

Р†РќРЎРЎР РЈРЉРІР†РЇ

$$\begin{array}{c} \text{PrP} \gg \text{CII} \text{ PjP}_{\mu} \text{PrP}^{\text{Sc}} \pm \text{PSPsPiPs} \text{ P} \cdot \text{P}^{\circ} \text{C}^{\text{I}} \text{C}, \text{PsC}^{\text{I}} \text{C}^{\text{r}} \text{PIP}^{\circ} \text{PSPSCII} \text{ P} \gg \text{C} - \text{PeP}^{\circ} \text{C}^{\text{I}} \text{C}^{\text{r}} \text{CHPePsPiPs} \\ \text{P} \cdot \text{P}^{\circ} \text{C}^{\text{I}} \text{PsP} \pm \text{C}^{\text{r}} \end{array}$$

P P•PḥP“P†PḡPḥ

(REAGILA^{B®})

$$P\check{Y}P\epsilon P\gg P^{\circ}P\Gamma:$$

РљР°РјРµСЂР»Рё С, РїРµСђРґ-РіРѕ 1,5 РјРі

$P''C-CHC\ddagger P^{\circ} CH_2P\mu C\ddagger PsPIP\ddot{e}PSP^{\circ}: PeP^{\circ}CHPePiCHP^{\circ}P\cdot PePS;$

1 PeP°PiCǱCǱP»P° PjC-CǱC,PēC,CH PeP°CǱPePiCǱP°P·PēPSCǱ 1,5 PjPi (CǱ PIPēPiP»CǱPrC-PeP°CǱPePiCǱP°P·PēPSCǱ PiC-PrCǱPsC...P»PsCǱPePrCǱ 1,635 PjPi).

$$P''PsPiPsPjC-P\PSC-C\overline{H}P\mu C\#PsPIP\ddot{e}PSP\ddot{e}:PeC\overline{H}PsC...PjP^{\circ}P\gg CH\ PeC\acute{r}PeC\acute{r}C\overline{H}C\acute{r}PrP\cdot C\PSP\ddot{e}PN_{\circ}$$

$$PiC\overline{H}P\mu P\P\P\mu P\gg P^{\circ}C,P\ddot{e}PSC-P\cdot PsPIP^{\circ}PSP\ddot{e}PN_{\circ},PiP^{\circ}PiPSC-C\overline{H}\ C\acute{r}C,P\mu P^{\circ}C\overline{H}P^{\circ}C,.$$

P_{9b}P^oPiC[†]CzP»Pë C,PIPμCT₅PrC-: C,PëC,P^oPSCr[†] PrC-PsPeC[†]PëPr, P• 171; P¶PμP»P^oC,PëPS; C‡PsC₅PSPëP»Ps C‡PsC₅PSPsPiPs PePsP»CHbPsC₅Cr[†] PrP»C¶ PrC₅Cr[†]PeCr[†]: P•P^oP»C-P•P^o PsPeC[†]PëPr C‡PsC₅PSPëPN_ë, P• 172; C€PμP»P^oPe; PμC,P^oPSPsP»; PIPsPrP^o; PiC₅PsPiC-P»PμPSPiP»C-PePsP»CHb; C-P•PsPiC₅PsPiP^oPSPsP»; P±Cr[†]C,P^oPSPsP»; P^oPjC-P^oPe PIPsPrPSPëPN_ë; PeP^oP»C-CT PiC-PrC₅PsPeC[†]PëPr.

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$$P''C-CHC\ddagger P^{\circ} CHP_{\mu}C\ddagger PsPIP\ddot{e}PSP^{\circ}: PeP^{\circ}CHP\ddot{e}P\ddot{i}CHP^{\circ}P\cdot P\ddot{e}PS;$$

1 PeP°PiCíCíP»P° PjC-CíC,PëC,CH PeP°CíPëPiCíP°P·PëPSCí 3 PjPi (Cí PIPëPiP»CíPrC-PeP°CíPëPiCíP°P·PëPSCí PiC-PrCíPSc...P»PsCíPëPrCí 3,270 PjPi).

