C,C-P»P°C...,
PrPμ P»P”P·5 PIC-PrPİPsPIC-PrP°C” P·P° CḡPsP·PİP°Pr C†P”PḡPḡ. P’PİP»PëPI
CfPëP»PrPμPSP°C,,C-P»Cf PSP° PμCḡPμPeC†C-Cḡ PjP°C” PİPμCḡPëC,,PμCḡPëC±PSPëPNë
C...P°CḡP°PeC,PμCḡ. PŸPëP»PrPμPSP°C,,C-P» PSPμ C±PëPSPëC,CH
P±PμP·PİPsCfPμCḡPμPrPSCḡPsC— CḡPμP»P°PeCfCfCḡC±PsC— PrC-C— PSP°
C-P·PsP»CHPSPiP°PSC- PeP°PIPμCḡPSPsP·PSC- C,C-P»P° P»CHPrPëPSPë, P°P»Pμ
PİPsC,CfPḡPSPs PİPsCfPëP»CḡC” CḡPsP·CfP»P°P±P»CHPIP°P»CHPSCf PrC-Cḡ NO PSP° C†Cḡ
C,PeP°PSPëPSCf. PμCḡPë P°PeC,PëPIP°C†C-C— PjPμC,P°P±PsP»C-C±PSPsPiPs C€P»CḡC...Cf
NO/C†P”PḡPḡ, C%Ps PIC-PrP±CfPIP°C”C,CHCfCḡ PİCḡPë CfPμPeCfCfP°P»CHPSC-PNë
CfC,PëPjCfP»CḡC†C-C—, C-PSPiC-P±CfPIP°PSPSCḡ CfPëP»PrPμPSP°C,,C-P»PsPj P»P”P·5
PİCḡPëP·PIPsPrPëC,CH PrPs PİC-PrPIPëC%PμPSPSCḡ CḡC-PIPSCḡ C†P”PḡPḡ Cf
PeP°PIPμCḡPSPsP·PSPëC... C,C-P»P°C.... PŸP°PePëPj C±PëPSPsPj, PrP»Cḡ C,PsPiPs C%PsP±
CfPëP»PrPμPSP°C,,C-P» CfPİCḡPëC±PëPSCḡPI PİPsC,CḡC-P±PSPëPNë
C,,P°CḡPjP°PePsP»PsPiC-C±PSPëPNë PμC,,PμPeC,, PSPμPsP±C...C-PrPSPμ
CfPμPeCfCfP°P»CHPSPμ P·P±CfPrPḡPμPSPSCḡ.

P’PİP»PëPI PSP° C,,P°CḡPjP°PePsPrPëPSP°PjC-PeCf. P”PsCfP»C-PrPḡPμPSPSCḡ *in vitro*

PrPsCfCfCfPiP°C" 18 PSPi/PjP» (38 PSPjPsP»CH). PŸC,CfPŸC-PSCH P·PiBf™ CŸP·CfPIP°PSPSCŸ P·P±C-P»PeP°PjPë PiP»P°P·PjPë PeCfjPsPIC- PSPµ P·P°P»PµP¶PëC,CH PIC-Pr P·P°PiP°P»CHPSPëC... PePsPSC†PµPSC,CfjP°C†C-PNë CfPëP»PrPµPSP°C,,C-P»Cf.

PJ P·PrPsCfjPsPIPëC... PrPsP±CfjPsPIPsP»CHC†C-PI, CŸPeC- P·P°CfC,PsCfjPsPICfPIP°P»Pë CfPëP»PrPµPSP°C,,C-P» PsPrPSPScfjP°P·PsPIPs PI PrPsP·C- 100 PjPi, C†PµCfjPµP· 90 C...PIPëP»PëPS PI PµCŸPeCfP»CŸC,C- PIPëP·PSP°C†P°P»PsCfCŸ PjPµPSC€Pµ 0,0002 % (PI CfPµCfjPµPrPSCHPsPjCf 188 PSPi) P·P°CfC,PsCfjPsPIP°PSPsC— PrPsP·Pë.

P'C-PSc,CfjP°PSCfC,,PsCfjPjP°C†C-CŸ. P·PµC,P°P±PsP»C-P·Pj CfPëP»PrPµPSP°C,,C-P»Cf P·PrC-PNëCfPSCfC"C,CHCfCŸ PiPsP»PsPIPsPëPj C†PëPSPsPj P· CfC†P°CfC,CH PjC-PeCfjPsCfjPsPjP°P»CHPSPëC... C-P·PsC,,PµCfjPjPµPSC,C-PI PiPµC†C-PSPePë CYP3A4 (PiPsP»PsPIPsPëPNë C€P»CŸC...) C,P° CYP2C9 (PrCfjCfPiPsCfjCŸPrPSPëPNë C€P»CŸC...). P"PsP»PsPIPsPëPNë C†PëCfjPeCfP»CHCfC†PëPNë PjPµC,P°P±PsP»C-C, CfC,PIPsCfjCfC"C,CHCfCŸ PI CfjPµP·CfP»CHC,P°C,C- N-PrPµPjPµC,PëP»CHPIP°PSPSCŸ CfPëP»PrPµPSP°C,,C-P»Cf. PŸPµP»PµPeC,PëPIPsC-CfC,CH PjPµC,P°P±PsP»C-C,Cf PIC-PrPSPScfjPSPs P·P·5 PiPsCfjC-PIPsCfC"C,CHCfCŸ P· CfPµP»PµPeC,PëPIPsC-CfC,CH CfPëP»PrPµPSP°C,,C-P»Cf (in vitro), P° P°PeC,PëPIPsC-CfC,CH PjPµC,P°P±PsP»C-C,Cf PIC-PrPSPScfjPSPs P·P·5 CfC,P°PSPsPIPëC,CH PiCfjPëP±P»PëP·PSPs 50 % P°PeC,PëPIPsCfC,C- PIPëC...C-PrPSPsC— CfjPµC†PsPIPëPSPë. PµP»P°P·PjPsPIC- PePsPSC†PµPSC,CfjP°C†C-C— C†CHPsPiPs PjPµC,P°P±PsP»C-C,Cf CfC,P°PSPsPIP»CŸC,CH PiCfjPëP±P»PëP·PSPs 40 % PePsPSC†PµPSC,CfjP°C†C-C— CfPëP»PrPµPSP°C,,C-P»Cf Cf PiP»P°P·PjC- PeCfjPsPIC-. N-PrPµPjPµC,PëP»CHPsPIP°PSPëPNë PjPµC,P°P±PsP»C-C, P·P°P·PSP°C" PiPsPrP°P»CHC€PsPiPs PjPµC,P°P±PsP»C-P·PjCf, P° PiPµCfjC-PsPr PNëPsPiPs PSP°PiC-PIPIPëPIPµPrPµPSPSCŸ CfC,P°PSPsPIPëC,CH PiCfjPëP±P»PëP·PSPs 4 PiPsPrPëPSPë.

P·P»C-PjC-PSP°C†C-CŸ. P—P°PiP°P»CHPSPëPNë PeP»C-CfjPµPSCf CfPëP»PrPµPSP°C,,C-P»Cf CfC,P°PSPsPIPëC,CH 41 P»/PiPsPr P· CfjPµP·CfP»CHC,P°C,PsPj C,PµCfjPjC-PSP°P»CHPSPsC— C,,P°P·Pë PSP°PiC-PICfjPsP·PiP°PrCf 3-5 Pi. PµC-CfP»CŸ PiPµCfjPsCfjP°P»CHPSPsPiPs P°P±Ps PIPScfC,CfjC-C€PSCHPsPiPµPSPSPsPiPs PIPiPµPrPµPSPSCŸ CfPëP»PrPµPSP°C,,C-P» PIPëPIPsPrPëC,CHCfCŸ Cf PIPëPiP»CŸPrC- PjPµC,P°P±PsP»C-C,C-PI PiPµCfjPµPIP°P¶PSPs P· PeP°P»PsPj (PiCfjPëP±P»PëP·PSPs 80 % PIPiPµPrPµPSPsC— PiPµCfjPsCfjP°P»CHPSPsC— PrPsP·Pë) C- PjPµPSC€PsCf PjC-CfjPsCf P· CfPµC†PµCf (PiCfjPëP±P»PëP·PSPs 13 % PIPiPµPrPµPSPsC— PiPµCfjPsCfjP°P»CHPSPsC— PrPsP·Pë).

P·P°CfjPjP°PePsPeC-PSPµC,PëPeP° PI PsCfjPsP±P»PëPIPsC... PiCfjCzPi PiP°C†C-C"PSC,C-PI.

PµP°C†C-C"PSC,Pë P»C-C,PSCHPsPiPs PIC-PeCz. PJ P·PrPsCfjPsPIPëC... PrPsP±CfjPsPIPsP»CHC†C-PI P»C-C,PSCHPsPiPs PIC-PeCf (PIC-PePsPj PIC-Pr 65 CfjPsPeC-PI) PIC-PrP·PSP°C†P°P»PsCfCŸ P·PSPëP¶PµPSPSCŸ PeP»C-CfjPµPSCfCf CfPëP»PrPµPSP°C,,C-P»Cf, C%Ps P·CfPjPsPIP»CHPIP°P»Ps PiC-PrPIPëC%PµPSPSCŸ PiP»P°P·PjPsPIPëC... PePsPSC†PµPSC,CfjP°C†C-PNë CfPëP»PrPµPSP°C,,C-P»Cf C,P° PNëPsPiPs P°PeC,PëPIPsPsPiPs N-PrPµPjPµC,PëP»CHPsPIP°PSPsPiPs PjPµC,P°P±PsP»C-C,Cf PiCfjPëP±P»PëP·PSPs PSP° 90 % PiPsCfjC-PIPsCŸPSPs P· PIC-PrPiPsPIC-PrPSPëPjPë PePsPSC†PµPSC,CfjP°C†C-CŸPjPë Cf P·PrPsCfjPsPIPëC... PrPsP±CfjPsPIPsP»CHC†C-PI PjPsP»PsPrC€PsPiPs PIC-PeCf

(18-45 CfjPsPeC-PI). PJ P·PiBf™ CŸP·PeCf P· PIC-PePsPIPëPjPë PIC-PrPjC-PSPSPScfC,CŸPjPë Cf P·PiBf™ CŸP·CfPIP°PSPSC- P· P±C-P»PeP°PjPë PiP»P°P·PjPë PeCfjPsPIC- PIC-PrPiPsPIC-PrPSPµ PiC-PrPIPëC%PµPSPSCŸ PiP»P°P·PjPsPIPsC— PePsPSC†PµPSC,CfjP°C†C-C— PIC-P»CHPSPsPiPs CfPëP»PrPµPSP°C,,C-P»Cf CfC,P°PSPsPIPëP»Ps PiCfjPëP±P»PëP·PSPs

PkPëCĤPpPsPIP° PSPμPŕPsCĬC,P°C,PSC-CĬC,CH. PJ PrPsP±CĤPsPIPsP»CHC†C-PI C-P·
PiPsCĤCŕC€PμPSPSCĬPjPë C,,CŕPSPeC†C-C— PSPëCĤPsPe P»PμPiPePsPiPs C,P°
PiPsPjC-CĤPSPsPiPs CĬC,CŕPiPμPSCĬ (PeP»C-CĤPμPSCĬ PeCĤPμP°C,PëPSC-PSCŕ 30-80
PjP»/C...PI) C,,P°CĤPjP°PePsPeC-PSPμC,PëPeP° CĬPëP»PrPμPSP°C,,C-P»Cŕ
P·P°P»PëC€P°P»P°CĬCĬ PSPμP·PjC-PSPμPSPsCĤ PiC-CĬP»CĬ PNePsPiPs
PsPrPSPsPeCĤP°C,PSPsPiPs PiPμCĤPsCĤP°P»CHbPSPsPiPs P·P°CĬC,PsCĬCŕPIP°PSPSCĬ Cŕ
PrPsP·C- 50 PjPi. PŸPμCĤPμPrPSC- AUC C,P° C_{max} N-PrPμPjPμC,PëP»CHbPsPIP°PSPsPiPs
PjPμC,P°P±PsP»C-C,Cŕ PiC-PrPIPëC‰CŕPIP°P»PëCĬCĬ PIC-PrPiPsPIC-PrPSPs PSP°
PjP°PeCĬPëPjCŕPj 126 % C,P° PjP°PeCĬPëPjCŕPj 73 % PiPsCĤC-PIPSCĬPSPs P·C,P°PePëPjPë
PiPsPeP°P·PSPëPeP°PjPë Cŕ PrPsP±CĤPsPIPsP»CHC†C-PI C,P°PePsPiPs P¶ PIC-PeCŕ P±PμP·
PiPsCĤCŕC€PμPSCĤ C,,CŕPSPeC†C-C— PSPëCĤPsPe. PhPrPSP°Pe C†PμCĤPμP· PIPëCĬPsPeCŕ
C-PSPrPëPIC-PrCŕP°P»CHbPSCŕ PIP°CĤC-P°P±PμP»CHbPSC-CĬC,CH C†C-
PIC-PrPjC-PSPSPsCĬC,C- PSPμ P±CŕP»Pë CĬC,P°C,PëCĬC,PëC†PSPs P·PSP°C†CŕC‰PëPjPë. PJ
PrPsP±CĤPsPIPsP»CHC†C-PI C-P· C,CĬP¶PePëPjPë PiPsCĤCŕC€PμPSPSCĬPjPë
C,,CŕPSPeC†C-C— PSPëCĤPsPe (PeP»C-CĤPμPSCĬ PeCĤPμP°C,PëPSC-PSCŕ PSPëP¶C†Pμ 30
PjP»/C...PI) PeP»C-CĤPμPSCĬ CĬPëP»PrPμPSP°C,,C-P»Cŕ P·PSPëP¶CŕPIP°PICĬCĬ, C‰Ps
PiCĤPëP·PiPsPrPëP»Ps PrPs CĬPμCĤPμPrPSC-C... PiC-PrPIPëC‰PμPSCĤ AUC C,P° C_{max}
PIC-PrPiPsPIC-PrPSPs PSP° 100 % C,P° 88 % PiPsCĤC-PIPSCĬPSPs P·
PrPsP±CĤPsPIPsP»CHC†CĬPjPë C,P°PePsPiPs P¶ PIC-PeCŕ P±PμP· PiPsCĤCŕC€PμPSCĤ
C,,CŕPSPeC†C-C— PSPëCĤPsPe. PĤCĤC-Pj C,PsPiPs, P·PSP°C†PμPSPSCĬ AUC C,P° C_{max} N-
PrPμPjPμC,PëP»CHbPsPIP°PSPsPiPs PjPμC,P°P±PsP»C-C,Cŕ P·PSP°C†CŕC‰Pμ
PiC-PrPIPëC‰CŕPIP°P»PëCĬCĬ PIC-PrPiPsPIC-PrPSPs PSP° 200 % C,P°
79 % PIC-PrPiPsPIC-PrPSPs.

РЛЬР»С-РSC-С±РSC- С...Р°СЃР°Р°С,РµСЃРёСЃЃС,РёРёРёРё.

$\text{P}_{\mu}\text{C}\tilde{\text{H}}\text{P}_{\mu}\text{P}\text{iP}^{\circ}\text{C}\tilde{\text{H}}\text{P}^{\circ}\text{C}$, $\text{P}\check{\text{Y}}\text{C}-\text{PSP}_{\mu}\text{P}\text{iC}\tilde{\text{H}}\text{P}^{\circ}\text{C}\tilde{\text{H}}\text{P}_{\mu}\text{P}\text{ePsPjP}_{\mu}\text{PSPrC}\acute{\text{r}}\text{C}^{\circ}\text{C}$, $\text{C}\tilde{\text{H}}\text{C}\acute{\text{I}}\text{C}\Pi$
 $\text{P}\cdot\text{P}^{\circ}\text{C}\acute{\text{I}}\text{C}$, $\text{PsC}\acute{\text{I}}\text{PsPIC}\acute{\text{r}}\text{PIP}^{\circ}\text{C}$, $\text{P}\check{\text{e}}\text{C}\ddagger\text{PsP}\gg\text{PsPIC}-\text{PeP}^{\circ}\text{PjP}\cdot\text{P}_{\mu}\text{C}\tilde{\text{H}}\text{P}_{\mu}\text{PeC}$, $\text{P}\check{\text{e}}\text{P}\gg\text{C}\tilde{\text{H}}\text{PSPsC}\tilde{\text{H}}$
 $\text{PrP}\check{\text{e}}\text{C}\acute{\text{I}}\text{C}$, „ $\text{C}\acute{\text{r}}\text{PSPeC}\ddagger\text{C}-\text{C}^{\circ}\text{C}\tilde{\text{H}}$, $\text{C}\Pi\text{PeP}^{\circ}\text{PIP}\check{\text{e}}\text{P}\cdot\text{PSP}^{\circ}\text{C}\ddagger\text{P}^{\circ}\text{C}^{\circ}\text{C}$, $\text{C}\tilde{\text{H}}\text{C}\acute{\text{I}}\text{C}\Pi$ $\text{C}\Pi\text{Pe}$
 $\text{PSP}_{\mu}\text{P}\cdot\text{PrP}^{\circ}\text{C}$, $\text{PSC}-\text{C}\acute{\text{I}}\text{C}$, $\text{C}\tilde{\text{H}}\text{PrPsC}\acute{\text{I}}\text{C}\Pi\text{PiC}$, $\text{P}\check{\text{e}}\text{P}^{\circ}\text{P}\pm\text{PsPiC}-\text{PrC}$, $\text{C}\tilde{\text{H}}\text{P}\check{\text{e}}\text{PjP}^{\circ}\text{C}$, $\text{P}\check{\text{e}}\text{P}_{\mu}\text{C}\tilde{\text{H}}\text{P}_{\mu}\text{PeC}\ddagger\text{C}-\text{C}\tilde{\text{H}}$
 $\text{C}\acute{\text{I}}\text{C}$, P°C , $\text{P}_{\mu}\text{PIP}\text{PsPiPsC}\ddagger\text{P}\gg\text{P}_{\mu}\text{PSP}^{\circ}$, $\text{PSP}_{\mu}\text{PsP}\pm\text{C}\dots\text{C}-\text{PrPSC}\acute{\text{r}}\text{PrP}\gg\text{C}\Pi\text{C}\acute{\text{r}}\text{C}\acute{\text{I}}\text{P}\check{\text{r}}\text{C}-\text{C}\epsilon\text{PSPsPiPs}$
 $\text{C}\acute{\text{I}}\text{C}$, P°C , $\text{P}_{\mu}\text{PIP}\text{PsPiPsP}^{\circ}\text{PeC}$, $\text{C}\acute{\text{r}}$.

$$P''P \gg Cl, PuC, PuPeC, PePIPSPsC - PrC - C - PiC_7PuPiP^{\circ}C_7P^{\circ}C, CrP\check{Y}C - PSPuPiC_7P^{\circ}PiPsC, C_7C - P \pm PSPu, C_7PuPeC_7CrP^{\circ}P \gg CHbPSPu, P \cdot P \pm CrPrP[PuPSPSCl.$$

РμCḡPsC, PëPīPsPeP°P·P°PSPSCII.

вḡ “ РμC-PrPIPëC%PμPSP° C†CṛC, P»PëPIC-ClṛC, Ch PrPs P°PeC, PëPIPSPsC— CḡPμC†PsPIPëPSPë P°P±Ps PrPs P±CṛPrCh-ClPePsC— P· PrPsPīPsPjC-PḡPSPëC... CḡPμC†PsPIPëPS PiCḡPμPiP°CḡP°C, Cṛ.

вḡ “ PhPrPSPsC†P°ClṛPSPμ P·P°ClṛC, PsClṛCṛPIP°PSPSCII P· PrPsPSPsCḡP°PjPë PsPeClṛPëPrCṛ P°P·PsC, Cṛ (C, P°PePëPjPë ClPe P°PjC-P»PSC-C, CḡPëC,) P°P±Ps PSC-C, CḡP°C, P°PjPë Cṛ P±CṛPrCh-ClPeC-PNḡ C,, PsCḡPjC-, PsClṛPeC-P»ChPePë PIC-PrPsPjPs, C%Ps ClṛPëP»PrPμPSP°C,, C-P» PIPīP»PëPIP°C” PSP° CḡP»ClC...Pë PjPμC, P°P±PsP»C-P·PjCṛ PsPeClṛPëPrCṛ P°P·PsC, Cṛ/C†PëPeP»C-C†PSPsPiPs PiCṛP°PSPsP·PëPSPjPsPSPsC,, PsClṛC,, P°C, Cṛ (C†P“PḡPḡ) C, P° PīPsC, PμPSC†C-CḡC” PiC-PīPsC, PμPSP·PëPIPSPëPNḡ PμC,, PμPeC, PSC-C, CḡP°C, C-PI.

вḡ “ PhPrPSPsC†P°ClṛPSPμ P·P°ClṛC, PsClṛCṛPIP°PSPSCII C-PSPiC-P±C-C, PsCḡC-PI PḡP”P·5 (Cṛ C, PsPjCṛ C†PëClṛP»C- ClṛPëP»PrPμPSP°C,, C-P»Cṛ) P·C- ClṛC, PëPjCṛP»ClC, PsCḡP°PjPë PiCṛP°PSC-P»P°C, C†PëPeP»P°P·Pë, C, P°PePëPjPë ClPe CḡC-PsC†PëPiCṛP°C,, PiCḡPsC, PëPīPsPeP°P·P°PSPμ, PsClṛPeC-P»ChPePë PjPsPḡPμ PiCḡPëP·PIPμClṛC, Pë PrPs ClṛPëPjPiC, PsPjP°C, PëC†PSPsC— PiC-PīPsC, PμPSP·C-C— (PrPëPI. CḡPsP·PrC-P» B«P·P·P°C”PjPsPrC-Cl P· C-PSCḡPëPjPë P»C-PeP°CḡClṛChPePëPjPë P·P°ClṛPsP±P°PjPë C, P° C-PSCḡC- PIPëPrPë PIP·P°C”PjPsPrC-PNḡB»).

вḡ “ PḢC, P°PSPë, PiCḡPë ClPePëC... PSPμ CḡPμPePsPjPμPSPPrPsPIP°PSP° ClṛPμPeClṛCṛP°P»ChPSP° P°PeC, PëPIPSC-ClṛC, Ch (PSP°PiCḡPëPeP»P°Pr C, ClPḡPeC- ClṛPμCḡC†PμPIPs-ClṛCṛPrPëPSPSC- CḡPsP·P»P°PrPë, C, P°PeC- ClPe PSPμClṛC, P°P±C-P»ChPSP° ClṛC, PμPSPsPeP°CḡPrC-Cl P°P±Ps ClṛPμCḡC†PμPIP° PSPμPrPsClṛC, P°C, PSC-ClṛC, Ch C, ClPḡPePsPiPs ClṛC, CṛPīPμPSCII).

вḡ “ P·C, CḡP°C, P° P·PsCḡCṛ PSP° PsPrPSPμ PsPePs PIPSP°ClṛP»C-PrPsPe PSPμP°CḡC, PμCḡC-P°P»ChPSPsC— PīPμCḡPμPrPSCḡPsC— C-CḡPμPjC-C†PSPsC— PSPμPICḡPsPīP°C, C-C— P·PsCḡPsPiPsPiPs PSPμCḡPIP° PSPμP·P°P»PμPḡPSPs PIC-Pr C, PsPiPs, PīPsPiḡḡ™ ClP·P°PSP° C†Cl PīP°C, PsP»PsPiC-Cl C-P· PīPsPīPμCḡPμPrPSC-Pj P·P°ClṛC, PsClṛCṛPIP°PSPSCII Pj C-PSPiC-P±C-C, PsCḡC-PI PḡP”P·5 C†Pë PSC-.

вḡ “ PḡP°ClṛPIPSC-ClṛC, Ch C, P°PePëC... P·P°C...PIPsCḡCḡPIP°PSCḡ ClPe PīPsCḡCṛCḡPμPSPSCII C,, CṛPSPeC†C-C— PīPμC†C-PSPePë C, ClPḡPePsPiPs ClṛC, CṛPīPμPSCII, P°CḡC, PμCḡC-P°P»ChPSP° PiC-PīPsC, PμPSP·C-Cl (P°CḡC, PμCḡC-P°P»ChPSPëPNḡ C, PëClṛPe PSPëPḡC†Pμ 90/50 PjPj CḡC,, ClṛC, .), PSPμC%PsPrP°PIPSPs PīPμCḡPμPSPμClṛPμPSPëPNḡ C-PSCṛCṛP»ChC, P°P±Ps C-PSC,, P°CḡPeC, PjC-PsPeP°CḡPrP° C, P° PIC-PrPsPjC- ClṛPīP°PrPePsPIC- PrPμPiPμPSPμCḡP°C, PëPIPSC- P·P°C...PIPsCḡCḡPIP°PSPSCII ClṛC-C, PeC-PIPePë, C, P°PeC- ClPe PīC-PiPjPμPSC, PSPëPNḡ CḡPμC, PëPSC-C, (PSPμPIPμP»PëPeP° PeC-P»ChPeC-ClṛC, Ch C, P°PePëC... PīP°C†C-C”PSC, C-PI PjP°C” PiPμPSPμC, PëC†PSC- CḡPsP·P»P°PrPë C,, PsClṛC,, PsPrC-PμClṛC, PμCḡP°P· ClṛC-C, PeC-PIPePë), PsClṛPeC-P»ChPePë P±PμP·PiPμPeP° ClṛPëP»PrPμPSP°C,, C-P»Cṛ PSPμ PrPsClṛP»C-PrPḡCṛPIP°P»P°ClṛCl Cṛ C, P°PePëC... PiC-PrPiCḡCṛPīP°C... PīP°C†C-C”PSC, C-PI.

P·P·P°C”PjPsPrC-Cl P· C-PSCḡPëPjPë P»C-PeP°CḡClṛChPePëPjPë P·P°ClṛPsP±P°PjPë C, P° C-PSCḡC- PIPëPrPë PIP·P°C”PjPsPrC-PNḡ.

$P \cdot C, P_{\mu}PeC, P\ddot{e} C - PSC\epsilon P\ddot{e}C \dots P \gg C - PeP^{\circ} C\tilde{H}C\tilde{I}CHPeP\ddot{e}C \dots P \cdot P^{\circ} C\tilde{I}PsP \pm C - PI PSP^{\circ}$
 $C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg$.

$P''PsC\tilde{I}P \gg C - PrP_{\mu}P_{\mu}PSPSC\tilde{I} in vitro$. $P_{\mu}P_{\mu}C, P^{\circ}P \pm PsP \gg C - P \cdot P_j C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I}$
 $PIC - PrP \pm C\tilde{I}PIP^{\circ}C''C, CHC\tilde{I}C\tilde{I} P\ddot{I}P_{\mu}C\tilde{H}P_{\mu}PIP^{\circ}P_{\mu}PSPs P \cdot C\tilde{I}C \pm P^{\circ}C\tilde{I}C, C\tilde{H} C - P \cdot PsC, PsC\tilde{H}P_jP\ddot{e} 3P_{\mu}4$
 $(PiPsP \gg PsPIPSP\ddot{e}PN_{\epsilon} C\epsilon P \gg C\tilde{I}C \dots) C, P^{\circ} C - P \cdot PsC, PsC\tilde{H}P_jP\ddot{e} 2P\check{Y}9 (PrC\tilde{H}C\tilde{I}PiPsC\tilde{H}C\tilde{I}PrPSP\ddot{e}PN_{\epsilon}$
 $C\epsilon P \gg C\tilde{I}C \dots) C \dagger P\ddot{e}C, PsC \dots C\tilde{H}PsP_jC\tilde{I} P 450 (CYP). P\check{Y}PsP_jC\tilde{I} C - PSPiC - P \pm C - C, PsC\tilde{H}P\ddot{e} C \dagger P\ddot{e}C \dots$
 $C - P \cdot PsC, P_{\mu}C\tilde{H}P_jP_{\mu}PSC, C - PI P_jPsP_{\mu}C\tilde{I}C, CH P \cdot PSP\ddot{e}P_{\mu}C\tilde{I}PIP^{\circ}C, P\ddot{e} PeP \gg C - C\tilde{H}P_{\mu}PSC\tilde{I}$
 $C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I}, P^{\circ} C - PSPPrC\tilde{I}PeC, PsC\tilde{H}P\ddot{e} C \dagger P\ddot{e}C \dots C - P \cdot PsC, P_{\mu}C\tilde{H}P_jP_{\mu}PSC, C - PI$
 $P_jPsP_{\mu}C\tilde{I}C, CH PiC - PrPIP\ddot{e}C \% C\tilde{I}PIP^{\circ}C, P\ddot{e} PeP \gg C - C\tilde{H}P_{\mu}PSC\tilde{I} C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I}.$

$P''PsC\tilde{I}P \gg C - PrP_{\mu}P_{\mu}PSPSC\tilde{I} in vivo$. $P_{\mu}PsPiC\tilde{I}P \gg C\tilde{I}C \dagger C - PN_{\epsilon}PSP\ddot{e}PN_{\epsilon}$
 $C, P^{\circ}C\tilde{H}P_jP^{\circ}PePsPeC - PSP_{\mu}C, P\ddot{e}C \dagger PSP\ddot{e}PN_{\epsilon} P^{\circ}PSP^{\circ}P \gg C - P \cdot PrP^{\circ}PSP\ddot{e}C \dots$
 $PeP \gg C - PSC - C \dagger PSP\ddot{e}C \dots PrPsC\tilde{I}P \gg C - PrP_{\mu}P_{\mu}PSC\tilde{H} PiC\tilde{H}PsPrP_{\mu}P_jPsPSC\tilde{I}C, C\tilde{H}C\tilde{I}PIP^{\circ}PI$
 $P \cdot PSP\ddot{e}P_{\mu}P_{\mu}PSPSC\tilde{I} PeP \gg C - C\tilde{H}P_{\mu}PSC\tilde{I}C\tilde{I} C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I} PiC\tilde{H}P\ddot{e} PN_{\epsilon}PsPiPs$
 $PsPrPSPsC \dagger P^{\circ}C\tilde{I}PSPsP_jC\tilde{I} P \cdot P^{\circ}C\tilde{I}C, PsC\tilde{I}C\tilde{I}PIP^{\circ}PSPSC - P \cdot C - PSPiC - P \pm C - C, PsC\tilde{H}P^{\circ}P_jP\ddot{e}$
 $CYP3A4 (C, P^{\circ}PeP\ddot{e}P_jP\ddot{e} C\tilde{I}Pe PeP_{\mu}C, PsPePsPSP^{\circ}P \cdot PsP \gg, P_{\mu}C\tilde{H}P\ddot{e}C, C\tilde{H}PsP_jC - C \dagger P\ddot{e}PS,$
 $C \dagger P\ddot{e}P_jP_{\mu}C, P\ddot{e}PrP\ddot{e}PS). P\check{I}PsC \dagger P^{\circ} PiC\tilde{H}P\ddot{e} PsPrPSPsC \dagger P^{\circ}C\tilde{I}PSPsP_jC\tilde{I}$
 $P \cdot P^{\circ}C\tilde{I}C, PsC\tilde{I}C\tilde{I}PIP^{\circ}PSPSC - C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I} C, P^{\circ} C - PSPiC - P \pm C - C, PsC\tilde{H}C - PI$
 $CYP3A4 P \cdot C\tilde{H}PsC\tilde{I}C, P^{\circ}PSPSC\tilde{I} C \dagger P^{\circ}C\tilde{I}C, PsC, P\ddot{e} PiPsP \pm C - C \dagger PSP\ddot{e}C \dots C\tilde{I}PIP\ddot{e}C \% PSP_{\mu}$
 $C\tilde{I}PiPsC\tilde{I}C, P_{\mu}C\tilde{H}C - PiP^{\circ}P \gg PsC\tilde{I}C\tilde{I}, C\tilde{I}P \gg C - Pr C\tilde{H}PsP \cdot PiP \gg C\tilde{I}PSC\tilde{I}C, P\ddot{e}$
 $P_jPsP_{\mu}P \gg P\ddot{e}PIC - C\tilde{I}C, CH P \cdot P^{\circ}C\tilde{I}C, PsC\tilde{I}C\tilde{I}PIP^{\circ}PSPSC\tilde{I} PiPsC \dagger P^{\circ}C, PePsPIPsC - PrPsP \cdot P\ddot{e}$
 $C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I} 25 P_jPi.$

$PhPrPSPsC \dagger P^{\circ}C\tilde{I}PSP_{\mu} P \cdot P^{\circ}C\tilde{I}C, PsC\tilde{I}C\tilde{I}PIP^{\circ}PSPSC\tilde{I} C - PSPiC - P \pm C - C, PsC\tilde{H}C\tilde{I} P^{\circ}P \dagger P \gg$
 $PiC\tilde{H}PsC, P_{\mu}P^{\circ}P \cdot P\ddot{e} C\tilde{H}P\ddot{e}C, PsPSP^{\circ}PIC - C\tilde{H}C\tilde{I}, PrC\tilde{I}P_{\mu}PiPsC, C\tilde{I}P_{\mu}PSPsPiPs$
 $C - PSPiC - P \pm C - C, PsC\tilde{H}C\tilde{I} P 450, C\tilde{I} C\tilde{I}C, P^{\circ}PSC - C\tilde{H}C - PIPSPsPIP^{\circ}P_{\mu}PSPsC -$
 $PePsPSC \dagger P_{\mu}PSC, C\tilde{H}P^{\circ}C \dagger C - C - (500 P_jPi 2 C\tilde{H}P^{\circ}P \cdot P\ddot{e} PSP^{\circ} PrPsP \pm C\tilde{I}) C, P^{\circ}$
 $C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I} (C\tilde{H}P^{\circ}P \cdot PsPIP^{\circ} PrPsP \cdot P^{\circ} 100 P_jPi) PiC\tilde{H}P\ddot{e}P \cdot PIPsPrP\ddot{e}P \gg Ps PrPs$
 $PiC - PrPIP\ddot{e}C \% P_{\mu}PSPSC\tilde{I} C_{max} C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I} PSP^{\circ} 300 \% (C\tilde{I} 4 C\tilde{H}P^{\circ}P \cdot P\ddot{e}) C, P^{\circ}$
 $PiC - PrPIP\ddot{e}C \% P_{\mu}PSPSC\tilde{I} PiP \gg P^{\circ}P \cdot P_jPsPIPsC - AUC C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I} PSP^{\circ} 1000 \% (PI$
 $11 C\tilde{H}P^{\circ}P \cdot C - PI). P\check{S}P_{\mu}C\tilde{H}P_{\mu}P \cdot 24 PiPsPrP\ddot{e}PSP\ddot{e} PiP \gg P^{\circ}P \cdot P_jPsPIC - C\tilde{H}C - PIPSC -$
 $C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I} PIC\tilde{I}P_{\mu} C \% P_{\mu} C\tilde{I}C, P^{\circ}PSPsPiP\ddot{e}P \gg P\ddot{e} PiC\tilde{H}P\ddot{e}P \pm P \gg P\ddot{e}P \cdot PSPs 200$
 $PSPi/P_jP \gg PiPsC\tilde{H}C - PIPSC\tilde{I}PSPs P \cdot C\tilde{H}C - PIPSP_{\mu}P_j PiC\tilde{H}P\ddot{e}P \pm P \gg P\ddot{e}P \cdot PSPs 5 PSPi/P_jP \gg,$
 $C \dots P^{\circ}C\tilde{H}P^{\circ}PeC, P_{\mu}C\tilde{H}PSP\ddot{e}P_j PrP \gg C\tilde{I} P \cdot P^{\circ}C\tilde{I}C, PsC\tilde{I}C\tilde{I}PIP^{\circ}PSPSC\tilde{I} C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I}$
 $PsPeC\tilde{H}P_{\mu}P_jPs, C \% Ps PIC - PrPiPsPIC - PrP^{\circ}C'' P \cdot PSP^{\circ}C \dagger PSPsP_jC\tilde{I} PIPiP \gg P\ddot{e}PIC\tilde{I}$
 $C\tilde{H}P\ddot{e}C, PsPSP^{\circ}PIC - C\tilde{H}C\tilde{I} PSP^{\circ} C\epsilon P\ddot{e}C\tilde{H}PsPeP\ddot{e}PN_{\epsilon} C\tilde{I}PiP_{\mu}PeC, C\tilde{H} C\tilde{I}C\tilde{I}P \pm C\tilde{I}C, C\tilde{H}P^{\circ}C, C - PI P 450.$
 $P\check{Y}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg PSP_{\mu} PIPiP \gg P\ddot{e}PIP^{\circ}C'' PSP^{\circ} C, P^{\circ}C\tilde{H}P_jP^{\circ}PePsPeC - PSP_{\mu}C, P\ddot{e}PeC\tilde{I}$
 $C\tilde{H}P\ddot{e}C, PsPSP^{\circ}PIC - C\tilde{H}C\tilde{I}. P - PsPiP \gg C\tilde{I}PrC\tilde{I} PSP^{\circ} C \dagger C - C, P^{\circ}C\tilde{H}P_jP^{\circ}PePsPeC - PSP_{\mu}C, P\ddot{e}C \dagger PSC -$
 $PrP^{\circ}PSC - PsPrPSPsC \dagger P^{\circ}C\tilde{I}PSP_{\mu} P \cdot P^{\circ}C\tilde{I}C, PsC\tilde{I}C\tilde{I}PIP^{\circ}PSPSC\tilde{I} C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I} C, P^{\circ}$
 $C\tilde{H}P\ddot{e}C, PsPSP^{\circ}PIC - C\tilde{H}C\tilde{I} PSP_{\mu} C\tilde{H}P_{\mu}PePsP_jP_{\mu}PSPPrC\tilde{I}C''C, CHC\tilde{I}C\tilde{I} (PrP\ddot{e}PI. C\tilde{H}PsP \cdot PrC - P \gg$
 $B \ll PhC\tilde{I}PsP \pm P \gg P\ddot{e}PIPsC\tilde{I}C, C - P \cdot P^{\circ}C\tilde{I}C, PsC\tilde{I}C\tilde{I}PIP^{\circ}PSPSC\tilde{I}B \gg); C\tilde{I} P \pm C\tilde{I}PrCHC - C\tilde{I}PePsP_jC\tilde{I}$
 $PIP\ddot{e}PiP \cdot PrPeC\tilde{I} P_jP^{\circ}PeC\tilde{I}P\ddot{e}P_jP^{\circ}P \gg CHPSP^{\circ} PrPsP \cdot P^{\circ} C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I} P \cdot P^{\circ}$
 $P_{\mu}PsPrPSP\ddot{e}C \dots PsP \pm C\tilde{I}C, P^{\circ}PIP\ddot{e}PS PSP_{\mu} PiPsPIP\ddot{e}PSPSP^{\circ} PiP_{\mu}C\tilde{H}P_{\mu}PIP\ddot{e}C \% C\tilde{I}PIP^{\circ}C, P\ddot{e} 25 P_jPi$
 $PiC\tilde{H}PsC, C\tilde{I}PiPsP_j 48 PiPsPrP\ddot{e}PS.$

$PhPrPSPsC \dagger P^{\circ}C\tilde{I}PSP_{\mu} P \cdot P^{\circ}C\tilde{I}C, PsC\tilde{I}C\tilde{I}PIP^{\circ}PSPSC\tilde{I} C - PSPiC - P \pm C - C, PsC\tilde{H}C\tilde{I} P^{\circ}P \dagger P \gg$
 $PiC\tilde{H}PsC, P_{\mu}P^{\circ}P \cdot P\ddot{e} C\tilde{I}P^{\circ}PePIC - PSP^{\circ}PIC - C\tilde{H}C\tilde{I}, C - PSPiC - P \pm C - C, PsC\tilde{H}C\tilde{I} CYP3A4, C\tilde{I} PrPsP \cdot C -,$
 $C \% Ps P \cdot P^{\circ}P \pm P_{\mu}P \cdot PiP_{\mu}C \dagger C\tilde{I}C'' C\tilde{H}C - PIPSPsPIP^{\circ}P_{\mu}PSC\tilde{I} PePsPSC \dagger P_{\mu}PSC, C\tilde{H}P^{\circ}C \dagger C - CH (1200 P_jPi 3$
 $C\tilde{H}P^{\circ}P \cdot P\ddot{e} PSP^{\circ} PrPsP \pm C\tilde{I}), C, P^{\circ} C\tilde{I}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C - P \gg C\tilde{I} (100 P_jPi C\tilde{H}P^{\circ}P \cdot PsPIPs)$

ΠΐCΉPēP·PIPsPrPēP»Ps PrPs ΠΐC-PrPIPēC%PμPSPSCΠ C_{max} CΉPēP»PrPμPSP°C,,C-P»CΉ PSP°
140 % C,P° P·P±C-P»CHC€PμPSPSCΠ CΉPēCΉC,PμPjPSPsC— PμPeCΉΠΐPsP·PēC†C-C— (AUC)
CΉPēP»PrPμPSP°C,,C-P»CΉ PSP° 210 %. PκPμ PIPēCΠIP»PμPSPs PIPΐP»PēPICΉ
CΉPēP»PrPμPSP°C,,C-P»CΉ PSP° C,,P°CΉPjP°PePsPeC-PSPμC,PēPeCΉ CΉP°PePIC-PSP°PIC-CΉCΉ
(PrPēPI. CΉPsP·PrC-P» B«PΨΠΐPsCΉC-P± P·P°CΉC,PsCΉCΉPIP°PSPSCΠ C,P° PrPsP·PēB»).

PμPμCΉPμPrP±P°C†P°C”C,CHCΉCΠ, C%Ps P±C-P»CHC€ ΠΐPsC,CΉPΠPSC-
C-PSPiC-P±C-C,PsCΉPē CYP3A4, C,P°PeC- CΠPe PePμC,PsPePsPSP°P·PsP» C,P°
C-C,CΉP°PePsPSP°P·PsP», P±CΉPrCΉC,CH PjP°C,Pē P±C-P»CHC€ PIPēCΉP°PΠPμPSPēPNē
PIΠP»PēPI.