$P''PsPiPsPjC-P\PSC-C\overline{H}P\mu C\#PsPIP\ddot{e}PSP\ddot{e}$: $PeC\overline{H}PsC...PjP^{\circ}P\gg CH$ $PeC\acute{r}PeC\acute{r}C\overline{H}C\acute{r}PrP\cdot C\PSP\ddot{e}PN_0$
 $PiC\overline{H}P\mu P\P P\mu P\gg P^{\circ}C,P\ddot{e}PSC-P\cdot PsPIP^{\circ}PSP\ddot{e}PN_0$, $PjP^{\circ}PiPSC-C\overline{H}$ $C\acute{r}C$, $P\mu P^{\circ}C\overline{H}P^{\circ}C,..$

*P*_h*P*^o*PiCl'CzP*»*Pë C,PIPμCTjPrC*–: *C,PëC,P°PSCr PrC*–*PsPeCl'PëPr*, *P*• 171; *P¶PμP*»*P°C,PëPS*;
Cl'PiPμC†C–*P°P*»*CHbPSPëPNë C†PμCTjPIPsPSPëPNë PhPÿ*, *P*• 129; *PrC*–*P°PjP°PSC,PsPIPëPNë*
Cl'PëPSC–*PNë FCF*, *P*• 133; *P·P°P*»*C*–*P·P° PsPeCl'PëPr P¶PsPIC,PëPNë*, *P*• 172; *C†PsCTjPSPëP*»*Ps*
C†PsCTjPSPsPiPs PePsP»*CHbPsCTjCr PrP*»*C‡ PrCTjCrPeCr*: *P·P°P*»*C*–*P·P° PsPeCl'PëPr*
C†PsCTjPSPëPNë, *P*• 172; *C€PμP*»*P°Pe*; *PμC,P°PSPsP*»; *PIPsPrP°*;
PiCTjPsPiC–*P*»*PμPSPiP*»*C*–*PePsP*»*CH*; *C*–*P·PsPiCTjPsPiP°PSPsP*»; *P±CrC,P°PSPsP*»; *P°PjC*–*P°Pe*
PIPsPrPSPëPNë; *PeP°P*»*C*–*CT PiC*–*PrCTjPsPeCl'PëPr*.

РљР°РјРґРґР»Рґ С,РІРґРґРґ- РіРґ 4,5 РјРј

$P''C-CHC\equiv P^{\circ}CH_2P\mu C\equiv PsPIP\ddot{e}PSP^{\circ}:PeP^{\circ}CHPePiCHP^{\circ}P\cdot PePS;$

1 PeP°PiCΓCíP»P° PjC-CΓC,PeC,CH PeP°CΓPëPiCΓP°P·PëPSCí 4,5 PjPi (Cí PIPëPiP»CΠPrC-PeP°CΓPëPiCΓP°P·PëPSCí PiC-PrCΓPsC...P»PsCΓPëPrCí 4,905 PjPi).

*P''PsPiPsPjC-P\PSC- CḡPμC#PsPIPëPSPë: PeCḡPsC...PjP°P»CH PeCíPeCíCḡCíPrP·CḡPSPëPNë
PiCḡPμPḡPμP»P°C,PëPSC-P·PsPIP°PSPëPNë, PjP°PiPSC-Cḡ CíC,PμP°CḡP°C,.*