PμCΉPē P·P°CΉC,PsCΉCΉPIP°PSPSC- CΉPēP»PrPμPSP°C,,C-P»CΉ (100 PjPi CΉP°P·PsPIPs) C,P°
PμCΉPēC,CΉPsPjC-C†PēPSCΉ, ΠΐPsPjC-CΉPSPsPiPs C-PSPiC-P±C-C,PsCΉCΉ CYP3A4, CΉ
CΉC-PIPSPsPIP°PΠPSPsPjCΉ CΉC,P°PSC- (500 PjPi 2 CΉP°P·Pē PSP° PrPsP±CΉ ΠΐCΉPsC,CΠΠΐPsPj
5 PrPSC-PI) CΉΠΐPsCΉC,PμCΉC-ΠP°P»PsCΉCΠ ΠΐC-PrPIPēC%PμPSPSCΠ CΉPēCΉC,PμPjPSPsC—
PμPeCΉΠΐPsP·PēC†C-C— CΉPēP»PrPμPSP°C,,C-P»CΉ PSP° 182 % (PΉUC). Pj P·PrPsCΉPsPIPēC...
PrPsP±CΉPsPIPsP»CHC†C-PI C†PsP»PsPIC-C†PsC— CΉC,P°C,C- PSPμ
CΉΠΐPsCΉC,PμCΉC-ΠP°P»PsCΉCΠ PIPΐP»PēPICΉ P°P·PēC,CΉPsPjC-C†PēPSCΉ (500 PjPi PSP°
PrPsP±CΉ ΠΐCΉPsC,CΠΠΐPsPj 3 PrC-P±) PSP° AUC, PΨ_{max}, T_{max}, PePsPSCΉC,P°PSC,CΉ
C€PIPēPrPePsCΉC,C- PμP»C-PjC-PSP°C†C-C— C,P° ΠΐPsPrP°P»CHC€PēPNē ΠΐPμCΉC-PsPr
PSP°ΠΐC-PIPIPēPIPμPrPμPSPSCΠ CΉPēP»PrPμPSP°C,,C-P»CΉ P°P±Ps PNēPsPiPs
PiPsP»PsPIPsPiPs C†PēCΉPeCΉP»CHCΉC†PsPiPs PjPμC,P°P±PsP»C-C,CΉ. PjPēPjPμC,PēPrPēPS
(C-PSPiC-P±C-C,PsCΉ C†PēC,PsC...CΉPsPjCΉ P 450 C,P° PSPμCΉΠΐPμC†PēC,,C-C†PSPēPNē
C-PSPiC-P±C-C,PsCΉ CYP3A4) CΉ PrPsP·C- 800 PjPi ΠΐCΉPē PsPrPSPsC†P°CΉPSPsPjCΉ
P·P°CΉC,PsCΉCΉPIP°PSPSC- C-P· CΉPēP»PrPμPSP°C,,C-P»PsPj CΉ PrPsP·C- 50 PjPi CΉ
P·PrPsCΉPsPIPēC... PrPsP±CΉPsPIPsP»CHC†C-PI ΠΐCΉPēP·PIPsPrPēPI PrPs
ΠΐC-PrPIPēC%PμPSPSCΠ ΠΐP»P°P·PjPsPIPsC— PePsPSC†PμPSC,CΉP°C†C-C—
CΉPēP»PrPμPSP°C,,C-P»CΉ PSP° 56 %.

P”CΉPμPNēΠΐC,,CΉCΉC,PsPIPēPNē CΉC-Pe C” CΉP»P°P±PePēPj C-PSPiC-P±C-C,PsCΉPsPj CYP3A4
CΉ CΉC,C-PSC†C- PePēC€PμC†PSPēPeCΉ C- PjPsPΠPμ CΉΠΐCΉPēC†PēPSCΠC,Pē ΠΐPsPjC-CΉPSPμ
ΠΐC-PrPIPēC%PμPSPSCΠ CΉC-PIPSCΠ CΉPēP»PrPμPSP°C,,C-P»CΉ CΉ ΠΐP»P°P·PjC- PeCΉPsPIC-.

PΉPrPSPsCΉP°P·PsPIPμ P·P°CΉC,PsCΉCΉPIP°PSPSCΠ P°PSC,P°C†PēPrC-PI (PjP°ΠΐPSC-CΉ
ΠΐC-PrCΉPsPeCΉPēPrCΉ/P°P»CΉPjC-PSC-CΉ ΠΐC-PrCΉPsPeCΉPēPrCΉ) PSPμ PIPΐP»PēPIP°P»Ps
PSP° P±C-PsPrPsCΉC,CΉΠΐPSC-CΉC,CH CΉPēP»PrPμPSP°C,,C-P»CΉ.

ΠΓPsC†P° PrPsCΉP»C-PrPΠPμPSCΉ CΉΠΐPμC†PēC,,C-C†PSPsC— PIP·P°C”PjPsPrC-C— P·
CΉCΉC-PjP° P»C-PeP°CΉCΉCHPePēPjPē P·P°CΉPsP±P°PjPē PSPμ ΠΐCΉPsPIPsPrPēP»Pē, P·P°
PrP°PSPēPjPē ΠΐPsΠΐCΉP»CΉC†C-PNēPSPsPiPs C,,P°CΉPjP°PePsPeC-PSPμC,PēC†PSPsPiPs
P°PSP°P»C-P·CΉ, C,,P°CΉPjP°PePsPeC-PSPμC,PēPeP° CΉPēP»PrPμPSP°C,,C-P»CΉ PSPμ
P·PjC-PSCΉPIP°P»P°CΉCΠ ΠΐCΉPē PNēPsPiPs PsPrPSPsC†P°CΉPSPsPjCΉ
P·P°CΉC,PsCΉCΉPIP°PSPSC- P· P»C-PeP°CΉCΉCHPePēPjPē P·P°CΉPsP±P°PjPē, C%Ps
PSP°P»PμPΠP°C,CH PrPs ΠΐCΉCΉΠΐPē C-PSPiC-P±C-C,PsCΉC-PI CYP2C9 (C,PsP»P±CΉC,P°PjC-Pr,
PIP°CΉC,,P°CΉPēPS, C,,PμPSC-C,PsC—PS), ΠΐCΉCΉΠΐPē C-PSPiC-P±C-C,PsCΉC-PI CYP2D6
(C,P°PePēC... CΠPe CΉPμP»PμPeC,PēPIPSC- C-PSPiC-P±C-C,PsCΉPē P·PIPsCΉPsC,PSPsPiPs
P·P°C...PsΠΐP»PμPSPSCΠ CΉPμCΉPsC,PsPSC-PSCΉ (PΨ†P—P—PΨ), C,CΉPēC†PēPeP»C-C†PSC-
P°PSC,PēPrPμΠΐCΉPμCΉP°PSC,Pē), ΠΐCΉCΉΠΐPē C,C-P°P·PēPrPSPēC... C,P°
C,C-P°P·PēPrPsPiPsPrC-P±PSPēC... PrC-CΉCΉPμC,PēPeC-PI, ΠΐPμC,P»CHPsPIPēC... C,P°
PeP°P»C-PNēP·P±PμCΉC-ΠP°CΉC†PēC... PrC-CΉCΉPμC,PēPeC-PI, C-PSPiC-P±C-C,PsCΉC-PI
P°PSPiC-PsC,PμPSP·PēPSPiPμCΉPμC,PIPsCΉCHPIP°P»CHPSPsPiPs C,,PμCΉPjPμPSC,CΉ,
P°PSC,P°ΠΐPsPSC-CΉC,C-PI PeP°P»CHC†C-CΉ, P°PSC,P°ΠΐPsPSC-CΉC,C-PI β-
P°PrCΉPμPSPsCΉPμC†PμΠΐC,PsCΉC-PI P°P±Ps C-PSPrCΉPeC,PsCΉC-PI PjPμC,P°P±PsP»C-P·PjCΉ

CYP450 (C,P°PePëC... CΠPe CḡPëC,,P°PjPīC-C†PëPS, P±P°CḡP±C-C,CrCḡP°C,Pë).

PJ C...PsPrC- PrPsCÍP»C-PrPΠPμPSPSCΠ P· CrC‡P°CÍC,Cḡ P·PrPsCḡPsPIPëC...
PrPsP±CḡPsPIPsP»CHC†C-PI-C‡PsP»PsPIC-PeC-PI PsPrPSPSc‡P°CÍPSPμ
P·P°CÍC,PsCÍCrPIP°PSPSCΠ P°PSC,P°PiPsPSC-CÍC,P° PμPSPPrPsC,PμP»C-PSCr
P±PsCÍPμPSC,P°PSCr (PiPsPjC-CḡPSPëPNḡ C-PSPrCrPeC,PsCḡ CYP3A4, CYP2C9 C,P°,
PjPsPΠP»PëPIPs, CYP2C19) Cr CḡC-PIPsPPIP°PΠPSPsPjCr CÍC,P°PSC- (125 PjPi 2 CḡP°P·Pë PSP°
PrPsP±Cr) C,P° CÍPëP»PrPμPSP°C,,C-P»Cr Cr CḡC-PIPsPPIP°PΠPSPsPjCr CÍC,P°PSC- (80 PjPi 3
CḡP°P·Pë PSP° PrPsP±Cr) PiCḡPëP·PIPsPrPëP»Ps PrPs P·PSPëPΠPμPSPSCΠ AUC C,P° C_{max}
CÍPëP»PrPμPSP°C,,C-P»Cr PSP° 62,6 % C,P° 55,4 % PIC-PrPiPsPIC-PrPSPs. PŷPsPjCr
PsPrPSPSc‡P°CÍPSPμ P·P°CÍC,PsCÍCrPIP°PSPSCΠ C,P°PePëC... PiPsC,CrPΠPSPëC...
C-PSPrCrPeC,PsCḡC-PI CYP3A4 CΠPe CḡPëC,,P°PjPīC-PS PjPsPΠPμ PiCḡPëP·PIPsPrPëC,Pë PrPs
P±C-P»CHC€ PIPëCḡP°PΠPμPSPsPiPs P·PSPëPΠPμPSPSCΠ PePsPSC†PμPSC,CḡP°C†C-C—
CÍPëP»PrPμPSP°C,,C-P»Cr Cr PiP»P°P·PjC- PeCḡPsPIC-.

PkC-PePsCḡP°PSPPrPëP» CΠPIP»CΠC” CÍPsP±PsCḡ PiC-P±CḡPëPr P°PeC,PëPIP°C,PsCḡP°
PeP°P»C-C”PIPëC... PeP°PSP°P»C-PI C,P° PSC-C,CḡP°C,Cr. PkC-C,CḡP°C,PSPëPNḡ
PePsPjPiPsPSPμPSC, P·CrPjPsPIP»CḡC” PjPsPΠP»PëPIC-CÍC,Cḡ PNḡPsPiPs
CÍPμCḡPNḡPsP·PSPSc— PIP·P°C”PjPsPrC-C— P· CÍPëP»PrPμPSP°C,,C-P»PsPj.

P·C,,PμPeC,Pë CÍPëP»PrPμPSP°C,,C-P»Cž PSP° C-PSC€C- P»C-PeP°CḡCÍCHPeC- P·P°CÍPsP±Pë

P”PsCÍP»C-PrPΠPμPSPSCΠ in vitro. PŷPëP»PrPμPSP°C,,C-P» Bḡ” CÍP»P°P±PePëPNḡ
C-PSPiC-P±C-C,PsCḡ C-P·PsC,,PsCḡPj 1Pḡ2, 2C9, 2C19, 2D6, 2E1 C,P° 3Pḡ4
C†PëC,PsC...CḡPsPjCr P 450 (P†Pḡ₅₀ > 150 PjPePjPsP»CH). PhCÍPeC-P»CHPePë PiC-PePsPIP°
PiP»P°P·PjPsPIP° PePsPSC†PμPSC,CḡP°C†C-CΠ CÍPëP»PrPμPSP°C,,C-P»Cr PrPsCḡC-PIPsCḡC”
PiCḡPëP±P»PëP·PSPs 1 PjPePjPsP»CH, PIPiP»PëPI PiCḡPμPiP°CḡP°C,Cr PŷC-PSPμPiCḡP° PSP°
PeP»C-CḡPμPSCÍ CÍCrP±CÍC,CḡP°C,C-PI C†PëC... C-P·PsC,,PμCḡPjPμPSC,C-PI
PjP°P»PsPNḡPjPsPIC-CḡPSPëPNḡ.

P’C-PrCÍCrC,PSC- PrP°PSC- C%PsPrPs PIP·P°C”PjPsPrC-C— CÍPëP»PrPμPSP°C,,C-P»Cr C,P°
C,P°PePëC... PSPμCÍPiPμC†PëC,,C-C‡PSPëC... C-PSPiC-P±C-C,PsCḡC-PI
C,,PsCÍC,,PsPrC-PμCÍC,PμCḡP°P·Pë, CΠPe C,PμPsC,,C-P»C-PS C,P° PrPëPiC-CḡPëPrP°PjPsP».

P”PsCÍP»C-PrPΠPμPSPSCΠ in vivo. PhCÍPeC-P»CHPePë PIC-PrPsPjPs, C%Ps
CÍPëP»PrPμPSP°C,,C-P» PIPiP»PëPIP°C” PSP° PjPμC,P°P±PsP»C-P·Pj PsPeCÍPëPrCr
P°P·PsC,Cr/C†PëPeP»C-C‡PSPsPiPs PiCrP°PSPsP·PëPSPjPsPSPsC,,PsCÍC,,P°C,Cr (C†P”PḡPα),
P±CrP»Ps PICÍC,P°PSPsPIP»PμPSPs, C%Ps CÍPëP»PrPμPSP°C,,C-P» PiPsC,PμPSC†C-CḡC”
PiC-PiPsC,PμPSP·PëPIPsC† PrC-Cḡ PSC-C,CḡP°C,C-PI, C,PsPjCr PNḡPsPiPs PsPrPSPSc‡P°CÍPSPμ
P·P°CÍC,PsCÍCrPIP°PSPSCΠ P· PrPsPSPsCḡP°PjPë PsPeCÍPëPrCr P°P·PsC,Cr P°P±Ps P·
PSC-C,CḡP°C,P°PjPë Cr P±CrPrCH-CΠPeC-PNḡ C,,PsCḡPjC- PiCḡPsC,PëPiPsPeP°P·P°PSPμ
(PrPëPI. CḡPsP·PrC-P» B«PμCḡPsC,PëPiPsPeP°P·P°PSPSCΠB»).

P C-PsC†PëPiCrP°C,. P”PsPeP»C-PSC-C‡PSC- PrPsCÍP»C-PrPΠPμPSPSCΠ
PiCḡPsPrPμPjPsPSCÍC,CḡCrPIP°P»Pë P°PrPëC,PëPIPsPëPNḡ CÍPëCÍC,PμPjPSPëPNḡ PμC,,PμPeC,
P·PSPëPΠPμPSPSCΠ P°CḡC,PμCḡC-P°P»CHPSPsPiPs C,PëCÍPeCr PiCḡPë
PsPrPSPSc‡P°CÍPSPsPjCr P·P°CÍC,PsCÍCrPIP°PSPSC- C-PSPiC-P±C-C,PsCḡC-PI PαP”P·5 C-P·
CḡC-PsC†PëPiCrP°C,PsPj. PḡP»C-PSC-C‡PSC- PrPsCÍP»C-PrPΠPμPSPSCΠ
PiCḡPsPrPμPjPsPSCÍC,CḡCrPIP°P»Pë, C%Ps CḡC-PsC†PëPiCrP°C, PiPsCÍPëP»CḡC”
PiC-PiPsC,PμPSP·PëPIPsC† PrC-Cḡ C-PSPiC-P±C-C,PsCḡC-PI PαP”P·5. PJ PiP°C†C-C”PSC,C-PI,
CΠPeC- P±CḡP°P»Pë CrC‡P°CÍC,CH Cr PrPsCÍP»C-PrPΠPμPSPSC-, PSPμ
CÍPiPsCÍC,PμCḡC-PiP°P»PsCÍCΠ PiPsP·PëC,PëPIPsPëPNḡ PeP»C-PSC-C‡PSPsPiPs PμC,,PμPeC,Cr

ΠΐCΉPē PsPrPSPsC‡P°CΓPSPsPjCř P·P°CΓC,PsCΓCřPIP°PSPSC- C-PSPIc-P±C-C,PsCΉC-PI PαP°P·5 C-P· CΉC-PsC‡PēPiCřP°C,PsPj. PμCΉPsC,PēPiPsPeP°P·P°PSPμ PsPrPSPsC‡P°CΓPSPμ P·P°CΓC,PsCΓCřPIP°PSPSCΠ CΉC-PsC‡PēPiCřP°C,Cř P· C-PSPIc-P±C-C,PsCΉP°PjPē PαP°P·5 (Cř C,PsPjCř C‡PēCΓP»C- CΓPēP»PrPμPSP°C,,C-P»PsPj) (PrPēPI. CΉPsP·PrC-P»B B«PμCΉPsC,PēPiPsPeP°P·P°PSPSCΠB»).

PħPrPSPsC‡P°CΓPSPμ P·P°CΓC,PsCΓCřPIP°PSPSCΠ CΓPēP»PrPμPSP°C,,C-P»Cř C,P° P±P»PsPeP°C,PsCΉC-PI α-P°PrCΉPμPSPsCΉPμC‡PμPiC,PsCΉC-PI PjPsP¶Pμ ΠΐCΉPēP·PIPμCΓC,Pē PrPs CΉPsP·PIPēC,PeCř CΓPēPjPiC,PsPjP°C,PēC‡PSPsC— PiC-PiPsC,PμPSP·C-C— Cř PrPμCΠPePēC... CΓC...PēP»CHΉPSPēC... PrPs C‡CHΉPsPiPs PiP°C‡C-C''PSC,C-PI. PŷP°PeP° CΉPμP°PeC‡C-CΠ PSP°PNēC‡P°CΓC,C-C€Pμ PIPēPSPēPeP°P»P° PiCΉPsC,CΠPiPsPj 4 PiPsPrPēPS PiC-CΓP»CΠ P·P°CΓC,PsCΓCřPIP°PSPSCΠ CΓPēP»PrPμPSP°C,,C-P»Cř. Pj C...PsPrC- 3 PrPsCΓP»C-PrP¶PμPSCH CΓPiPμC‡PēC,,C-C‡PSPsC— PIP·P°C''PjPsPrC-C— P»C-PeP°CΉCΓCHPePēC... P·P°CΓPsP±C-PI P±P»PsPeP°C,PsCΉ α-P°PrCΉPμPSPsCΉPμC‡PμPiC,PsCΉC-PI PrPsPeCΓP°P·PsP·PēPS (4 PjPi C,P° 8 PjPi) C,P° CΓPēP»PrPμPSP°C,,C-P» (25 PjPi, 50 PjPi C,P° 100 PjPi) P·P°CΓC,PsCΓPsPICřPIP°P»Pē PsPrPSPsC‡P°CΓPSPs PiP°C‡C-C''PSC,P°Pj C-P· PrPsP±CΉPsCΠPeC-CΓPSPsCΉ PiC-PiPμCΉPiP»P°P·C-C''CΉ PiPμCΉPμPrPjC-C...CřCΉPsPIPsC— P·P°P»PsP·Pē, CΓC,P°P±C-P»C-P·P°C‡C-CΠ CΓC,P°PSCř CΠPePēC... P±CřP»P° PrPsCΓCΠPiPSCřC,P° ΠΐCΉPē P·P°CΓC,PsCΓCřPIP°PSPSC- PrPsPeCΓP°P·PsP·PēPSCř. Pj C,P°PePēC... PiP°C‡C-C''PSC,C-PI CΓPiPsCΓC,PμCΉC-PiP°P»PsCΓCΠ CΓPμCΉPμPrPSC'' PrPsPrP°C,PePsPIPμ P·PSPēP¶PμPSPSCΠ P°CΉC,PμCΉC-P°P»CHΉPSPsPiPs C,PēCΓPeCř Cř PiPsP»PsP¶PμPSPSC- P»PμP¶P°C‡Pē PSP° 7/7 PjPj CΉC,. CΓC,.. 9/5 PjPj CΉC,. CΓC,.. C,P° 8/4 PjPj CΉC,. CΓC,.. C,P° CΓPμCΉPμPrPSC'' P·PSPēP¶PμPSPSCΠ P°CΉC,PμCΉC-P°P»CHΉPSPsPiPs C,PēCΓPeCř Cř PiPsP»PsP¶PμPSPSC- CΓC,PsCΠC‡Pē PSP° 6/6 PjPj CΉC,. CΓC,.. 11/4 PjPj CΉC,. CΓC,.. 4/5 PjPj CΉC,. CΓC,.. PIC-PrPiPsPIC-PrPSPs. PμCΉPē PsPrPSPsC‡P°CΓPSPsPjCř P·P°CΓC,PsCΓCřPIP°PSPSC- CΓPēP»PrPμPSP°C,,C-P»Cř C,P° PrPsPeCΓP°P·PsP·PēPSCř Cř PiP°C‡C-C''PSC,C-PI, CΓC,P°P±C-P»C-P·P°C‡C-CΠ CΓC,P°PSCř CΠPePēC... P±CřP»P° PrPsCΓCΠPiPSCřC,P° ΠΐCΉPē P·P°CΓC,PsCΓCřPIP°PSPSC- PrPsPeCΓP°P·PsP·PēPSCř, C-PSPsPrC- PiPsPIC-PrPsPjP»CΠP»Pē ΠΐCΉPs CΉPsP·PIPēC,PsPe CΓPēPjPiC,PsPjP°C,PēC‡PSPsC— PsCΉC,PsCΓC,P°C,PēC‡PSPsC— PiC-PiPsC,PμPSP·C-C—. Pj C‡PēC... PiPsPIC-PrPsPjP»PμPSPSCΠC... PNēC€P»PsCΓCΠ ΠΐCΉPs PIPēPiP°PrPePē P·P°PiP°PjPsCΉPsC‡PμPSPSCΠ C,P° CΓC,P°PS PiPμCΉPμPr PSPμPiCΉPēC,PsPjPSC-CΓC,CΉ, P°P»Pμ P±PμP· CΓPēPSPePsPiPμ.

PκPμ CΓPiPsCΓC,PμCΉC-PiP°P»PsCΓCΠ P¶PsPrPSPēC... P·PSP°C‡CřC%PēC... PIP·P°C''PjPsPrC-PNē ΠΐCΉPē PsPrPSPsC‡P°CΓPSPsPjCř P·P°CΓC,PsCΓCřPIP°PSPSC- CΓPēP»PrPμPSP°C,,C-P»Cř (50 PjPi) C- C,PsP»P±CřC,P°PjC-PrCř (250 PjPi) P°P±Ps PIP°CΉC,,P°CΉPēPSCř (40 PjPi), C%Ps PjPμC,P°P±PsP»C-P·CřCΉC,CHCΓCΠ CYP2C9.

PŷPēP»PrPμPSP°C,,C-P» (50 PjPi) PSPμ ΠΐCΉPēP·PIPsPrPēPI PrPs PiPsPrPsPIP¶PμPSPSCΠ C‡P°CΓCř PeCΉPsPIPsC,PμC‡C-, CΓPiCΉPēC‡PēPSPμPSPsC— P·P°CΓC,PsCΓCřPIP°PSPSCΠPj P°C‡PμC,PēP»CΓP°P»C-C‡PēP»PsPIPsC— PePēCΓP»PsC,Pē (150 PjPi).

PŷPēP»PrPμPSP°C,,C-P» (50 PjPi) PSPμ PiPsC,PμPSC‡C-CΉPIP°PI PiC-PiPsC,PμPSP·PēPIPSCř PrC-CΉ P°P»PePsPiPsP»CΉ Cř P·PrPsCΉPsPIPēC... PrPsP±CΉPsPIPsP»CHC‡C-PI ΠΐCΉPē CΓPμCΉPμPrPSCΉPsPjCř PjP°PeCΓPēPjP°P»CHΉPSPsPjCř CΉC-PIPS- PμC,P°PSPsP»Cř PI PeCΉPsPIC- 80 PjPi/PrP».

Pj PiP°C‡C-C''PSC,C-PI, CΠPeC- P·P°CΓC,PsCΓPsPICřPIP°P»Pē CΓPēP»PrPμPSP°C,,C-P», PSPμ CΓPiPsCΓC,PμCΉC-PiP°P»PsCΓCΠ P¶PsPrPSPēC... PIC-PrPjC-PSPSPsCΓC,PμPNē ΠΐCΉPsC,,C-P»CΉ PiPsP±C-C‡PSPēC... PμC,,PμPeC,C-PI PiPsCΉC-PIPSCΠPSPs P· PiP»P°C‡PμP±Ps ΠΐCΉPē PsPrPSPsC‡P°CΓPSPsPjCř P·P°CΓC,PsCΓCřPIP°PSPSC- C,P°PePēC... PeP»P°CΓC-PI

PiC-PiPsC, P_μPSP·PēPIPSPēC... P»C-PeP°C₇C₇CHPePēC... P·P°C₇PSP±C-PI C₇Pe
 PrC-CrC₇P_μC, PēPePē, P±P»PsPeP°C, PsC₇Pē β-P°PrC₇P_μPSPsC₇P_μC†P_μPiC, PsC₇C-PI,
 C-PSPiC-P±C-C, PsC₇Pē P_hP_μP_α, P°PSC, P°PiPsPSC-C₇C, Pē P°PSPiC-PsC, P_μPSP·PēPSC₇ P†P†,
 P°PSC, PēPiC-PiP_μC₇C, P_μPSP·PēPIPSC- P»C-PeP°C₇C₇CHPeC- P·P°C₇PSP±Pē
 (C₇C₇PrPēPSPsC₇P_μC·CēPēC₇C₇PIP°P»CHPSC- C, P° C†P_μPSC, C₇P°P»CHPSPsC— PrC-C—),
 P±P»PsPeP°C, PsC₇Pē P°PrC₇P_μPSP_μC₇PiC-C†PSPēC... PSP_μPN₂C₇PSPSC-PI,
 P±P»PsPeP°C, PsC₇Pē PeP°P»CHC†C-C”PIPēC... PeP°PSP°P»C-PI C, P° P±P»PsPeP°C, PsC₇Pē α-
 P°PrC₇P_μPSPsC₇P_μC†P_μPiC, PsC₇C-PI. P_j C₇PiP_μC†C-P°P»CHPSPsP_jC₇
 PrPsC₇P»C-PrP_μPSPSC- PIP·P°C”P_jPsPrC-C— PiC₇Pē PsPrPSPsC†P°C₇PSPsP_jC₇
 P·P°C₇C, PsC₇C₇PIP°PSPSC- C₇PēP»PrP_μPSP°C,, C-P»C₇ (100 P_jPi) C, P° P°P_jP»PsPrPēPiC-PSC₇
 PiP°C†C-C”PSC, P°P_j P· P°C₇C, P_μC₇C-P°P»CHPSPsC₇ PiC-PiP_μC₇C, P_μPSP·C-C”C₇
 C₇PiPsC₇C, P_μC₇C-PiP°P»PsC₇C₇ PrPsPrP°C, PePsPIP_μ P·PSPēP_μPSPSC₇
 C₇PēC₇C, PsP»C-C†PSPsPiPs P°C₇C, P_μC₇C-P°P»CHPSPsPiPs C, PēC₇PeC₇ PI
 PiPsP»PsP_μPSPSC- P»P_μP_μP°C†Pē PSP° 8 P_jP_j C₇C,, C₇C,, P’C-PrPiPsPIC-PrPSP_μ
 P·PSPēP_μPSPSC₇ PrC-P°C₇C, PsP»C-C†PSPsPiPs P°C₇C, P_μC₇C-P°P»CHPSPsPiPs C, PēC₇PeC₇
 C₇C, P°PSPsPIPēP»Ps 7 P_jP_j C₇C,, C₇C,, P—P° PIP_μP»PēC†PēPSPsC₇ C†C- PrPsPrP°C, PePsPIC-
 P·PSPēP_μPSPSC₇ P°C₇C, P_μC₇C-P°P»CHPSPsPiPs C, PēC₇PeC₇ P±C₇P»Pē
 PiPsC₇C-PIPSC₇PSPSPēP_jPē C-P· C, PēP_jPē, C%Ps C₇PiPsC₇C, P_μC₇C-PiP°P»PēC₇C₇ PiC₇Pē
 P·P°C₇C, PsC₇C₇PIP°PSPSC- P»PēCēP_μ C₇PēP»PrP_μPSP°C,, C-P»C₇ C₇ P·PrPsC₇PSPiPēC...
 PrPsP±C₇PSPiPsP»CHC†C-PI (PrPēPI. C₇PSP·PrC-P» B«PαP°C₇P_jP°PePsP»PsPiC-C†PSC-
 PIP»P°C₇C, PēPIPsC₇C, C-B»).

PŸPēP»PrP_μPSP°C,, C-P» C₇ PrPsP·C- 100 P_jPi PSP_μ PIPiP»PēPIP°PI PSP°
 C,, P°C₇P_jP°PePsPeC-PSP_μC, PēC†PSC- PiPsPeP°P·PSPēPePē C-PSPiC-P±C-C, PsC₇C-PI
 P’P†P>-PiC₇P_μC, P_μP°P·Pē, C₇P°PePIC-PSP°PIC-C₇C₇ C, P° C₇PēC, PsPSP°PIC-C₇C₇, C₇PeC- C”
 C₇C₇P±C₇C, C₇P°C, P°P_jPē CYP3A4.

P_j P·PrPsC₇PSPiPēC... PrPsP±C₇PSPiPsP»CHC†C-PI C†PsP»PsPIC-C†PsC— C₇C, P°C, C-
 P·P°C₇C, PsC₇C₇PIP°PSPSC₇ C₇PēP»PrP_μPSP°C,, C-P»C₇ C₇ C₇C-PIPSPsPIP°P_μPSPsP_jC₇
 C₇C, P°PSC-

(80 P_jPi 3 C₇P°P·Pē PSP° PrPsP±C₇) PiC₇PēP·PIPsPrPēP»Ps PrPs PiC-PrPIPēC%P_μPSPSC₇ AUC
 C, P° C_{max} P±PsC₇P_μPSC, P°PSC₇ (125 P_jPi 2 C₇P°P·Pē PSP° PrPsP±C₇) PSP° 49,8 % 42 %
 PIC-PrPiPsPIC-PrPSPs.

P_hPrPSPsC†P°C₇PSP_μ P·P°C₇C, PsC₇C₇PIP°PSPSC₇ C₇PēP»PrP_μPSP°C,, C-P»C₇ P·
 P»C-PeP°C₇C₇CHPePēP_jPē P·P°C₇PSP±P°P_jPē, C%Ps P_jC-C₇C, C₇C, CH
 C₇P°PeC₇P±C-C, C₇PēP»/PIP°P»C₇P°C₇C, P°PS P±C₇P»Ps PiPsPIb₇™ C₇P·P°PSPs P·
 PiPsC₇PēP»P_μPSPSC₇P_j PiC-PiPsC, P_μPSP·C-C— C₇ PiP°C†C-C”PSC, C-PI, PiPsC₇C-PIPSC₇PSPs
 C-P· P·P°C₇C, PsC₇C₇PIP°PSPSC₇P_j P»PēCēP_μ
 C₇P°PeC₇P±C-C, C₇PēP»C₇/PIP°P»C₇P°C₇C, P°PSC₇. PŸPsP_jC₇ C₇P»C-Pr P·
 PsP±P_μC₇P_μP_μPSC-C₇C, C₇ C₇PSP·PiPsC†PēPSP°C, Pē P·P°C₇C, PsC₇C₇PIP°PSPSC₇
 C₇PēP»PrP_μPSP°C,, C-P»C₇ PiP°C†C-C”PSC, P°P_j, C₇PeC- PsC, C₇PēP_jC₇C, CH
 C₇P°PeC₇P±C-C, C₇PēP»/PIP°P»C₇P°C₇C, P°PS.

P_hC₇PSP±P»PēPIP_sC₇C, C- P·P°C₇C, PsC₇C₇PIP°PSPSC₇.

PŸP»C-Pr PiC₇PSPiP_μC₇C, Pē P_jP_μPrPēC†PSPēPN₂ P°PSP°P_jPSP_μP· C- C,, C-P·PēC†PSP_μ
 PsP±C₇C, P_μP_μP_μPSPSC₇ PrP»C₇ PrC-P°PiPSPsC₇C, PēPePē P_μC₇P_μPeC, PēP»CHPSPsC—
 PrPēC₇C,, C₇PSPeC†C-C— C- PIPēP·PSP°C†PēC, Pē PiPsC, P_μPSC†C-PN₂PSC-

РiРµCҗCЄPsPіCҗPёC±PёPSPё PrPs C,,P°CҗPjP°PePsP»PsPiC-C±PSPsPiPs P»C-PeCрPіP°PSPSCЦ.

PαP°PeC,PsCҗPё CҗPёP·PёPeCз CҗPµCҗC±PµPIPs-CҗCзPrPёPSPSPёC...

P·P°C...PIPsCҗCҗPіP°PSCҖ. PhCҗPeC-P»CHPePё CҗPµPeCҗCрP°P»CHPSP°

P°PeC,PёPIPSC-CҗC,CH CҗCрPіCҗPsPIPsPrP¶CҗC" C,CHCҗCЦ PіPµPIPSPёPj CҗPёP·PёPePsPj P·

P±PsPeCр CҗPµCҗC±CЦ, PrPs PіPsC±P°C,PeCр P±CрPrCH-CЦPePsPiPs P»C-PeCрPіP°PSPSCЦ

PµCҗPµPeC,PёP»CHPSPsC— PrPёCҗC,,CрPSPeCҗC-C— P»C-PeP°Cҗ PjP°C" PsCҗC-PSPёC,Pё

CҗC,P°PS CҗPµCҗC±PµPIPs-CҗCрPrPёPSPSPsC— CҗPёCҗC,PµPjPё PіP°CҗC-C"PSC,P°.

PҮPёP»PrPµPSP°C,,C-P» C±PёPSPёC,CH CҗCрPrPёPSPsCҗPsP·CёPёCҗCҗPіP°P»CHPSCр

PrC-CҖ, C°%Ps PіCҗPsCЦPіP»CЦC" C,CHCҗCЦ P»PµPiPePёPj C,P° PePsCҗPsC,PePsC±P°CҗPSPёPj

P·PSPёP¶PµPSPSCЦPj P°CҗC,PµCҗC-P°P»CHPSPsPiPs C,PёCҗPeCр (PrPёPі. CҗPsP·PrC-P»

B«PαP°CҗPjP°PePsPrPёPSP°PjC-PeP°B»). P"Ps PіCҗPёP·PSP°C±PµPSPSCЦ

CҗPёP»PrPµPSP°C,,C-P»Cр P»C-PeP°Cҗ PjP°C" CҗPµC,PµP»CHPSPs P·PіP°P¶PёC,Pё, C±Pё

PjPsP¶Pµ C,P°PePёPNё PµC,,PµPeC, PSPµCҗPіCҗPёCЦC,P»PёPIPs PіPіP»PёPіP°C,Pё PSP°

PіP°CҗC-C"PSC,C-PI C-P· PіPµPIPSPёPjPё PsCҗPSPsPIPSPёPjPё

P·P°C...PIPsCҗCҗPіP°PSPSCЦPjPё, PsCҗPsP±P»PёPIPs Cр PePsPjP±C-PSP°CҗC-C— C-P·

CҗPµPeCҗCрP°P»CHPSPsCҖ P°PeC,PёPIPSC-CҗC,CH. P"Ps PіP°CҗC-C"PSC,C-PI C-P·

PіC-PrPіPёC°%PµPSPsCҖ C±CҗC,P»PёPіC-CҗC,CH PrPs PіP°P·PsPrPёP»P°C,P°C,PsCҗC-PI

PSP°P»PµP¶P°C,CH PіP°CҗC-C"PSC,Pё P· PsP±CҗC,CҗCрPeCҗC-C"CH PіPёPіC-PrPSPsPiPs

C,CҗP°PeC,Cр P»C-PIPsPiPs CёP»CрPSPsC±PeP° (PSP°PіCҗPёPeP»P°Pр, CҗC,PµPSPsP·

P°PsCҗC,Pё, PіC-PіPµCҗC,CҗPsC,,C-C±PSP° PsP±CҗC,CҗCрPeC,PёPIPSP°

PeP°CҗPrC-PsPjC-PsPіP°C,C-CЦ) P°P±Ps PіP°CҗC-C"PSC,Pё P· CҗC-PrPeC-CҗPSPёPj

CҗPёPSPPrCҗPsPjPsPj PjCрP»CHC,PёCҗPёCҗC,PµPjPSPsC— P°C,CҗPsC,,C-C—, PsPrPSPёPj C-P·

PіCҗPsCЦPіC-PI CЦPePsC— C" C,CЦP¶PePµ PіPsCҗCрCёPµPSPSCЦ CҗPµPiCрP»CЦCҗC-C—

P°CҗC,PµCҗC-P°P»CHPSPsPiPs C,PёCҗPeCр P· P±PsPeCр PіPµPіPµC,P°C,PёPIPSPsC—

PSPµCҗPіPsPIPsC— CҗPёCҗC,PµPjPё.

PҮC-PSPµPіCҗP° PіPsC,PµPSCҗC-CҖC" PіC-PіPsC,PµPSP·PёPIPSPёPNё PµC,,PµPeC,

PSC-C,CҗP°C,C-PI (PrPёPі. CҗPsP·PrC-P» B«PµCҗPsC,PёPіPsPeP°P·P°PSPSCЦB»).

Pj PіPsCҗC,PjP°CҗPePµC,PёPSPiPsPIPsPjCр PіPµCҗC-PsPrC- PіPsPіC-PrPsPjP»CЦP»Pё PіCҗPs

C,CЦP¶PeC- PіPsP±C-C±PSC- CҗPµP°PeCҗC-C— P· P±PsPeCр CҗPµCҗC±PµPIPs-

CҗCрPrPёPSPSPsC— CҗPёCҗC,PµPjPё, PіPeP»CҖC±P°CҖC±Pё C-PSC,,P°CҗPeC,

PjC-PsPeP°CҗPrP°, PSPµCҗC,P°P±C-P»CHPSCр CҗC,PµPSPsPeP°CҗPrC-CҖ, CҗP°PіC,PsPіCр

CҗPµCҗC±PµPіCр CҗPjPµCҗC,CH, CёP»CрPSPsC±PePsPіCр P°CҗPёC,PjC-CҖ,

C±PµCҗPµP±CҗPsPіP°CҗPeCрP»CЦCҗPSC- PeCҗPsPIPsPіPёP»PёPіPё, C,CҗP°PSP·PёC,PsCҗPSCр

C-CёPµPjC-C±PSCр P°C,P°PeCр, P°CҗC,PµCҗC-P°P»CHPSCр PіC-PіPµCҗC,PµPSP·C-CҖ C,P°

P°CҗC,PµCҗC-P°P»CHPSCр PіC-PіPsC,PµPSP·C-CҖ, CЦPeC- P·P° C±P°CҗPsPj

P·P±C-PіP°P»PёCҗCЦ C-P· P·P°CҗC,PsCҗCрPіP°PSPSCЦPj PіCҗPµPіP°CҗP°C,Cр

PҮC-PSPµPіCҗP°. Pj P±C-P»CHCёPsCҗC,C- PіP°CҗC-C"PSC,C-PI, P°P»Pµ PSPµ Cр PіCҗC-C...,

C-CҗPSCрPіP°P»Pё C,,P°PeC,PsCҗPё CҗPёP·PёPeCр CҗPµCҗC±PµPIPs-CҗCрPrPёPSPSPёC...

P·P°C...PIPsCҗCҗPіP°PSCҖ. P'P°PіP°C,Ps C,P°PePёC... PіPsP±C-C±PSPёC... CҗPµP°PeCҗC-PNё

CҗPіPsCҗC,PµCҗC-PіP°P»PsCҗCЦ PіC-Pr C±P°Cҗ P°P±Ps PsPrCҗP°P·Cр PіC-CҗP»CЦ

CҗC,P°C,PµPIPsPiPs P°PeC,Cр C- P»PёCёPµ PeC-P»CHPeP° C,CҗP°PіPёP»PsCҗCЦ

PSPµPіPрPsPіP·C- PіC-CҗP»CЦ P·P°CҗC,PsCҗCрPіP°PSPSCЦ PіCҗPµPіP°CҗP°C,Cр

PҮC-PSPµPіCҗP° P±PµP· CҗPµPeCҗCрP°P»CHPSPsC— P°PeC,PёPIPSPsCҗC,C-. PҮPsPjCр

PSPµPjPsP¶P»PёPIPs PіPёP·PSP°C±PёC,Pё, C±Pё PіPsPіBҗ™ CЦP·P°PSPёPNё CҗPsP·PіPёC,PsPe

C,P°PePёC... PіPsP±C-C±PSPёC... CҗPµP°PeCҗC-PNё P±PµP·PіPsCҗPµCҗPµPrPSCҖPs P·

C,,P°PeC,PsCҗP°PjPё CҗPёP·PёPeCр, C±Pё C—C... CҗPsP·PіPёC,PsPe P·CрPjPsPіP»PµPSPёPNё

C-PSCёPёPjPё C±PёPSPSPёPeP°PjPё.

PµCҗC-P°PіC-P·Pj. P—P°CҗPsP±Pё PrP»CЦ P»C-PeCрPіP°PSPSCЦ PµCҗPµPeC,PёP»CHPSPsC—

PrPëCíC,,CíPSPeC†C-C—, Cí C,PsPjCí C†PëCíP»C- C- CíPëP»PrPµPSP°C,,C-P», CíP»C-Pr
PíCĤPëP·PSP°C†P°C,Pë P· PsP±PµCĤPµP¶PSC-CíC,CĤ PíP°C†C-C”PSC,P°Pj P·
P°PSP°C,PsPjC-C†PSPëPjPë PrPµC,,PsCĤPjP°C†C-CĬPjPë PíPµPSC-CíP° (C,P°PePëPjPë CĬPe
P°PSPiCíP»CĬC†C-CĬ, PeP°PIPµCĤPSPsP·PSPëPñ C,,C-P±CĤPSP· P°P±Ps C...PIPScĤPSP±P°
PµPµPñCĤPSPSC-) P°P±Ps PíP°C†C-C”PSC,P°Pj P·C- CíC,P°PSP°PjPë, C%Ps
CíPíCĤPëCĬCĤC,CĤ CĤPSP·PIPëC,PeCí PíCĤC-P°PíC-P·PjCí (C,P°PePëPjPë CĬPe
CíPµCĤPíPSPíPSPrC-P±PSPsPeP»C-C,PëPSPSP° P°PSPµPjC-CĬ, PjPSPsP¶PëPSPSP°
PjC-C”P»PsPjP° P°P±Ps P»PµPñPePµPjC-CĬ).