*PḡP°PiCíCáP»Pë C,PIPμCḡPrC-: C,PëC,P°PSCí PrC-PsPeCíPëPr, P• 171; PḡPμP»P°C,PëPS;
CíPiPμC†C-P°P»CHPSPëPNë C#PμCḡPIPSPSPëPNë PhPŸ, P• 129; PrC-P°PjP°PSC,PsPIPëPNë
CíPëPSC-PNë FCF, P• 133; P·P°P»C-P·P° PsPeCíPëPr PḡPsPIC,PëPNë, P• 172; C#PsCḡPSPëP»Ps
P±C-P»PsPiPs PePsP»CHPsCḡCí PrP»Cḡ PrCḡCíPeCí: PiP»P°P·CíCḡCH
C,,P°CḡPjP°C†PμPIC,PëC#PSP° (CëPμP»P°PeCí CḡPsP·C#PëPS PI PμC,P°PSPsP»C-); C,PëC,P°PSCí
PrC-PsPeCíPëPr, P• 171; C-P·PsPiCḡPsPiP°PSPsP»; P±CíC,P°PSPsP»; P°PjC-P°Pe PIPsPrPSPëPNë;
PiCḡPsPiC-P»PμPSPiP»C-PePsP»CH; CíPëPjPμC,PëPePsPS.*

PḡP°PiCíCáP»Pë C,PIPμCḡPrC- PiPs 6 PjPi

P''C-CḡC#P° CḡPμC#PsPIPëPSP°: PeP°CḡPëPiCḡP°P·PëPS;

*1 PeP°PiCíCáP»P° PjC-CíC,PëC,CH PeP°CḡPëPiCḡP°P·PëPSCí 6 PjPi (Cí PIPëPiP»CḡPrC-
PeP°CḡPëPiCḡP°P·PëPSCí PiC-PrCḡPsC...P»PsCḡPëPrCí 6,540 PjPi).*

*P''PsPiPsPjC-P\PSC- CḡPμC#PsPIPëPSPë: PeCḡPsC...PjP°P»CH PeCíPeCíCḡCíPrP·CḡPSPëPNë
PiCḡPμPḡPμP»P°C,PëPSC-P·PsPIP°PSPëPNë, PjP°PiPSC-Cḡ CíC,PμP°CḡP°C,.*

*PḡP°PiCíCáP»Pë C,PIPμCḡPrC-: C,PëC,P°PSCí PrC-PsPeCíPëPr, P• 171; PḡPμP»P°C,PëPS;
CíPiPμC†C-P°P»CHPSPëPNë C#PμCḡPIPSPSPëPNë PhPŸ, P• 129; PrC-P°PjP°PSC,PsPIPëPNë
CíPëPSC-PNë FCF, P• 133; C#PsCḡPSPëP»Ps C#PsCḡPSPsPiPs PePsP»CHPsCḡCí PrP»Cḡ
PrCḡCíPeCí: P·P°P»C-P·P° PsPeCíPëPr C#PsCḡPSPëPNë, P• 172; CëPμP»P°Pe; PμC,P°PSPsP»;
PIPsPrP°; PiCḡPsPiC-P»PμPSPiP»C-PePsP»CH; C-P·PsPiCḡPsPiP°PSPsP»; P±CíC,P°PSPsP»;
P°PjC-P°Pe PIPsPrPSPëPNë; PeP°P»C-Cḡ PiC-PrCḡPsPeCíPëPr.*

P·C-PeP°CḡCíCHPeP° C,,PsCḡPjP°. PḡP°PiCíCáP»Pë C,PIPμCḡPrC-.

PhCíPSPsPIPSC- C,,C-P·PëPePs-C...C-PjC-C#PSC- PIP»P°CíC,PëPIPsCíC,C-:

*PḡP°PiCíCáP»Pë PiPs 1,5 PjPi. PŸPIPμCḡPrC- PḡPμP»P°C,PëPSPsPIC- PeP°PiCíCáP»Pë,
CḡPsP·PjC-Cḡ B,,- 4. PḡCḡPëCëPμC#PeP° PeP°PiCíCáP»Pë Bḡ" P±C-P»P° PSPμPiCḡPsP·PsCḡP°,
PePsCḡPiCíCí PeP°PiCíCáP»Pë Bḡ" P±C-P»PëPNë PSPμPiCḡPsP·PsCḡPëPNë. PkP° PePsCḡPiCíCí
PeP°PiCíCáP»Pë PSP°PSPμCíPμPSPs PSP°PiPëCí B«GR 1.5B» C#PsCḡPSPëPj PePsP»CHPsCḡPsPj.
P'PjC-CíC, PeP°PiCíCáP» Bḡ" PiPsCḡPsCëPsPe P±C-P»PsPiPs P°P±Ps PḡPsPIC,CíPIP°C,Ps-
P±C-P»PsPiPs PePsP»CHPsCḡCí.*