PµC-CíP»CĬ PIPëC...PsPrCí PíCĤPµPíP°CĤP°C,Cí PSP° CĤPëPSPsPe PíPSPIC-PrPsPjP»CĬP»Pë
PíCĤPSP PIPëPíP°PrPePë PíPSPrPsPIP¶PµPSPsC— PµCĤPµPeC†C-C— C,P° PíCĤC-P°PíC-P·PjCí.
PíPeC%Ps PµCĤPµPeC†C-CĬ C,CĤPëPIP°C” P±C-P»CĤCëPµ 4 PíPSPrPëPS, PíP°C†C-C”PSC,P°Pj
CíP»C-Pr PSpµPíP°PñPSPs P·PIPµCĤPSCíC,PëCíCĬ P·P° PjPµPrPëC†PSPsCĤ
PrPsPíPSPjPSPíPSCĤ. Pj CĤP°P·C- PIC-PrCíCíC,PSPsCíC,C- PSpµPíP°PñPSPsPíPSP
P»C-PeCíPIP°PSPSCĬ PíCĤC-P°PíC-P·Pj PjPSP¶Pµ PíCĤPëP·PIPµCíC,Pë PrPs
PíPSCëPePsPrP¶PµPSPSCĬ C,PeP°PSPëPS PíPµPSC-CíP° C,P° PrPs CíC,C-PñPePsC—
PIC,CĤP°C,Pë PíPSC,PµPSC†C-C—.

PhPrPSPsC†P°CíPSPµ P·P°CíC,PsCíCáPIP°PSPSCĬ P· C-PSCëPëPjPë C-PSPiC-P±C-C,PsCĤP°PjPë
PαP”P·5 P°P±Ps C-PSCëPëPjPë PíCĤPµPíP°CĤP°C,P°PjPë PrP»CĬ P»C-PeCáPIP°PSPSCĬ
PµCĤPµPeC,PëP»CĤPSPsC— PrPëCíC,,CáPSPeC†C-C—. P’PµP·PíPµPeCí C,P°
PµC,,PµPeC,PëPIPSC-CíC,CĤ PsPrPSPsC†P°CíPSPsPíPSP P·P°CíC,PsCíCíPIP°PSPSCĬ
CíPëP»PrPµPSP°C,,C-P»Cí P· C-PSCëPëPjPë C-PSPiC-P±C-C,PsCĤP°PjPë PαP”P·5 P°P±Ps
C-PSCëPëPjPë PíCĤPµPíP°CĤP°C,P°PjPë PrP»CĬ P»C-PeCíPIP°PSPSCĬ
PiC-PíPµCĤC,PµPSP·C-C— P»PµPíPµPSPµPIPSC— P°CĤC,PµCĤC-C—, CĬPeC- PjC-CíC,CĬC,CĤ
CíPëP»PrPµPSP°C,,C-P» (PSP°PíCĤPëPeP»P°Pr P PµPIP°C†PëPs), C†Pë P· C-PSCëPëPjPë
PíCĤPµPíP°CĤP°C,P°PjPë PrP»CĬ P»C-PeCíPIP°PSPSCĬ PµCĤPµPeC,PëP»CĤPSPsC—
PrPëCíC,,CíPSPeC†C-C— PSpµ PIPëPIC†P°P»Pë. PŷPsPjCí P·P°CíC,PsCíCíPIP°PSPSCĬ
C,P°PePëC... PePsPjP±C-PSP°C†C-Pñ PSpµ CĤPµPePsPjPµPSPrPsPIP°PSPµ.

P’PíP»PëPI PSP° P·C-CĤ. PŷPíPSPSC,P°PSPSC- PíPSPIC-PrPsPjP»PµPSPSCĬ PíCĤPSP
PIPëPSPëPePSPµPSPSCĬ PrPµC,,PµPeC,C-PI P·PsCĤCí PSP°PrC...PsPrPëP»Pë
P°CíPSC†C-PñPsPIP°PSPs C-P· P·P°CíC,PsCíCíPIP°PSPSCĬPj
CíPëP»PrPµPSP°C,,C-P»Cí C,P° C-PSCëPëC... C-PSPiC-P±C-C,PsCĤC-PI PαP”P·5
(PrPëPI. CĤPSP·PrC-P» B«PµPsP±C-C†PSC- CĤPµP°PeC†C-C—B»). PµCĤPSP
PIPëPíP°PrPePë PSpµP°CĤC,PµCĤC-P°P»CĤPSPsC— PíPµCĤPµPrPSCĤPsC—
C-CëPµPjC-C†PSPsC— PSpµPICĤPSPíP°C,C-C— P·PsCĤPSPsPíPSP PSpµCĤPIP°, C%Ps C”
CĤC-PrPeC-CíPSPëPj CíC,P°PSPsPj, PSP°PrC...PsPrPëP»Pë CíPíPSPSC,P°PSPSC-
PíPSPIC-PrPsPjP»PµPSPSCĬ C,P° PíPSPIC-PrPsPjP»CĬP»PsCíCĬ Cí
PSP°PíP»CĬPrPsPíPSPjCí PrPsCíP»C-PrP¶PµPSPSC- P°CíPSC†C-PñPsPIP°PSPs C-P·
P·P°CíC,PsCíCíPIP°PSPSCĬPj CíPëP»PrPµPSP°C,,C-P»Cí C,P° C-PSCëPëC...
C-PSPiC-P±C-C,PsCĤC-PI PαP”P·5 (PrPëPI. CĤPSP·PrC-P» B«PµPsP±C-C†PSC-
CĤPµP°PeC†C-C—B»).

PµP°C†C-C”PSC,C-PI CíP»C-Pr PíPSPíPµCĤPµPrPëC,Pë, C%Ps Cí
CĤP°P·C- CĤP°PíC,PsPíPSPíPSP PíPSCĤCíCëPµPSPSCĬ P·PsCĤCí
P·P°CíC,PsCíPSPiCíPIP°PSPSCĬ PíCĤPµPíP°CĤP°C,Cí PŷC-PSPµPíCĤP° CíP»C-Pr
PíCĤPëPíPëPSPëC,Pë C,P° PSpµPíP°PñPSPs P·PIPµCĤPSCíC,PëCíCĬ PrPs
P»C-PeP°CĤCĬ (PrPëPI. CĤPSP·PrC-P» B«PµCĤPSPC,PëPíPSPeP°P·P°PSPSCĬB»).

PhPrPSPsC†P°CíPSPµ P·P°CíC,PsCíCáPIP°PSPSCĬ P· CĤPëC,PsPSP°PIC-CĤPSPj.
PhPrPSPsC†P°CíPSPµ P·P°CíC,PsCíCíPIP°PSPSCĬ CíPëP»PrPµPSP°C,,C-P»Cí C,P°

P·P°CÍC,PsCÍCíPíP°PSPSCÍPj C-PSPiC-P±C-C,PsCÍC-PI PαP°P·5 C±Pë P· C-PSC€PëPjPë
C,,P°PeC,PsCÍP°PjPë PSPμPjPsP¶P»PëPIPs.

*PhPrPSPsC±P°CÍPSPμ P·P°CÍC,PsCÍCáPIP°PSPSCÍ C-P· PiC-PiPsC,PμPSP·PëPIPSPëPjPë
PiCÍPμPiP°CÍP°C,P°PjPë. PÝC-PSPμPiCÍP° C±PëPSPëC,CH CÍPëCÍC,PμPjPSCÍ
CÍCíPrPëPSPsCÍPsP·C€PëCÍPíPíP°P»CHPSCÍ PrC-CH C,P° PjPsP¶Pμ CÍ
PiPsPrP°P»CHC€PsPjCí P·PSPëP¶CíPíP°C,Pë P°CÍC,PμCÍC-P°P»CHPSPëPNë C,PëCÍPe Cí
PíP°C†C-C°PSC,C-PI, CÍPeC- P·P°CÍC,PsCÍPsPICíCÍC,CH PiC-PiPsC,PμPSP·PëPIPSC-
P»C-PeP°CÍCÍCHPeC- P·P°CÍPsP±Pë. P' PsPeCÍPμPjPsPjCí PrPsCÍP»C-PrP¶PμPSPSC-
P»C-PeP°CÍCÍCHPePsC— PIP·P°C°PjPsPrC-C— PsPrPSPsC±P°CÍPSPμ
P·P°CÍC,PsCÍCíPíP°PSPSCÍ P°PjP»PsPrPëPiC-PSCÍ (5 PjPi P°P±Ps 10 PjPi) C,P°
PiCÍPμPiP°CÍP°C,Cí PÝC-PSPμPiCÍP° (100 PjPi) PiPμCÍPμPsCÍP°P»CHPSPs
CÍPiPsCÍC,PμCÍC-PiP°P»PsCÍCÍ CÍPμCÍPμPrPSC° PrPsPrP°C,PePsPiPμ P·PSPëP¶PμPSPSCÍ
CÍPëCÍC,PsP»C-C±PSPsPiPs C,PëCÍPeCí PSP° 8 PjPj CÍC,.. CÍC,.. C,P°
PrC-P°CÍC,PsP»C-C±PSPsPiPs BÍ° PSP° 7 PjPj CÍC,.. CÍC,.*

*P—P°C...PIPsCÍCÍPíP°PSPSCÍ, C°Ps PiPμCÍPμPrP°CÍC,CHCÍCÍ CÍC,P°C,PμPIPëPj
C€P»CÍC...PsPj. P—P°CÍC,PsCÍCíPíP°PSPSCÍ PiCÍPμPiP°CÍP°C,Cí PÝC-PSPμPiCÍP°
PSPμ P·P°C...PëC°P°C° PIC-Pr P·P°C...PIPsCÍCÍPíP°PSCÍ, C°Ps
PiPμCÍPμPrP°CÍC,CHCÍCÍ CÍC,P°C,PμPIPëPj C€P»CÍC...PsPj. PÝP»C-Pr
CÍPsP·PiP»CÍPSCÍC,Pë PjPsP¶P»PëPIC-CÍC,CH C-PSC,,PsCÍPjCíPíP°PSPSCÍ
PíP°C†C-C°PSC,C-PI C°PsPrPs PSPμPsP±C...C-PrPSPëC... P·P°PiPsP±C-P¶PSPëC...
P·P°C...PsPrC-PI PrP»CÍ P·P°C...PëCÍC,Cí PIC-Pr P·P°C...PIPsCÍCÍPíP°PSCÍ, C°Ps
PiPμCÍPμPrP°CÍC,CHCÍCÍ CÍC,P°C,PμPIPëPj C€P»CÍC...PsPj, PiPeP»CÍC±P°CÍC±Pë
PIC-CÍCíCÍ C-PjCíPSPsPrPμC,,C-C†PëC,Cí P»CÍPrPëPSPë.*

*P—P°CÍC,PsCÍCáPIP°PSPSCÍ Cá PiPμCÍC-PsPr PIP°PiC-C,PSPsCÍC,C- P°P±Ps
PiPsPrCáPIP°PSPSCÍ PiCÍCáPrPrCÍ.*

PμCÍPμPiP°CÍP°C, PÝC-PSPμPiCÍP° PSPμ PiCÍPëP·PSP°C±PμPSPëPNë PrP»CÍ
P·P°CÍC,PsCÍCíPíP°PSPSCÍ P¶C-PSPeP°Pj.

*P—PrP°C,PSC-CÍC,CH PIPiP»PëPIP°C,Pë PSP° C€PIPëPrPeC-CÍC,CH CÍPμP°PeC†C-C—
PiCÍPë PePμCÍCáPIP°PSPSC- P°PIC,PsC,CÍP°PSCÍPiPsCÍC,PsPj P°P±Ps C-PSC€PëPjPë
PjPμC...P°PSC-P·PjP°PjPë.*

PÝC-PSPμPiCÍP° PjPsP¶Pμ PjP°C,Pë PSPμP·PSP°C±PSPëPNë PIPiP»PëPI PSP°
PePμCÍCíPíP°PSPSCÍ P°PIC,PsC,CÍP°PSCÍPiPsCÍC,PsPj P°P±Ps C-PSC€PëPjPë
PjPμC...P°PSC-P·PjP°PjPë. PhCÍPeC-P»CHPePë Cí C...PsPrC- PrPsCÍP»C-PrP¶PμPSCÍ
P·P°CÍC,PsCÍCíPíP°PSPSCÍ CÍPëP»PrPμPSP°C,,C-P»Cí PiPsPIC-PrPsPjP»CÍP»Pë PiCÍPs
PIPëPiP°PrPePë P·P°PiP°PjPsCÍPsC±PμPSPSCÍ C,P° PiPsCÍCíC€PμPSPSCÍ P· P±PsPeCí
PsCÍPiP°PSC-PI P·PsCÍCí, PiPμCÍPμPr C,PëPj CÍPe CÍC-PrP°C,Pë P·P° PePμCÍPjPs
C,CÍP°PSCÍPiPsCÍC,PSPsPiPs P·P°CÍPsP±Cí P°P±Ps PiCÍP°C†CÍPíP°C,Pë P·
PjPμC...P°PSC-P·PjP°PjPë, PiP°C†C-C°PSC,P°Pj PSPμPsP±C...C-PrPSPs P·BÍ™ CÍCÍCíPíP°C,Pë,
CÍPePsCÍ C° C—C...PSCÍ C-PSPrPëPIC-PrCíP°P»CHPSP° CÍPμP°PeC†C-CÍ PSP°
P·P°CÍC,PsCÍCíPíP°PSPSCÍ PiCÍPμPiP°CÍP°C,Cí PÝC-PSPμPiCÍP°.

PŸPİPsCİC-P± P·P°CİC,PsCİCžPIP°PSPSCİ C,P° PrPsP·Pë.

PıCĤPıPİP°CĤP°C, P·P°CİC,PsCİCıPICıPIP°C,Pë PİPıCĤPsCĤP°P»CHBPSPs.

P''PsCĤPsCİP»C-. P PıPePsPjPıPSPPrPsPIP°PSP° PrPsP·P° PİCĤPıPİP°CĤP°C,Cı PŸC-PSPıPİCĤP°CİC,P°PSPsPIPëC,CH 50 PjPi, P·P°CİC,PsCİCıPICıPIP°C,Pë Cı CĤP°P·C-PSPıPsP±C...C-PrPSPsCİC,C- PİCĤPëP±P»PëP·PSPs P·P° PİPsPrPëPSCı PrPs CİPıPeCİCıP°P»CHBPSPsC— P°PeC,PëPIPSPsCİC,C-. P—P°P»PıPıPSPs PIC-Pr PıC,,PıPeC,PëPIPSPsCİC,C- C,P° PİPıCĤPıPSPsCİPëPjPsCİC,C- PİCĤPıPİP°CĤP°C,Cı PrPsP·Cı PjPsPıPSP° P·P±C-P»CHCëPëC,Pë PrPs 100 PjPi P°P±Ps P·PjPıPSCëPëC,Pë PrPs 25 PjPi. PıP°PeCİPëPjP°P»CHBPSP° CĤPıPePsPjPıPSPPrPsPIP°PSP° PrPsP·P° CİC,P°PSPsPIPëC,CH 100 PjPi. PŸP°CİC,PsC,P° P·P°CİC,PsCİCıPIP°PSPSCİ PjP°PeCİPëPjP°P»CHBPSPsC— CĤPıPePsPjPıPSPPrPsPIP°PSPsC— PrPsP·Pë PİCĤPıPİP°CĤP°C,Cı CİC,P°PSPsPIPëC,CH 1 CĤP°P·PSP° PrPsP±Cı. PıCĤPë P·P°CİC,PsCİCıPIP°PSPSC- PİCĤPıPİP°CĤP°C,Cı PŸC-PSPıPİCĤP°PİC-Pr CıP°Cİ PİCĤPëPNëPsPjCı C—PıC- PrC-Cİ PİCĤPıPİP°CĤP°C,Cı PjPsPıPıPSP°CİC,P°PIP°C,Pë PİC-P·PSC-CëPı, PSC-Pı PİCĤPë PNëPsPİPs P·P°CİC,PsCİCıPIP°PSPSC-PSP°C,C°ıPı.

PıP°CıC-C''PSC,Pë P»C-C,PSCHPsPiPs PIC-PeCž. PıPıPsP±C...C-PrPSC-CİC,CH PePsCĤPıPeCıC— PrPsP·Pë PrP»Cİ PİP°CıC-C''PSC,C-PI P»C-C,PSCHPsPiPs PIC-PeCı (≥ 65 CĤPsPeC-PI) PIC-PrCİCıC,PSCİ.

PıP°CıC-C''PSC,Pë P·PSPëCĤPePsPIPsCĤ PSPıPrPsCİC,P°C,PSC-CİC,CĤ. PıP°CıC-C''PSC,P°Pj C-P·PSPëCĤPePsPIPsCĤ PSPıPrPsCİC,P°C,PSC-CİC,CĤ P»PıPİPePsPiPs C,P° PİPsPjC-CĤPSPsPiPs CİC,CıPİPıPSCİ (PeP»C-CĤPıPSCı PeCĤPıP°C,PëPSC-PSCı 30-80 PjP»/C...PI) CĤPıPePsPjPıPSPPrPsPIP°PSP° PrPsP·P° PİCĤPıPİP°CĤP°C,Cı C'' C,P°PePsCĤ Pı, CİPe PSP°PIPıPrPıPSPs PİPëC°ıPı Cı CĤPsP·PrC-P»C- B«P''PsCĤPsCİP»C-B».

PıCİPeC-P»CHPePë Cı PİP°CıC-C''PSC,C-PI C-P·PSPëCĤPePsPIPsCĤ PSPıPrPsCİC,P°C,PSC-CİC,CĤ C,CİPıPePsPiPs CİC,CıPİPıPSCİ (PeP»C-CĤPıPSCı PeCĤPıP°C,PëPSC-PSCı < 30 PjP»/C...PI) PeP»C-CĤPıPSCı CİPëP»PrPıPSP°C,,C-P»Cı P·PSPëPıPıPSPëPNë, CİP»C-Pr CĤPsP·PİP»CİPSCıC,Pë PjPsPıP»PëPIC-CİC,CH P·P°CİC,PsCİCıPIP°PSPSCİ PrPsP·Pë 25 PjPi. P—P°P»PıPıPSPs PIC-Pr PıC,,PıPeC,PëPIPSPsCİC,C- C,P° PİPıCĤPıPSPsCİPëPjPsCİC,C- PİCĤPıPİP°CĤP°C,Cı PİCĤPë PSPıPsP±C...C-PrPSPsCİC,C- PrPsP·Cı PjPsPıPSP° P·P±C-P»CHCëPëC,Pë PİPsCİC,CıPİPsPIPs PrPs 50 PjPi C,P° PrPs 100 PjPi.

PıP°CıC-C''PSC,Pë C-P·PİPıCıC-PSPePsPIPsCĤ PSPıPrPsCİC,P°C,PSC-CİC,CĤ. PıCİPeC-P»CHPePë Cı PİP°CıC-C''PSC,C-PI C-P·PİPıCıC-PSPePsPIPsCĤ PSPıPrPsCİC,P°C,PSC-CİC,CĤ (PSP°PİCĤPëPeP»P°Pr CıPëCĤPsP·PsPj) PeP»C-CĤPıPSCı CİPëP»PrPıPSP°C,,C-P»Cı P·PSPëPıPıPSPëPNë, CİP»C-Pr CĤPsP·PİP»CİPSCıC,Pë PjPsPıP»PëPIC-CİC,CH P·P°CİC,PsCİCıPIP°PSPSCİ PrPsP·Pë 25 PjPi. P—P°P»PıPıPSPs PIC-Pr PıC,,PıPeC,PëPIPSPsCİC,C- C,P° PİPıCĤPıPSPsCİPëPjPsCİC,C- PİCĤPıPİP°CĤP°C,Cı PİCĤPë PSPıPsP±C...C-PrPSPsCİC,C- PrPsP·Cı PjPsPıPSP° P·P±C-P»CHCëPëC,Pë PİPsCİC,CıPİPsPIPs PrPs 50 PjPi C,P° PrPs 100 PjPi.

PıP°CıC-C''PSC,Pë, CİPeC- P·P°CİC,PsCİCıPICžCĤC,CH C-PSCëC- P»C-PeP°CĤCİCĤPeC- P·P°CİPsP±Pë. PİPeC°ıPs PİP°CıC-C''PSC,Pë PsPrPSPsCıP°CİPSPs P·P°CİC,PsCİCıPICıCĤC,CH

C-PSPIc-P±C-C,PsCḡPë CYP3A4 (PrPëPI. CḡPsP·PrC-P» B«P'P·P°C"PjPsPrC-Cḡ P· C-PSC€PëPjPë P»C-PeP°CḡCḡCHPePëPjPë P·P°CḡPsP±P°PjPë C,P° C-PSC€C- PIPëPrPë PIP·P°C"PjPsPrC-PNëB»), CḡP»C-Pr CḡPsP·PiP»CḡPSCrC,Pë PjPsPḡP»PëPIC-CḡC,CH P·P°CḡC,PsCḡCḡPIP°PSPSCḡ PİPsC±P°C,PePsPIPsC— PrPsP·Pë 25 PjPi (P·P° PIPëPSCḡC,PePsPj CḡPëC,PsPSP°PIC-CḡCḡ, P·P°CḡC,PsCḡCḡPIP°PSPSCḡ CḡPePsPiPs PsPrPSPsC±P°CḡPSPs C-P· CḡPëP»PrPµPSP°C,,C-P»PsPj PSPµ CḡPµPePsPjPµPSPrCḡC"C,CHCḡCḡ, PrPëPI. CḡPsP·PrC-P» B«PhCḡPsP±P»PëPIPsCḡC,C- P·P°CḡC,PsCḡCḡPIP°PSPSCḡB»).