*PḡP°PiCíCáP»Pë PiPs 3 PjPi. PŸPIPμCḡPrC- PḡPμP»P°C,PëPSPsPIC- PeP°PiCíCáP»Pë,
CḡPsP·PjC-Cḡ B,,- 4. PḡCḡPëCëPμC#PeP° PeP°PiCíCáP»Pë Bḡ" P·PμP»PμPSP°
PSPμPiCḡPsP·PsCḡP°, PePsCḡPiCíCí PeP°PiCíCáP»Pë Bḡ" P±C-P»PëPNë
PSPμPiCḡPsP·PsCḡPëPNë. PkP° PePsCḡPiCíCí PeP°PiCíCáP»Pë PSP°PSPμCíPμPSPs PSP°PiPëCí
B«GR 3B» C#PsCḡPSPëPj PePsP»CHPsCḡPsPj. P'PjC-CíC, PeP°PiCíCáP» Bḡ" PiPsCḡPsCëPsPe
P±C-P»PsPiPs P°P±Ps PḡPsPIC,CíPIP°C,Ps-P±C-P»PsPiPs PePsP»CHPsCḡCí.*

*PḡP°PiCíCáP»Pë PiPs 4,5 PjPi. PŸPIPμCḡPrC- PḡPμP»P°C,PëPSPsPIC- PeP°PiCíCáP»Pë,
CḡPsP·PjC-Cḡ B,,- 4. PḡCḡPëCëPμC#PeP° PeP°PiCíCáP»Pë Bḡ" P·PμP»PμPSP°
PSPμPiCḡPsP·PsCḡP°, PePsCḡPiCíCí PeP°PiCíCáP»Pë Bḡ" P·PμP»PμPSPëPNë
PSPμPiCḡPsP·PsCḡPëPNë. PkP° PePsCḡPiCíCí PeP°PiCíCáP»Pë PSP°PSPμCíPμPSPs PSP°PiPëCí
B«GR 4.5B» P±C-P»PëPj PePsP»CHPsCḡPsPj. P'PjC-CíC, PeP°PiCíCáP» Bḡ" PiPsCḡPsCëPsPe*

$P \pm C - P \gg PsPiPs$ $P^\circ P \pm Ps$ $P \parallel PsPIC, CrPIP^\circ C, Ps - P \pm C - P \gg PsPiPs$ $PePsP \gg CHPsC\bar{H}C\bar{r}$.