P— PjPµC,PsCḡ PjC-PSC-PjC-P·P°CḡC-C— PjPsPḡP»PëPIPsPiPs CḡPsP·PIPëC,PeCḡ PİPsCḡC,CḡCḡP°P»CHPSPsC— PiC-PİPsC,PµPSP·C-C— Cḡ PİP°CḡC-C"PSC,C-PI, CḡPeC- P·P°CḡC,PsCḡPsPICḡCḡC,CH P±P»PsPeP°C,PsCḡPë α-P°PrCḡPµPSPsCḡPµCḡPµPİC,PsCḡC-PI, C—C... CḡC,P°PS PİPsC,CḡC-P±PSPs CḡC,P°P±C-P»C-P·CḡPIP°C,Pë P·P° PrPsPİPsPjPsPiPsCḡ P±P»PsPeP°C,PsCḡC-PI α-P°PrCḡPµPSPsCḡPµCḡPµPİC,PsCḡC-PI PrPs PİPsC±P°C,PeCḡ P·P°CḡC,PsCḡPsPICḡPIP°PSPSCḡ CḡPëP»PrPµPSP°C,,C-P»Cḡ. PḡP°PePsPḡ CḡP»C-Pr CḡPsP·PiP»CḡPSCrC,Pë PjPsPḡP»PëPIC-CḡC,CH P·P°CḡC,PsCḡCḡPIP°PSPSCḡ PİPsC±P°C,PePsPIPsC— PrPsP·Pë 25 PjPi (PrPëPI. CḡPsP·PrC-P»Pë B«PhCḡPsP±P»PëPIPsCḡC,C- P·P°CḡC,PsCḡCḡPIP°PSPSCḡB» C- B«P'P·P°C"PjPsPrC-Cḡ P· C-PSC€PëPjPë P»C-PeP°CḡCḡCHPePëPjPë P·P°CḡPsP±P°PjPë C,P° C-PSC€C- PIPëPrPë PIP·P°C"PjPsPrC-PNëB»).

P"C-C,Pë.

PµCḡPµPİP°CḡP°C, PSPµ PİPsPeP°P·P°PSPëPNë PrPs P·P°CḡC,PsCḡCḡPIP°PSPSCḡ PsCḡPsP±P°Pj PIC-PePsPj PrPs 18 CḡPsPeC-PI.

PµPµCḡPµPrPsP·CḡPIP°PSPSCḡ.

Pj C...PsPrC- PrPsCḡP»C-PrPḡPµPSCḡ P· CḡC±P°CḡC,Cḡ PrPsP±CḡPsPIPsP»CHCḡC-PI PİC-Pr C±P°Cḡ P·P°CḡC,PsCḡCḡPIP°PSPSCḡ CḡP°P·PsPIPsC— PrPsP·Pë CḡPëP»PrPµPSP°C,,C-P»Cḡ PrPs 800 PjPi PİPsP±C-C±PSC- CḡPµP°PeCḡC-C— P±CḡP»Pë PİPsPrC-P±PSPëPjPë PrPs C,PëC..., C%oPs CḡPİPsCḡC,PµCḡC-PiP°P»PëCḡCḡ PİCḡPë P·P°CḡC,PsCḡCḡPIP°PSPSC- CḡPëP»PrPµPSP°C,,C-P»Cḡ Cḡ PSPëPḡC±PëC... PrPsP·P°C..., P°P»Pµ PIPëPSPëPeP°P»Pë C±P°CḡC,C-C€Pµ C,P° P±CḡP»Pë P±C-P»CHC€ C,CḡPḡPePëPjPë. P—P°CḡC,PsCḡPsPICḡPIP°PSPSCḡ CḡPëP»PrPµPSP°C,,C-P»Cḡ Cḡ PrPsP·C- 200 PjPi PSPµ PİCḡPëP·PIPsPrPëP»Ps PrPs PİC-PrPIPëC%oPµPSPSCḡ PµC,,PµPeC,PëPIPsCḡC,C-, P°P»Pµ CḡPİCḡPëC±PëPSCḡP»Ps P·CḡPsCḡC,P°PSPSCḡ PeC-P»CHPePsCḡC,C- PIPëPİP°PrPeC-PI CḡPsP·PIPëC,PeCḡ PİPsP±C-C±PSPëC... CḡPµP°PeCḡC-PNë (PİPsP»PsPIPsPiPs P±PsP»Cḡ, PİCḡPëPİP»PëPIC-PI, P·P°PİP°PjPsCḡPsC±PµPSPSCḡ, PrPëCḡPİPµPİCḡC-C—, P·P°PeP»P°PrPµPSPsCḡC,C- PSPsCḡP°, PİPsCḡCḡC€PµPSCḡ P· P±PsPeCḡ PsCḡPİP°PSC-PI P·PsCḡCḡ).

Pj CḡP°P·C- PİPµCḡPµPrPsP·CḡPIP°PSPSCḡ PİCḡPë PSPµPsP±C...C-PrPSPsCḡC,C- CḡP»C-Pr PIPrP°C,PëCḡCḡ PrPs P·PIPëC±P°PNëPSPëC... PİC-PrC,CḡPëPjCḡCḡC±PëC... P·P°C...PsPrC-PI. PµCḡPëCḡPePsCḡPµPSPSCḡ PeP»C-CḡPµPSCḡCḡ CḡPëP»PrPµPSP°C,,C-P»Cḡ PİCḡPë PİPµPjPsPrC-P°P»C-P·C- PjP°P»PsPNëPjPsPIC-CḡPSPµ PIPSP°CḡP»C-PrPsPe PIPëCḡPsPePsPiPs CḡC,CḡPİPµPSCḡ P·PIBḡ™ CḡP·CḡPIP°PSPSCḡ PİCḡPµPİP°CḡP°C,Cḡ P· P±C-P»PeP°PjPë PİP»P°P·PjPë PeCḡPsPIC- C,P° PIC-PrCḡCḡC,PSPsCḡC,C- PµP»C-PjC-PSP°CḡC-C— CḡPëP»PrPµPSP°C,,C-P»Cḡ C-P· CḡPµC±PµCḡ.

PμPsP±C-C±PSC- CḡPμP°PeC†C-C—.

PK°PNᵇC†P°C†C,C-C€Pμ PīPsPIC-PrPsPjP»CḡP»Pē PīCḡPs C,P°PeC- PīPsP±C-C±PSC-
CḡPμP°PeC†C-C— CḡPe PīPsP»PsPIPSpēPNᵇ P±C-P»CH, PīCḡPēPīP»PēPIPē,
PrPēC†PīPμPīC†C-Cḡ, P·P°PeP»P°PrPμPSC-C†C,CH PSPsC†P°, P±C-P»CH Cr C†PīPēPSC-,
P·P°PīP°PjPsCḡPsC†PμPSPSCḡ, PSC†PrPsC,C†, PīPsCḡC†C€PμPSPSCḡ P·PsCḡC†,
C†C-P°PSPsPīC†C-Cḡ C,P° P·P°C,C†PjP°PSPμPSPSCḡ P·PsCḡC†.

PμPsP±C-C±PSC- CḡPμP°PeC†C-C—, C%ᵇPs C†PīPsC†C,PμCḡC-PiP°P»PēC†Cḡ PīC-Pr C†P°C†
PrPsC†P»C-PrPḡPμPSCḡ, PSP°PIPμPrPμPSPs PSPēPḡC†Pμ PIC-PrPīPsPIC-PrPSPs PrPs
PeP»P°C†PēC,,C-PeP°C†C-C— C,P° C†P°C†C,PsC,Pē: PrC†PḡPμ C†P°C†C,Ps (≥ 1/10), C†P°C†C,Ps
(≥ 1/100-< 1/10), PSPμC†P°C†C,Ps (≥ 1000-< 1/100) C,P° CḡC-PrPePs (≥ 1/10000-< 1/1000).

P†PSC,,PμPeC†C-C— C,P° C-PSPIP°P·C-C—: PSPμC†P°C†C,Ps Bḡ“ CḡPēPSC-C,,.

*P— P±PsPeCž C-PjCžPSPSPsC— C†PēC†C,PμPjPē: PSPμC†P°C†C,Ps Bḡ“
PiC-PīPμCḡC†C†C,P»PēPIC-C†C,CH.*

*P— P±PsPeCž PSPμCḡPIPsPIPsC— C†PēC†C,PμPjPē: PrC†PḡPμ C†P°C†C,Ps Bḡ“
PīPsP»PsPIPSpēPNᵇ P±C-P»CH; C†P°C†C,Ps Bḡ“ P·P°PīP°PjPsCḡPsC†PμPSPSCḡ;
PSPμC†P°C†C,Ps Bḡ“ C†PsPSP»PēPIC-C†C,CH, PiC-PīPμC†C,PμP·C-Cḡ; CḡC-PrPePs Bḡ“
C-PSC†C†P»CHC,, C,CḡP°PSP·PēC,PsCḡPSP° C-C€PμPjC-C†PSP° P°C,P°PeP°,
C†C†PrPsPjPē*, CḡPμC†PēPrPēPIPē C†C†PrPsPj*, C†PēPSPePsPīPμ.*

*P— P±PsPeCž PsCḡPiP°PSC-PI P·PsCḡCž: C†P°C†C,Ps Bḡ“ PīPsCḡC†C€PμPSPSCḡ
C†PīCḡPēPNᵇPSCḡC,C,Cḡ PePsP»CHPsCḡC†**, CḡPsP·P»P°PrPē P·PsCḡC†,
P·P°C,CHPjP°CḡPμPSPSCḡ P·PsCḡC†; PSPμC†P°C†C,Ps Bḡ“ CḡPsP·P»P°PrPē
C†P»CHPsP·PsPIPēPrC-P»PμPSPSCḡ***, P±C-P»CH PI PsC†P°C...,
C,,PsC,PsC,,PsP±C-Cḡ, C,,PsC,PsPīC†C-Cḡ, PiC-PīPμCḡPμPjC-Cḡ PsC†PμPNᵇ,
CḡC†PeCḡP°PIC-C†C,CH P·PsCḡC†, PePsPSBḡ™ CHPSPeC,PēPIC-C,,; CḡC-PrPePs Bḡ“
PSPμP°CḡC,PμCḡC-P°P»CHPSP° PīPμCḡPμPrPSCḡ C-C€PμPjC-C†PSP°
PSPμPNᵇCḡPsPīP°C,C-Cḡ P·PsCḡPsPIPsPīPs PSPμCḡPIP°*, PsPeP»CHP·C-Cḡ C†C†PrPēPS
C†C-C,PeC-PIPePē*, CḡPμC,PēPSP°P»CHPSPēPNᵇ PeCḡPsPIPsPIPēP»PēPI,
P°CḡC,PμCḡC-PsC†PeP»PμCḡPsC,PēC†PSP° CḡPμC,PēPSPsPīP°C,C-Cḡ,
PīPsCḡC†C€PμPSPSCḡ P· P±PsPeC† C†C-C,PeC-PIPePē, PiP»P°C†PePsPjP°,
PrPμC,,PμPeC,Pē PīPsP»Cḡ P·PsCḡC†, PrPēPīP»PsPīC-Cḡ, P·PSPēPḡPμPSPSCḡ
PīPsC†C,CḡPsC,Pē P·PsCḡC†, PjC-PsPīC-Cḡ, P°C†C,PμPSPsPīC-Cḡ, PīP»P°PIP°CḡC†C-
PīPsPjC†C,PSC-PSPSCḡ C†PeP»PēC†C,PsPīPs C,C-P»P°, PīPsCḡC†C€PμPSPSCḡ P·
P±PsPeC† CḡP°PNᵇPrC†PḡPSPsC— PsP±PsP»PsPSPePē, PjC-PrCḡC-P°P·, PīPsCḡPIP°
C†CḡCḡC†PēC... PeCḡC†PiC-PI PSP°PIPePsP»Ps PrPḡPμCḡPμP»P° C†PIC-C,P»P°
(PiP°P»Ps) Cr PīPsP»C- P·PsCḡC†, PSP°P±CḡCḡPe PsC†PμPNᵇ,
PīCḡPēPīC†C...P»C-C†C,CH PsC†PμPNᵇ, PīPsCḡC†C€PμPSPSCḡ P· P±PsPeC† PsC†PμPNᵇ,
PiC-PīPμCḡPμPjC-Cḡ PePsPSBḡ™ CHPSPeC,PēPIPē, PīPsPrCḡP°P·PSPμPSPSCḡ
PsC†PμPNᵇ, P°PSPsPjP°P»CHPSC- PIC-PrC†C†C,C,Cḡ PI PsC†P°C..., PSP°P±CḡCḡPe
PīPsPIC-Pe, P·PSPμP±P°CḡPIP»PμPSPSCḡ C†PeP»PμCḡPē.*

P— P±PsPeCž PsCḡPiP°PSC-PI C†P»CžC...Cž: PSPμC†P°C†C,Ps Bḡ“

$P \cdot P^\circ \ddot{P} P^\circ P_j P_s C_{\overline{H}} P_s C_{\overline{H}} P_{\mu} P S P S C_{\overline{H}}$, $P r P \cdot P I C - P S$ $C r$ $P I C r C \dots P^\circ C \dots$; $C_{\overline{H}} C - P r P e P_s$ $\overline{H}$ “
 $P i P \gg C r C \dots P_s C, P^\circ$.

$P - P_{\pm} P_s P e C \acute{z} C \acute{I} P_{\mu} C_{\overline{H}} C_{\overline{H}} P_{\mu} P I P_s - C \acute{I} C \acute{z} P r P \ddot{e} P S P S P_s C - C \acute{I} P \ddot{e} C \acute{I} C, P_{\mu} P_j P \ddot{e}$: $C_{\overline{H}} P^\circ C \acute{I} C, P_s$ $\overline{H}$ “
 $P i C_{\overline{H}} P \ddot{e} P i P \gg P \ddot{e} P i P \ddot{e}$; $P S P_{\mu} C_{\overline{H}} P^\circ C \acute{I} C, P_s$ $\overline{H}$ “ $C, P^\circ C \dots C - P e P^\circ C_{\overline{H}} P r C - C_{\overline{H}}$, $P i P_s C \acute{I} P \ddot{e} P \gg P_{\mu} P S P_{\mu}$
 $C \acute{I} P_{\mu} C_{\overline{H}} C_{\overline{H}} P_{\mu} P_{\pm} P \ddot{e} C, C, C_{\overline{H}}$, $P^\circ C_{\overline{H}} C, P_{\mu} C_{\overline{H}} C - P^\circ P \gg C_{\overline{H}} P S P^\circ$
 $P i C - P i P_{\mu} C_{\overline{H}} C, P_{\mu} P S P \cdot C - C_{\overline{H}} / P i C - P i P_s C, P_{\mu} P S P \cdot C - C_{\overline{H}}$; $C_{\overline{H}} C - P r P e P_s$ $\overline{H}$ “ $C_{\overline{H}} P^\circ P i C, P_s P i P^\circ$
 $C \acute{I} P_{\mu} C_{\overline{H}} C_{\overline{H}} P_{\mu} P i P^\circ$ $C \acute{I} P_j P_{\mu} C_{\overline{H}} C, C_{\overline{H}}^*$, $C - P S C, P^\circ C_{\overline{H}} P e C$, $P_j C - P_s P e P^\circ C_{\overline{H}} P r P^\circ$, $C \epsilon P \gg C r P S P_s C_{\overline{H}} P e P_s P i P^\circ$
 $P^\circ C_{\overline{H}} P \ddot{e} C, P_j C - C_{\overline{H}}^*$, $C, C - P_{\pm} C_{\overline{H}} P \ddot{e} P \gg C_{\overline{H}} C_{\overline{H}} C - C_{\overline{H}}$ $P i P_{\mu} C_{\overline{H}} P_{\mu} P r C \acute{I} P_{\mu} C_{\overline{H}} P r C_{\overline{H}}$,
 $P S P_{\mu} C \acute{I} C, P^\circ P_{\pm} C - P \gg C_{\overline{H}} P S P^\circ$ $C \acute{I} C, P_{\mu} P S P_s P e P^\circ C_{\overline{H}} P r C - C_{\overline{H}}$.

$P - P_{\pm} P_s P e C \acute{z} P r P \ddot{e} C \dots P^\circ P \gg C_{\overline{H}} P S P_s C - C \acute{I} P \ddot{e} C \acute{I} C, P_{\mu} P_j P \ddot{e}$: $C_{\overline{H}} P^\circ C \acute{I} C, P_s$ $\overline{H}$ “
 $P \cdot P^\circ P e P \gg P^\circ P r P_{\mu} P S C - C \acute{I} C, C_{\overline{H}}$ $P S P_s C \acute{I} P^\circ$; $P S P_{\mu} C_{\overline{H}} P^\circ C \acute{I} C, P_s$ $\overline{H}$ “ $P S P_s C \acute{I} P_s P i P^\circ$
 $P e C_{\overline{H}} P_s P i P_s C, P_{\mu} C_{\overline{H}} P^\circ$, $P \cdot P^\circ P e P \gg P^\circ P r P_{\mu} P S C - C \acute{I} C, C_{\overline{H}}$ $P i C_{\overline{H}} P \ddot{e} P r P^\circ C, P e P_s P i P \ddot{e} C \dots$
 $P i P^\circ P \cdot C r C \dots P S P_s C \acute{I} P^\circ$; $C_{\overline{H}} C - P r P e P_s$ $\overline{H}$ “ $P I C - P r C_{\overline{H}} C r C, C, C_{\overline{H}}$ $C \acute{I} C, P \ddot{e} C \acute{I} P S P_{\mu} P S P S C_{\overline{H}}$ $C r$
 $P i P_s C_{\overline{H}} P \gg C -$, $P S P^\circ P_{\pm} C_{\overline{H}} C_{\overline{H}} P e$ $C \acute{I} P \gg P \ddot{e} P \cdot P_s P i P_s C - P_s P_{\pm} P_s P \gg P_s P S P e P \ddot{e}$ $P S P_s C \acute{I} P^\circ$,
 $C \acute{I} C r C \dots C - C \acute{I} C, C_{\overline{H}}$ $C r$ $P S P_s C \acute{I} C -$.

$P - P_{\pm} P_s P e C \acute{z} C, C_{\overline{H}} P^\circ P i P S P_s P i P_s C, C_{\overline{H}} P^\circ P e C, C \acute{z}$: $C_{\overline{H}} P^\circ C \acute{I} C, P_s$ $\overline{H}$ “ $P S C r P r P_s C, P^\circ$,
 $P r P \ddot{e} C \acute{I} P i P_{\mu} P i C \acute{I} C - C_{\overline{H}}$; $P S P_{\mu} C_{\overline{H}} P^\circ C \acute{I} C, P_s$ $\overline{H}$ “ $P i P^\circ C \acute{I} C, C_{\overline{H}} P_s P_{\mu} P \cdot P_s C, P^\circ P i P_{\mu} P^\circ P \gg C_{\overline{H}} P S P^\circ$
 $C_{\overline{H}} P_{\mu} C, P \gg C_{\overline{H}} P e C \acute{I} P S P^\circ$ $C \dots P i P_s C_{\overline{H}} P_s P_{\pm} P^\circ$, $P_{\pm} P \gg C_{\overline{H}} P i P^\circ P S P S C_{\overline{H}}$, $P_{\pm} C - P \gg C_{\overline{H}}$ $C r$
 $P i P_{\mu} C_{\overline{H}} C \dots P S C - P N_e$ $C_{\overline{H}} P^\circ C \acute{I} C, P \ddot{e} P S C - P \P P \ddot{e} P i P_s C, P^\circ$, $C \acute{I} C r C \dots C - C \acute{I} C, C_{\overline{H}}$ $C r$ $C_{\overline{H}} P_s C, C -$;
 $C_{\overline{H}} C - P r P e P_s$ $\overline{H}$ “ $P i C - P i P_{\mu} C \acute{I} C, P_{\mu} P \cdot C - C_{\overline{H}}$ $C_{\overline{H}} P_s C, P_s P i P_s C - P i P_s C_{\overline{H}} P_s P \P P S P \ddot{e} P S P \ddot{e}$.

$P - P_{\pm} P_s P e C \acute{z} C \epsilon P e C - C_{\overline{H}} P \ddot{e} C, P^\circ P i C - P r C \epsilon P e C - C_{\overline{H}} P S P_s C - C, P e P^\circ P S P \ddot{e} P S P \ddot{e}$: $P S P_{\mu} C_{\overline{H}} P^\circ C \acute{I} C, P_s$ $\overline{H}$ “
 $P i P \ddot{e} C \acute{I} P \ddot{e} P i P^\circ P S P S C_{\overline{H}}$; $C_{\overline{H}} C - P r P e P_s$ $\overline{H}$ “ $C \acute{I} P \ddot{e} P S P r C_{\overline{H}} P_s P_j P \ddot{Y} C, C - P i P_{\mu} P S C \acute{I} P^\circ -$
 $P^\circ P \P P_s P S C \acute{I} P_s P S P^\circ^*$, $C, P_s P e C \acute{I} P \ddot{e} C_{\overline{H}} P S P \ddot{e} P N_e$ $P_{\mu} P i C - P r P_{\mu} C_{\overline{H}} P_j P^\circ P \gg C_{\overline{H}} P S P \ddot{e} P N_e$
 $P S P_{\mu} P e C_{\overline{H}} P_s P \gg C - P^*$.