$P_{\bar{b}}P^\circ PiC\bar{r}C\bar{z}P \gg P\bar{e} PiPs$ 6 $PjPi$. $P\bar{y}PIP_{\mu}C\bar{H}PrC - P \parallel P_{\mu}P \gg P^\circ C, P\bar{e}PSPsPIC - PeP^\circ PiC\bar{r}C\bar{r}P \gg P\bar{e}$, $C\bar{H}PsP \cdot PjC - C\bar{H}$ $\bar{b}_{\bar{r}}$ - 3. $P_{\bar{b}}C\bar{H}P\bar{e}C\bar{e}P_{\mu}C\bar{z}PeP^\circ$ $PeP^\circ PiC\bar{r}C\bar{r}P \gg P\bar{e}$ $\bar{b}\bar{H}$ " $C_{\bar{r}}, C - PsP \gg P_{\mu}C, PsPIP^\circ$ $PSP_{\mu}PiC\bar{H}PsP \cdot PsC\bar{H}P^\circ$, $PePsC\bar{H}PiC\bar{r}C\bar{r}$ $PeP^\circ PiC\bar{r}C\bar{r}P \gg P\bar{e}$ $\bar{b}\bar{H}$ " $P \pm C - P \gg P\bar{e}PN_{\bar{e}}$ $PSP_{\mu}PiC\bar{H}PsP \cdot PsC\bar{H}P\bar{e}PN_{\bar{e}}$. $P\bar{k}P^\circ$ $PePsC\bar{H}PiC\bar{r}C\bar{r}$ $PeP^\circ PiC\bar{r}C\bar{r}P \gg P\bar{e}$ $PSP^\circ PSP_{\mu}C\bar{r}P_{\mu}PSPs$ $PSP^\circ PiP\bar{e}C\bar{r}$ $B \ll GR$ 6B» $C\bar{z}PsC\bar{H}PSP\bar{e}Pj$ $PePsP \gg CHPsC\bar{H}PsPj$. $P \cdot PjC - C\bar{r}C$, $PeP^\circ PiC\bar{r}C\bar{r}P \gg \bar{b}\bar{H}$ " $PiPsC\bar{H}PsC\bar{e}PsPe$ $P \pm C - P \gg PsPiPs$ $P^\circ P \pm Ps$ $P \parallel PsPIC, CrPIP^\circ C, Ps - P \pm C - P \gg PsPiPs$ $PePsP \gg CHPsC\bar{H}C\bar{r}$.

$P_{\bar{r}}P^\circ C\bar{H}PjP^\circ PePsC, P_{\mu}C\bar{H}P^\circ PiP_{\mu}PIC, P\bar{e}C\bar{z}PSP^\circ$ $PiC\bar{H}C\bar{r}PiP^\circ$. $P_{\bar{r}}C\bar{r}P\bar{e}C...PsP \gg P_{\mu}PiC, P\bar{e}C\bar{z}PSC - P \cdot P^\circ C\bar{r}PsP \pm P\bar{e}$, $C - PSC\bar{e}C - P^\circ PSC, P\bar{e}PiC\bar{r}P\bar{e}C...PsC, P\bar{e}C\bar{z}PSC - P \cdot P^\circ C\bar{r}PsP \pm P\bar{e}$.

$P_{\bar{b}}PsPr$ $P\bar{h}P\bar{y}PI^\Gamma$ N05A X15.

$P_{\bar{r}}P^\circ C\bar{H}PjP^\circ PePsP \gg PsPiC - C\bar{z}PSC - PIP \gg P^\circ C\bar{r}C, P\bar{e}PIPsC\bar{r}C, C -$

$P_{\bar{r}}P^\circ C\bar{H}PjP^\circ PePsPrP\bar{e}PSP^\circ PjC - PeP^\circ$.