$P - P_{\pm} P_s P e C \acute{z} P e C - C \acute{I} C, P e P_s P i P_s - P_j e \overline{H}^{\text{TM}} C_{\overline{H}} P \cdot P_s P i P_s C - C \acute{I} P \ddot{e} C \acute{I} C, P_{\mu} P_j P \ddot{e}$: $P S P_{\mu} C_{\overline{H}} P^\circ C \acute{I} C, P_s$ $\overline{H}$ “
 $P_j C - P^\circ P \gg P i C - C_{\overline{H}}$, $P_{\pm} C - P \gg C_{\overline{H}}$ $C r$ $P e C - P S C_{\overline{H}} C - P i P e P^\circ C \dots$

$P - P_{\pm} P_s P e C \acute{z} C \acute{I} P_{\mu} C_{\overline{H}} P_s P i P \ddot{e} P r C - P \gg C_{\overline{H}} P S P_s C - C \acute{I} P \ddot{e} C \acute{I} C, P_{\mu} P_j P \ddot{e}$: $P S P_{\mu} C_{\overline{H}} P^\circ C \acute{I} C, P_s$ $\overline{H}$ “
 $P i P_{\mu} P_j P^\circ C, C r C_{\overline{H}} C - C_{\overline{H}}$.

$P - P_{\pm} P_s P e C \acute{z} C_{\overline{H}} P_{\mu} P i C_{\overline{H}} P_s P r C \acute{z} P e C, P \ddot{e} P i P S P_s C - C \acute{I} P \ddot{e} C \acute{I} C, P_{\mu} P_j P \ddot{e}$: $C_{\overline{H}} C - P r P e P_s$ $\overline{H}$ “
 $P e C_{\overline{H}} P_s P i P_s C, P_{\mu} C_{\overline{H}} P^\circ$ $P \cdot C - C \acute{I} C, P^\circ C, P_{\mu} P i P_s P i P_s$ $C_{\overline{H}} P \gg P_{\mu} P S P^\circ$, $P i C_{\overline{H}} C - P^\circ P i C - P \cdot P_j^*$,
 $P i P_{\mu} P_j P^\circ C, P_s C \acute{I} P i P_{\mu} C_{\overline{H}} P_j C - C_{\overline{H}}$, $P i P_s P r P_s P i P \P P_{\mu} P S P^\circ$ $P_{\mu} C_{\overline{H}} P_{\mu} P e C_{\overline{H}} C - C_{\overline{H}}$.

$P - P^\circ P i P^\circ P \gg C_{\overline{H}} P S C - C_{\overline{H}} P_s P \cdot P \gg P^\circ P r P \ddot{e}$: $P S P_{\mu} C_{\overline{H}} P^\circ C \acute{I} C, P_s$ $\overline{H}$ “ $P_{\pm} C - P \gg C_{\overline{H}}$ $C r$ $P i C_{\overline{H}} C r P r C_{\overline{H}} C \dots$,
 $P i C - P r P i P \ddot{e} C \% P_{\mu} P S P^\circ$ $C \acute{I} C, P_s P_j P \gg C_{\overline{H}} P i P^\circ P S C - C \acute{I} C, C_{\overline{H}}$, $P I C - P r C_{\overline{H}} C r C, C, C_{\overline{H}}$ $P \P P^\circ C_{\overline{H}} C r$;
 $C_{\overline{H}} C - P r P e P_s$ $\overline{H}$ “ $P i P_s P r C_{\overline{H}} P^\circ P \cdot P S P_{\mu} P S P S C_{\overline{H}}$.

$P \gg P^\circ P_{\pm} P_s C_{\overline{H}} P^\circ C, P_s C_{\overline{H}} P S C - P i P_s P e P^\circ P \cdot P S P \ddot{e} P e P \ddot{e}$: $P S P_{\mu} C_{\overline{H}} P^\circ C \acute{I} C, P_s$ $\overline{H}$ “ $P i C - P r P i P \ddot{e} C \% P_{\mu} P S P^\circ$
 $C_{\overline{H}} P^\circ C \acute{I} C, P_s C, P^\circ$ $C \acute{I} P_{\mu} C_{\overline{H}} C_{\overline{H}} P_{\mu} P i P \ddot{e} C \dots$ $C \acute{I} P e P_s C_{\overline{H}} P_s C_{\overline{H}} P_{\mu} P S C_{\overline{H}}$.

* $P_{\mu} P_s P i C - P r P_s P_j P \gg C_{\overline{H}} P \gg P \ddot{e} P \gg P \ddot{e} C \epsilon P_{\mu}$ $P i C - P r$ $C_{\overline{H}} P^\circ C \acute{I} P r P_s C \acute{I} P \gg C - P r P \P P_{\mu} P S P S C_{\overline{H}}$ $P i C - C \acute{I} P \gg C_{\overline{H}}$
 $P i P \ddot{e} C \dots P_s P r C r$ $P i C_{\overline{H}} P_{\mu} P i P^\circ C_{\overline{H}} P^\circ C, C r$ $P S P^\circ$ $C_{\overline{H}} P \ddot{e} P S P_s P e$.

** $P_{\mu} P_s C_{\overline{H}} C r C \epsilon P_{\mu} P S P S C_{\overline{H}}$ $C \acute{I} P i C_{\overline{H}} P \ddot{e} P N_e P S C_{\overline{H}} C, C, C_{\overline{H}}$ $P e P_s P \gg C_{\overline{H}} P_s C_{\overline{H}} C r$: $C \dots P \gg P_s C_{\overline{H}} P_s P i C \acute{I} C - C_{\overline{H}}$,
 $C \dots C_{\overline{H}} P_s P_j P^\circ C, P_s P i C \acute{I} C - C_{\overline{H}}$, $C_{\overline{H}} C - P^\circ P S P_s P i C \acute{I} C - C_{\overline{H}}$, $P_{\mu} C_{\overline{H}} P \ddot{e} C, C_{\overline{H}} P_s P i C \acute{I} C - C_{\overline{H}}$,

PeCfP°PSC,PsPfcC-Cf.

*** PµPsCfCfCfPµPSPSCf CfP»CHPsP·PsPIPëPrC-P»PµPSPSCf: CfCfC...C-CfC,CH PI PsC±P°C..., PµPsCfCfCfPµPSPSCf CfP»CHPsP·PsPIPëPrC-P»PµPSPSCf C,P° Pfc-PrPIPëC%PµPSPSCf CfP»CHPsP·PsPIPëPrC-P»PµPSPSCf.

PkPëP¶C±PµP·P°P·PSP°C±PµPSC- CfPIPëC%P° CfPµPsCfC,PµCfC-PiP°P»PëCfCf Cf C...PsPrC- PrPsCfP»C-PrP¶PµPSCf; PfcCfPëC±PëPSPSPëPNë PIP·P°C" PjPsP·PIb¶™ CfP·PsPe PSPµ PIPëP·PSP°C±PµPSPëPNë. PµPsPIC-PrPsPjP»PµPSPSCf PIPëP»CfC±P°P»Pë CfPIPëC%P°, C%Ps PjP°P»Pë PIC-CfPµPiC-PrPSPëPNë P·PIb¶™ CfP·PsPe C-P· P·P°CfC,PsCfCfPIP°PSPSCfPj PfcCfPµPiP°CfP°C,Cf. PfcPIPëC%P°, CfPeC- PSPµ P±CfP»Pë P·P°P·PSP°C±PµPSC-, P±CfP»Pë P»PµPiPePëPjPë C,P° PµPsPIC-PrPsPjP»PµPSPSCf P±CfP»Pë PrCfP¶Pµ PSPµC,PsC±PSPëPjPë, C%PsP± PjP°C,Pë P·PSP°C±PµPSPSCf.

P—P°PiP°P»CHPSC-: PSP°P±CfCfPe PsP±P»PëC±CfCf, CfPµP°PeC†C-C— C,,PsC,PsC±CfC,P»PëPIPsCfC,C-, CëPsPe, P°CfC,PµPSC-Cf, P±C-P»CH, CfP°Pfc,PsPIPµ PiP°PrC-PSPSCf, P±C-P»CH Cf P¶PëPIPsC,C-, CfP°Pfc,PsPIPµ PiPsCëPePsPrP¶PµPSPSCf.

P— P±PsPeCé CfPµCfCfPµPIPs-CfCéPrPëPSPSPsC— CfPëCfC,PµPjPë: CfC,PµPSPsPeP°CfPrC-Cf, PjV-P±P»PsPeP°PrP°, PjC-PiCfPµPSCf, PµPsCfC,CfCfP°P»CHPSP° PiC-PµPsC,PµPSP·C-Cf, C-CëPµPjC-Cf PjC-PsPeP°CfPrP°, C,CfPsPjP±PsP· CfCfPrPëPS PiPsP»PsPIPsPiPs PjPsP·PeCf, CfP°Pfc,PsPIP° P·CfPiPëPSPeP° CfPµCfCfCf, PµPsCfCfCfPµPSPSCf CfPµP·CfP»CHC,P°C,C-PI PSP° P·PbP", PeP°CfPrC-PsPjC-PsPiP°C,C-Cf.

P— P±PsPeCé C,CfP°PIPsPiPs C,CfP°PeC,Cé: PiP»PsCfPëC,, PePsP»C-C,, PrPëCfC,,P°PiC-Cf, PiP°CfC,CfPëC,, PiP°CfC,CfPsPµPSC,PµCfPëC,, PµP·PsC,, P°PiC-C,, CfC,PsPjP°C,PëC,, PµPsCfCfCfPµPSPSCf CfPµP·CfP»CHC,P°C,C-PI PiPµC±C-PSPePsPIPëC... PfcCfPsP±, CfPµPeC,P°P»CHPSP° PeCfPsPIPsC,PµC±P°, PiC-PSPiC-PIC-C,,.

P— P±PsPeCé CfPëCfC,PµPjPë PeCfPsPIC-: P°PSPµPjC-Cf, P»PµPNëPePsPiPµPSC-Cf.

P— P±PsPeCé PjPµC,P°P±PsP»C-P·PjCé C,P° C...P°CfC±CéPIP°PSPSCf: CfPfcCfP°PiP°, PSP°P±CfCfPe, PµPsPrP°PiCfP°, PSPµCfC,P°P±C-P»CHPSPëPNë PrC-P°P±PµC,, PiC-PiPµCfPiP»C-PePµPjC-Cf, PiPµCfPëC,,PµCfPëC±PSC- PSP°P±CfCfPePë, PiC-PiPµCfCfCfPëPePµPjC-Cf, PiC-PiPsPiP»C-PePµPjC-Cf, PiC-PiPµCfPSP°C,CfC-C" PjC-Cf.

P— P±PsPeCé PeC-CfC,PePsPIPs-Pjeb¶™ CfP·PsPIPsC— CfPëCfC,PµPjPë: P°CfC,CfPëC,, P°CfC,CfPsP°, CfPsP·CfPëPI CfCfC...PsP¶PëP»P»Cf, C,PµPSPsCfPëPSPsPIC-C,, P±C-P»CH Cf PeC-CfC,PeP°C..., PjC-P°CfC,PµPSC-Cf, CfPëPSPsPIC-C,,.

P— P±PsPeCé PSPµCfPIPsPIPsC— CfPëCfC,PµPjPë: P°C,P°PeCfC-Cf, PSPµPICfP°P»PiC-Cf, PSPµPNëCfPsPiP°C,C-Cf, PiP°CfPµCfC,PµP·C-Cf, C,CfPµPjPsCf, PIPµCfC,PëPiPs, PrPµPiCfPµCfC-Cf, P±PµP·CfPsPSPSCf, P°PSPsPjP°P»CHPSC- CfPSPsPIPëPrC-PSPSCf, P·PSPëP¶PµPSPSCf CfPµC,,P»PµPeCfC-PI.

P— P±PsPeCé PrPëC...P°P»CHPSPsC— CfPëCfC,PµPjPë: P°CfC,PjP°, PrPëCfPiPSPsPµ, P»P°CfPëPSPiC-C,, C,,P°CfPëPSPiC-C,, CfPëPSCfCfPëC,, P±CfPsPSC...C-C,, PµPsCfPëP»PµPSPµ CfP»PëPSPsPIPëPrC-P»PµPSPSCf, PµPsCfPëP»PµPSPSCf PeP°CëP»Cf.

P— P±PsPeCé CëPeC-CfPë: PeCfPsPiPëPIb¶™ CfPSPeP°, PiPµCfPiPµCf, CfPiPµCfP±C-P¶, PiC-C,P»PëPIC-CfC,CH, PIPëCfP°P·PePë CëPeC-CfPë, PePsPSC,P°PeC,PSPëPNë PrPµCfPjP°C,PëC,, PµPeCfC,,PsP»C-P°C,PëPIPsPëPNë PrPµCfPjP°C,PëC,,.

PŸPiPµC†PëC,,C-C‡PSC- PIC-PrC‡C¿C,C,CŬ: CŦP°PïC,PsPIPµ P·PSPëP¶PµPSPSCŬ C‡Pë PIC,CŦP°C,P° CŦP»CŦC...CŦ, P±C-P»CH CŦ PICŦC...P°C..., PeCŦPsPIPsPIPëP»PëPI CŦ PsPePs, PeP°C,P°CŦP°PeC,P°, CŦCŦC...C-CŦC,CH PsC‡PµPNë.

P— P±PsPeC¿ C¿CŦPsPiPµPSC-C,P°P»CHPSPsC— CŦPëCŦC,PµPjPë: C†PëCŦC,PëC,, PSC-PeC,CŦCŦC-CŬ, PïC-PrPIPëC%PµPSP° C‡P°CŦC,PsC,P° CŦPµC‡PsPIPëPïCŦCŦP°PSCĤ, P·P±C-P»CHC€PµPSPSCŬ PjPsP»PsC‡PSPëC... P·P°P»PsP·, PSPµC,CŦPëPjP°PSPSCŬ CŦPµC‡C-, PïPsCŦCŦC€PµPSPSCŬ PµCŬPeCŦP»CŬC†C-C—, PSP°P±CŦCŬPe CŦC,P°C,PµPIPëC... PsCŦPiP°PSC-PI, P°PSPsCŦPiP°P·PjC-CŬ.

PµC-CŦP»CŬ PIPëC...PsPrCŦ PïCŦPµPïP°CŦP°C,CŦ PSP° CŦPëPSPsPe P±CŦP»Pë C-PrPµPSC,PëC,,C-PePsPIP°PSC- PSPëP¶C‡PµP·P°P·PSP°C‡PµPSC- PïPsP±C-C‡PSC- CŦPµP°PeC†C-C—. PĥCŦPeC-P»CHPePë PïCŦPs C,P°PeC- CŦPµP°PeC†C-C— PïPsPIC-PrPsPjP»CŬCĤC,CH PrPsP±CŦPsPIC-P»CHPSPs C,P° PïPsPIC-PrPsPjP»PµPSPSCŬ PSP°PrC...PsPrCŬC,CH PIC-Pr PïPsPïCŦP»CŬC†C-C— PSPµPIPëP·PSP°C‡PµPSPsC— C‡PëCŦPµP»CHPSPsCŦC-, PSPµ P·P°PIP¶PrPë PjPsP¶PSP° PrPsCŦC,PsPIC-CŦPSPs PsC†C-PSPëC,Pë C—C... C‡P°CŦC,PsC,CŦ C,P° PICŦC,P°PSPsPIPëC,Pë PïCŦPëC‡PëPSPSPëPNë P·PiBŦ™ CŬP·PsPe P· PµPeCŦPïPsP·PëC†C-C”CĤ P»C-PeP°CŦCŦCHPePsPiPs P·P°CŦPsP±CŦ. PïC- CŬPIPëC%P° P±CŦP»Pë P·P°P·PSP°C‡PµPSC- CŬPe C‡PµCŦPµP· C—C... CŦPµCŦPNëPsP·PSC-CŦC,CH, C‡P°CŦC,PsC,CŦ PïPsPIC-PrPsPjP»PµPSPSCŬ, PIC-PrCŦCŦC,PSC-CŦC,CH C‡C-C,PePsPiPs P°P»CHC,PµCŦPSP°C,PëPIPSsPiPs P·PiBŦ™ CŬP·PeCŦ, C,P°Pe C- C‡PµCŦPµP· PePsPjP±C-PSP°C†C-CĤ C†PëC... C,,P°PeC,PsCŦC-PI.

PŸPµCŦC†PµPIPs-CŦC¿PrPëPSPSC- C,P° C†PµCŦPµP±CŦPsPIP°CŦPeC¿P»CŬCŦPSC- CŬPIPëC%P°. PµPsPIC-PrPsPjP»CŬP»Pë PïCŦPs CŦPµCŦPNëPsP·PSC- CŦPµCŦC†PµPIPs- CŦCŦPrPëPSPSC-, C†PµCŦPµP±CŦPsPIP°CŦPeCŦP»CŬCŦPSC- C,P° CŦCŦPrPëPSPSC- CŬPIPëC%P°, PIPeP»CHC‡P°CHC‡Pë C†PµCŦPµP±CŦPsPIP°CŦPeCŦP»CŬCŦPSCŦ PeCŦPsPIPsC,PµC‡CŦ, CŦCŦP±P°CŦP°C...PSPsC—PrP°P»CHPSCŦ C,P° PIPSCŦC,CŦC-C€PSCĤPsC†PµCŦPµP±CŦP°P»CHPSCŦ PeCŦPsPIPsC,PµC‡CŦ C,P° P»PµPiPµPSPµPICŦ PeCŦPsPIPsC,PµC‡CŦ, C%Ps P±CŦP»Pë PïPsPiBŦ™ CŬP·P°PSC- CŦ C‡P°CŦC- C-P· P·P°CŦC,PsCŦCŦPIP°PSPSCŬPj PïCŦPµPïP°CŦP°C,CŦ PŸC-PSPµPiCŦP°. P·C-P»CHC€C-CŦC,CH PïP°C†C-C”PSC,C-PI, P°P»Pµ PSPµ PICŦC-, PjP°P»Pë C-CŦPSCŦCHC‡C- C,,P°PeC,PsCŦPë CŦPµCŦC†PµPIPs-CŦCŦPrPëPSPSPsPiPs CŦPëP·PëPeCŦ. PµPsPIC-PrPsPjP»CŬP»Pë, C%Ps P±P°PiP°C,Ps P· C†PëC... CŬPIPëC% PIPëPSPëPeP»Pë PïC-Pr C‡P°CŦ P°P±Ps PsPrCŦP°P·CŦ PïC-CŦP»CŬ CŦPµPeCŦCŦP°P»CHPSPsC— P°PeC,PëPIPSsCŦC-, C,P° PrPµPeC-P»CHPeP° CŬPIPëC% PIPëPSPëPeP»Pë PsPrCŦP°P·CŦ PïC-Pr C‡P°CŦ P·P°CŦC,PsCŦCŦPIP°PSPSCŬ PïCŦPµPïP°CŦP°C,CŦ PŸC-PSPµPiCŦP° P±PµP· CŦPµPeCŦCŦP°P»CHPSPsC— P°PeC,PëPIPSsCŦC-, P†PSC€C- CŬPIPëC%P° PIPëPSPëPeP»Pë PïCŦPsC,CŬPiPsPj PSP°CŦC,CŦPïPSPëC... PiPsPrPëPS C‡Pë PrPSC-PI PïC-CŦP»CŬ P·P°CŦC,PsCŦCŦPIP°PSPSCŬ PïCŦPµPïP°CŦP°C,CŦ PŸC-PSPµPiCŦP° C,P° CŦPµPeCŦCŦP°P»CHPSPsC— P°PeC,PëPIPSsCŦC-, PĥPµPjPsP¶P»PëPIPs PICŦC,P°PSPsPIPëC,Pë, C‡Pë PjP°CHC,CH C†C- CŬPIPëC%P° PïCŦCŬPjPëPNë P·PiBŦ™ CŬP·PsPe C-P· P·P°CŦC,PsCŦCŦPIP°PSPSCŬPj PïCŦPµPïP°CŦP°C,CŦ, C-P· CŦPµPeCŦCŦP°P»CHPSPsCH P°PeC,PëPIPSC-CŦC,CĤ, P· PSP°CŬIPSPëPjPë C,,P°PeC,PsCŦP°PjPë CŦPëP·PëPeCŦ C‡Pë P· PePsPjP±C-PSP°C†C-C”CĤ C†PëC... C,,P°PeC,PsCŦC-PI P°P±PsB P· C-PSC€PëPjPë C,,P°PeC,PsCŦP°PjPë.

P— P±PsPeC¿ PeCŦPsPIPsPSPsCŦPSPsC— C,P° P»C-PjC,,P°C,PëC‡PSPsC— CŦPëCŦC,PµPj: PIP°P·PsPsPeP»CHP·PëPIPSPëPNë PeCŦPëP· Pj PSPµPIPµP»PëPePsPjCŦ P·P°PIC‡P°CŦPSPs PïCŦPëPïPëPSPµPSPsPjCŦ PrPsCŦP»C-PrP¶PµPSPSC- P·P°CŦC,PsCŦCŦPIP°PSPSCŬ PïCŦPµPïP°CŦP°C,CŦ P PµPIP°C†PëPs (CŦPëP»PrPµPSP°C,,C-P») PrP»CŬ PïP°C†C-C”PSC,C-PI

C-P· P»PμPiPμPSPμPIPsCḡ P°CḡC,PμCḡC-P°P»CHbPSPsCḡ PiC-PiPμCḡC,PμPSP·C-C''Cḡ,
PIC,PsCḡPëPSPSPsCḡ C%PsPrPs CḡPμCḡPiPsPiPsPrC-P±PSPsPeP»C-C,PëPSPSPsC—
P°PSPμPjC-C—, PiCḡPë P·P°CḡC,PsCḡCḡPIP°PSPSC- CḡPëP»PrPμPSP°C,,C-P»Cḡ PiCḡPs
CḡPsP·PIPëC,PsPe PIP°P·PsPsPeP»CḡP·PëPIPSPëC... PeCḡPëP·C-PI, C%Ps
PiPsC,CḡPμP±CḡPIP°P»Pë PiPsCḡPiC-C,P°P»C-P·P°CḡC-C—, PiPsPIC-PrPsPjP»CḡP»Pë
CḡP°CḡC,C-CëPμ, PSC-Pḡ PiCḡPë P·P°CḡC,PsCḡPsPICḡPIP°PSPSC- PiP»P°CḡPμP±Ps.
PḡP»C-PSC-CḡPSPμ P·PSP°CḡPμPSPSCḡ CḡC-C''C— C-PSC,,PsCḡPjP°CḡC-C— PrP»Cḡ
PiP°CḡC-C''PSC,C-PI, CḡPeC- P·P°CḡC,PsCḡPsPICḡCḡC,CHb PiCḡPμPiP°CḡP°C, PḡC-PSPμPiCḡP°
P· PjPμC,PsCḡ P»C-PeCḡPIP°PSPSCḡ PμCḡPμPeC,PëP»CHbPSPsC— PrPëCḡC,,CḡPSPeCḡC-C—, C''
PSPμPIC-PrPsPjPëPj.

*PḡPμCḡPIPsPIP° CḡPëCḡC,PμPjP°: C,CḡPëPIPsPḡPSC-CḡC,CHb, C,CḡP°PSP·PëC,PsCḡPSP°
PiP»PsP±P°P»CHbPSP° P°PjPSPμP·C-Cḡ.*

PḡPiPμCḡPëC,,C-CḡPSC- PIC-PrCḡCḡC,C,Cḡ.

*PḡP»CḡC.... PμC-CḡP»Cḡ PIPëC...PsPrCḡ PiCḡPμPiP°CḡP°C,Cḡ PSP° CḡPëPSPsPe
PiPsPIC-PrPsPjP»CḡP»Pë PiCḡPs PIPëPiP°PrPePë CḡP°PiC,PsPIPsPiPs P·PSPëPḡPμPSPSCḡ
P°P±Ps PIC,CḡP°C,Pë CḡP»CḡC...Cḡ, PiPsPiḡḡ™ CḡP·P°PSC- Cḡ CḡP°CḡC- C-P·
P·P°CḡC,PsCḡCḡPIP°PSPSCḡPj PiCḡPμPiP°CḡP°C,Cḡ PḡC-PSPμPiCḡP°. Pj PrPμCḡPePëC...
PIPëPiP°PrPeP°C... PiPsPIC-PrPsPjP»CḡP»Pë PiCḡPs PSP°CḡPIPS- CḡC,CHb PjPμPrPëCḡPSPëC...
CḡC,P°PSC-PI C,P° C-PSCëPëC... C,,P°PeC,PsCḡC-PI, C%Ps PjPsPiP»Pë PIC-PrC-PiCḡP°C,Pë
CḡPsP»CHb Cḡ CḡPsP·PIPëC,PeCḡ PiPsP±C-CḡPSPëC... CḡPμP°PeCḡC-PNḡ P· P±PsPeCḡ
CḡP»CḡC...Cḡ. Pj P±P°PiP°C,CHbPsC... PIPëPiP°PrPeP°C... C-PSC,,PsCḡPjP°CḡC-Cḡ C%PsPrPs
PiPsPrP°P»CHbCëPsPiPs PjPμPrPëCḡPSPsPiPs CḡPiPsCḡC,PμCḡPμPḡPμPSPSCḡ
PIC-PrCḡCḡC,PSCḡ. P'PëP·PSP°CḡPëC,Pë, CḡPë CḡC- CḡPIPëC%P° PiCḡCḡPjPs
PiPsPiḡḡ™ CḡP·P°PSC- C-P· P·P°CḡC,PsCḡCḡPIP°PSPSCḡPj PiCḡPμPiP°CḡP°C,Cḡ
PḡC-PSPμPiCḡP°, P· PSP°CḡPIPSPëPjPë C,,P°PeC,PsCḡP°PjPë CḡPëP·PëPeCḡ PIC,CḡP°C,Pë
CḡP»CḡC...Cḡ, P· PePsPjP±C-PSP°CḡC-C''Cḡ CḡPëC... C,,P°PeC,PsCḡC-PI CḡPë P· C-PSCëPëPjPë
C,,P°PeC,PsCḡP°PjPë, PSPμPjPsPḡP»PëPIPs.*

*P—C-Cḡ. PḡPëPjCḡP°CḡPsPIP° PIC,CḡP°C,P° P·PsCḡCḡ, PiPsCḡPμCḡPIPsPSC-PSPSCḡ PsCḡPμPNḡ,
PiPμCḡC-PSPSCḡ PI PsCḡP°C..., PiC-PrPIPëC%PμPSPSCḡ PIPSCḡC,CḡC-CëPSCḡPsPsCḡPSPsPiPs
C,PëCḡPeCḡ, PSP°P±CḡCḡPe CḡC-C,PeC-PIPePë, CḡCḡPrPëPSPSC-
P·P°C...PIPsCḡCḡPIP°PSPSCḡ CḡC-C,PeC-PIPePë CḡPë PeCḡPsPIPsC,PμCḡP°,
PIC-PrCëP°CḡCḡPIP°PSPSCḡ CḡPeP»PëCḡC,PsPiPs C,C-P»P°.*

*PμC-CḡP»Cḡ PIPëC...PsPrCḡ PiCḡPμPiP°CḡP°C,Cḡ PSP° CḡPëPSPsPe CḡC-PrPePs
PiPsPIC-PrPsPjP»CḡP»PsCḡCḡ PiCḡPs PIPëPiP°PrPePë PSPμP°CḡC,PμCḡC-P°P»CHbPSPsC—
PiPμCḡPμPrPSCḡPsC— C-CëPμPjC-CḡPSPsC— PSPμPICḡPsPiP°C,C-C— P·PsCḡPsPIPsPiPs
PSPμCḡPIP°, C%Ps C'' PiCḡPëCḡPëPSPsCḡ P·PSPëPḡPμPSPSCḡ P·PsCḡCḡ,
PIPeP»CḡCḡP°CḡCḡPë PiPsCḡC,C-PNḡPSCḡ PIC,CḡP°C,Cḡ P·PsCḡCḡ, CḡPeC- P±CḡP»Pë
PiPsPiḡḡ™ CḡP·P°PSC- Cḡ CḡP°CḡC- C-P· P·P°CḡC,PsCḡCḡPIP°PSPSCḡPj
C-PSPiC-P±C-C,PsCḡC-PI PḡP''P·5, PIPeP»CḡCḡP°CḡCḡPë PiCḡPμPiP°CḡP°C, PḡC-PSPμPiCḡP°.
P'P°PiP°C,Ps P· PiP°CḡC-C''PSC,C-PI, P°P»Pμ PSPμ PICḡC-, PjP°P»Pë PSP°CḡPIPS-
P°PSP°C,PsPjC-CḡPSC- P°P±Ps CḡCḡPrPëPSPSC- C,,P°PeC,PsCḡPë CḡPëP·PëPeCḡ
CḡPsP·PIPëC,PeCḡ PSPμP°CḡC,PμCḡC-P°P»CHbPSPsC— PiPμCḡPμPrPSCḡPsC—
C-CëPμPjC-CḡPSPsC— PSPμPICḡPsPiP°C,C-C— P·PsCḡPsPIPsPiPs PSPμCḡPIP°,
PIPeP»CḡCḡP°CḡCḡPë (P°P»Pμ PSPμPsP±PsPiḡḡ™ CḡP·PePsPIPs PsP±PjPμPḡCḡCḡCḡPëCḡCḡ)
C,P°PeC-: PSPëP·CHbPePμ CḡPiC-PIPIC-PrPSPsCëPμPSPSCḡ PrC-P°PjPμC,CḡP°
PμPeCḡPeP°PIP°CḡC-C— C,P° PrPëCḡPeP° P·PsCḡPsPIPsPiPs PSPμCḡPIP° (P·P°CḡC,C-PNḡPSPëPNḡ
PrPëCḡPe P·PsCḡPsPIPsPiPs PSPμCḡPIP°), PIC-Pe PiPsPSP°Pr 50 CḡPsPeC-PI,*

$P^{\circ}C\bar{H}C, P_{\mu}C\bar{H}C - P^{\circ}P \gg CH\bar{H}PSP^{\circ} \text{ PiC} - \bar{P}iP_{\mu}C\bar{H}C, P_{\mu}PSP^{\circ}C - C\bar{H}, P^{\circ}C \dots PIPsC\bar{H}C\bar{H}PIP^{\circ}PSPSC\bar{H}$
 $PePsC\bar{H}PsPSP^{\circ}C\bar{H}PSP\bar{e}C \dots P^{\circ}C\bar{H}C, P_{\mu}C\bar{H}C - PN_{\bar{e}}, \text{ PiC} - \bar{P}iP_{\mu}C\bar{H}P \gg C - \bar{P}iC - PrP_{\mu}PjC - C\bar{H} \text{ C}, P^{\circ}$
 $\bar{P}iP^{\circ}P \gg C - PSPSC\bar{H}. P\bar{k}P_{\mu}PjPsP\bar{H}P \gg P\bar{e}PIPs \text{ PIP}\bar{e}P \cdot PSP^{\circ}C\bar{H}P\bar{e}C, P\bar{e}, C\bar{H}P\bar{e} \text{ C}\bar{H}C - C\bar{H}PIP\bar{e}C\%P^{\circ}$
 $\bar{P}iC\bar{H}C\bar{H}PjPs \bar{P}iPsPIB\bar{H}^{\text{TM}}C\bar{H}P \cdot P^{\circ}PSC - C - P \cdot P^{\circ}C\bar{H}C, PsC\bar{H}C\bar{H}PIP^{\circ}PSPSC\bar{H}Pj$
 $C - PSPiC - P\pm C - C, PsC\bar{H}C - PI \text{ P}\alpha P^{\circ}P \cdot 5 \text{ C}\bar{H}P\bar{e} \text{ P} \cdot PSP^{\circ}C\bar{H}PIPSP\bar{e}PjP\bar{e} \text{ P}^{\circ}PSP^{\circ}C, PsPjC - C\bar{H}PSP\bar{e}PjP\bar{e}$
 $P^{\circ}P\pm Ps \text{ C}\bar{H}C\bar{H}PrP\bar{e}PSPSP\bar{e}PjP\bar{e} \text{ C}, P^{\circ}PeC, PsC\bar{H}P^{\circ}PjP\bar{e} \text{ C}\bar{H}P\bar{e}P \cdot P\bar{e}PeC\bar{H}, C\bar{H}P\bar{e} \text{ P} \cdot$
 $PePsPjP\pm C - PSP^{\circ}C\bar{H}C - C^{\circ}C\bar{H} \text{ C}\bar{H}P\bar{e}C \dots PIC\bar{H}C - C \dots C, P^{\circ}PeC, PsC\bar{H}C - PI, C\bar{H}P\bar{e} \text{ P} \cdot C - PSC\bar{H}P\bar{e}PjP\bar{e}$
 $C, P^{\circ}PeC, PsC\bar{H}P^{\circ}PjP\bar{e}.$

$P - PIC - C, C\bar{H}PIP^{\circ}PSPSC\bar{H} \bar{P}iC\bar{H}Ps \bar{P}iC - PrPsP \cdot C\bar{H}C\bar{H}PIP^{\circ}PSC - \bar{P}iPsP\pm C - C\bar{H}PSC - C\bar{H}P_{\mu}P^{\circ}PeC\bar{H}C - C -$
 $P - PIC - C, C\bar{H}PIP^{\circ}PSPSC\bar{H} \bar{P}iC\bar{H}Ps \bar{P}iC - PrPsP \cdot C\bar{H}C\bar{H}PIP^{\circ}PSC - \bar{P}iPsP\pm C - C\bar{H}PSC - C\bar{H}P_{\mu}P^{\circ}PeC\bar{H}C - C -$
 $\bar{P}iC - C\bar{H}P \gg C\bar{H} \text{ C}\bar{H}P_{\mu}C^{\circ}C\bar{H}C, C\bar{H}P^{\circ}C\bar{H}C - C - P \gg C - PeP^{\circ}C\bar{H}C\bar{H}PePsPiPs \text{ P} \cdot P^{\circ}C\bar{H}PsP\pm C\bar{H} \text{ PjP}^{\circ}C^{\circ}$
 $PIP^{\circ}P\bar{H}P \gg P\bar{e}PIP_{\mu} \text{ P} \cdot PSP^{\circ}C\bar{H}P_{\mu}PSPSC\bar{H}. P_{\bar{i}}P_{\mu} \text{ PrP}^{\circ}C^{\circ} \text{ P} \cdot PjPsPiC\bar{H} \bar{P}iC\bar{H}PsPIPsPrP\bar{e}C, P\bar{e}$
 $P\pm P_{\mu}P \cdot \bar{P}iP_{\mu}C\bar{H}P_{\mu}C\bar{H}PIPSP\bar{e}PN_{\bar{e}} \text{ PjPsPSC} - C, PsC\bar{H}P\bar{e}PSPi \text{ C}\bar{H}P\bar{e}C - PIPIC - PrPSPsC\bar{H}P_{\mu}PSPSC\bar{H}$
 $PePsC\bar{H}P\bar{e}C\bar{H}C, C - C - C\bar{H}P\bar{e}P \cdot P\bar{e}PeC - PI, \bar{P}iPsPIB\bar{H}^{\text{TM}}C\bar{H}P \cdot P^{\circ}PSP\bar{e}C \dots C - P \cdot$
 $P^{\circ}C\bar{H}C, PsC\bar{H}C\bar{H}PIP^{\circ}PSPSC\bar{H}Pj \text{ C}\bar{H}PePsPiPs \text{ P} \gg C - PeP^{\circ}C\bar{H}C\bar{H}PePsPiPs \text{ P} \cdot P^{\circ}C\bar{H}PsP\pm C\bar{H}.$
 $P \gg C - PeP^{\circ}C\bar{H}C\bar{H}Pj \text{ C}\bar{H}P \gg C - Pr \text{ P} \cdot PIC - C, C\bar{H}PIP^{\circ}C, P\bar{e} \bar{P}iC\bar{H}Ps \text{ P}\pm C\bar{H}PrC\bar{H}C - C\bar{H}PeC -$
 $\bar{P}iC - PrPsP \cdot C\bar{H}C\bar{H}PIP^{\circ}PSC - \bar{P}iPsP\pm C - C\bar{H}PSC - C\bar{H}P_{\mu}P^{\circ}PeC\bar{H}C - C - PIC - Pr\bar{P}iPsPIC - PrPSPs \text{ PrPs}$
 $PIP\bar{e}PjPsPi \text{ P} \cdot P^{\circ}PePsPSPsPrP^{\circ}PIC\bar{H}C, PIP^{\circ}.$

$P\bar{H}P_{\mu}C\bar{H}PjC - PS \bar{P}iC\bar{H}P\bar{e}PrP^{\circ}C, PSPsC\bar{H}C, C -$.

$P\bar{H}P^{\circ}P\pm P \gg P_{\mu}C, PeP\bar{e} \bar{P}iPs \text{ 25 PjPi: 3 C}\bar{H}PsPeP\bar{e}.$

$P\bar{H}P^{\circ}P\pm P \gg P_{\mu}C, PeP\bar{e} \bar{P}iPs \text{ 50 PjPi C}, P^{\circ} \bar{P}iPs \text{ 100 PjPi: 4 C}\bar{H}PsPeP\bar{e}.$

$PjPjPsPIP\bar{e} \text{ P} \cdot P\pm P_{\mu}C\bar{H}C - \bar{P}iP^{\circ}PSPSC\bar{H}. B$

$P - P\pm P_{\mu}C\bar{H}C - \bar{P}iP^{\circ}C, P\bar{e} \bar{P}iC\bar{H}P\bar{e} \text{ C}, P_{\mu}Pj\bar{P}iP_{\mu}C\bar{H}P^{\circ}C, C\bar{H}C\bar{H}C - PSP_{\mu} \text{ PIP}\bar{e}C\%P_{\mu} \text{ 25 B}^{\circ}P\bar{H} \text{ PI}$
 $PsC\bar{H}P\bar{e}PiC - PSP^{\circ}P \gg CH\bar{H}PSC - PN_{\bar{e}} \text{ C}\bar{H}P\bar{e}P^{\circ}PePsPIC\bar{H}C -$

$P - P\pm P_{\mu}C\bar{H}C - \bar{P}iP^{\circ}C, P\bar{e} \text{ C}\bar{H} PSP_{\mu}PrPsC\bar{H}C, C\bar{H}P\bar{e}PSPsPjC\bar{H} \text{ PrP} \gg C\bar{H} \text{ PrC} - C, P_{\mu}PN_{\bar{e}} \text{ PjC} - C\bar{H}C\bar{H}C -$

$Pj\bar{P}iP^{\circ}PePsPIP\bar{e}P^{\circ}.$

$P_{\bar{H}}Ps \text{ 1 C}, P^{\circ}P\pm P \gg P_{\mu}C, C\bar{H}C - C\bar{H} \text{ P}\pm P \gg C - C\bar{H}C, P_{\mu}C\bar{H}C - , \bar{P}iPs \text{ 1 P}\pm P \gg C - C\bar{H}C, P_{\mu}C\bar{H}C\bar{H} \text{ C}\bar{H}$
 $PeP^{\circ}C\bar{H}C, PsPSPSC - PN_{\bar{e}} \text{ C}\bar{H}P\bar{e}P^{\circ}PePsPIC\bar{H}C -$

$P_{\bar{H}}Ps \text{ 4 C}, P^{\circ}P\pm P \gg P_{\mu}C, PeP\bar{e} \text{ C}\bar{H} \text{ P}\pm P \gg C - C\bar{H}C, P_{\mu}C\bar{H}C - , \bar{P}iPs \text{ 1 P}\pm P \gg C - C\bar{H}C, P_{\mu}C\bar{H}C\bar{H} \text{ C}\bar{H}$
 $PeP^{\circ}C\bar{H}C, PsPSPSC - PN_{\bar{e}} \text{ C}\bar{H}P\bar{e}P^{\circ}PePsPIC\bar{H}C -$

$P\bar{H}P^{\circ}C, P_{\mu}PiPsC\bar{H}C - C\bar{H} PIC - Pr\bar{P}iC\bar{H}C\bar{H}PeC\bar{H}.$

$$P-P^{\circ} \quad C\bar{H}P_{\mu}C^{\dagger}P_{\mu}P^{\ddagger}C, P_sP_j.$$

P'PëCᵗᵃPₛP±PSPëPε.

$$\text{P}\acute{\text{K}}\text{P}\text{sP}\pm\text{P}_{\mu}\text{P}\gg\text{P}\dagger\text{P}\gg\text{P}^{\circ}\text{C}\ddagger\text{P}\ddot{\text{Y}}\text{P}^{\circ}\text{P}\text{S}\text{P}^{\circ}\text{C}-\text{P}\text{I}\text{P}_{\mu}\text{P}\ddot{\text{Y}}\text{C}-\text{P}\text{r}\text{P}\P\text{P}^{\circ}\text{C}\text{H}\text{P}_{\mu}\text{C},\text{P}_{\text{h}}.\text{P}\ddot{\text{E}}.$$

**Р_НС-СГ⁺С†Р_μР·Р_{SP}°С...Р_SР_ГР_¶Р_μР_{SP}SC‡ PIPëC‡Р_SР±Р_{SP}ëР_εР° С,Р° Р°Р_ГС‡Р_РСГ⁺Р°
Р_ГС-СГ⁺С‡С‡ PïC‡Р_SР_{IP}°Р_ГР_¶Р_μР_{SP}SC‡ Р_NР_SР_иР_S Р_ГС-С‡Р_Р»С‡Р_{SP}Р_SСГ⁺С,С-.**

Р_ЛPIP°C₇C, P°P» P_YP°PSPeP°PeP»P°C₇, PiC₇. P•CГPеC- PhPeC±P°PePsPrP¶P°, B,, -299, 81100 Pj. P"CHP·PrP¶Pμ, P_YCrC₇PμC±C±PөPSP°.