$P_{\bar{b}}P_{\mu}C...P^\circ PSC - P \cdot Pj$ $PrC - C -$

$P_{\bar{b}}P_{\mu}C...P^\circ PSC - P \cdot Pj$ $PrC - C -$ $PeP^\circ C\bar{H}P\bar{e}PiC\bar{H}P^\circ P \cdot P\bar{e}PSC\bar{r}$ $C\%_{\bar{o}}P_{\mu}$ PSP_{μ} $PiPsPIPSC - C\bar{r}C, C\bar{H}$ $PIC - PrPsPjP\bar{e}PN_{\bar{e}}$. $P_{\bar{r}}C\bar{H}PsC, P_{\mu}$ $C, P_{\mu}C\bar{H}P^\circ PiP_{\mu}PIC, P\bar{e}C\bar{z}PSP\bar{e}PN_{\bar{e}}$ $P_{\mu}C_{\bar{r}}, P_{\mu}PeC$, $PeP^\circ C\bar{H}P\bar{e}PiC\bar{H}P^\circ P \cdot P\bar{e}PSC\bar{r}$ $PjPsP \parallel P_{\mu}$ $PsPiPsC\bar{r}P_{\mu}C\bar{H}P_{\mu}PrPePsPIC\bar{r}PIP^\circ C, P\bar{e}C\bar{r}C\bar{r}$ $C\bar{z}P_{\mu}C\bar{H}P_{\mu}P \cdot$ $PePsPjP \pm C - PSP^\circ C\bar{r}C - C\bar{H}$ $PrC - C -$ $C\bar{z}P^\circ C\bar{r}C, PePsPIPsPiPs$ $P^\circ PiPsPSC - C\bar{r}C, P^\circ$ PSP° $D_3, D_2 -$ $PrPsC_{\bar{r}}, P^\circ PjC - PSPsPIC - (P \cdot PSP^\circ C\bar{z}P_{\mu}PSPSC\bar{r}$ Ki $PIC - PrPiPsPIC - PrPSPs$ PI $PjP_{\mu}P \parallel P^\circ C...$ $0,085\bar{b}\bar{H}$ " $0,3$ $PSP_{\bar{b}}$ $PiC\bar{H}PsC, P\bar{e}$ $0,49\bar{b}\bar{H}$ " $0,71$ $PSP_{\bar{b}}$) C, P° $5-HT_{1A} - C\bar{r}P_{\mu}C\bar{H}PsC, PsPSC - PSPsPIC -$ $C\bar{H}P_{\mu}C\bar{r}P_{\mu}PiC, PsC\bar{H}P\bar{e}$ $(P \cdot PSP^\circ C\bar{z}P_{\mu}PSPSC\bar{r}$ Ki PI $PjP_{\mu}P \parallel P^\circ C...$ $1,4\bar{b}\bar{H}$ " $2,6$ $PSP_{\bar{b}}$) $C - PrC - C -$ $P^\circ PSC, P^\circ PiPsPSC - C\bar{r}C, P^\circ$ PSP° $5-HT_{2B}, 5-HT_{2A} - C\bar{r}P_{\mu}C\bar{H}PsC, PsPSC - PSPsPIC - C, P^\circ$ $H_1 -$ $PiC - C\bar{r}C, P^\circ PjC - PSPsPIC - C\bar{H}P_{\mu}C\bar{r}P_{\mu}PiC, PsC\bar{H}P\bar{e}$ $(P \cdot PSP^\circ C\bar{z}P_{\mu}PSPSC\bar{r}$ Ki $PIC - PrPiPsPIC - PrPSPs$ PI $PjP_{\mu}P \parallel P^\circ C...$ $0,58\bar{b}\bar{H}$ " $1,1$ $PSP_{\bar{b}}$, $18,8$ $PSP_{\bar{b}}$ C, P° $23,3$ $PSP_{\bar{b}}$). $P_{\bar{b}}P^\circ C\bar{H}P\bar{e}PiC\bar{H}P^\circ P \cdot P\bar{e}PS$ $PjP^\circ C$ " $PSP\bar{e}P \cdot CHPeC\bar{r}$ $P^\circ C_{\bar{r}}, C - PSPSC - C\bar{r}C, CH$ $C\%_{\bar{o}}PsPrPs$ $5-HT_{2C} - C\bar{r}P_{\mu}C\bar{H}PsC, PsPSC - PSPsPIP\bar{e}C...$ C, P° $\alpha_1 - P^\circ PrC\bar{H}P_{\mu}PSPsC\bar{H}P_{\mu}C\bar{r}P_{\mu}PiC, PsC\bar{H}C - PI$ $(P \cdot PSP^\circ C\bar{z}P_{\mu}PSPSC\bar{r}$ Ki $PIC - PrPiPsPIC - PrPSPs$ 134 $PSP_{\bar{b}}$ C, P° 155 $PSP_{\bar{b}}$). $P_{\bar{b}}P^\circ C\bar{H}P\bar{e}PiC\bar{H}P^\circ P \cdot P\bar{e}PS$ PSP_{μ} $PjP^\circ C$ " $P \cdot PSP^\circ C\bar{z}PSPsC -$ $P^\circ C_{\bar{r}}, C - PSPSPsC\bar{r}C, C - C\%_{\bar{o}}PsPrPs$ $C...PsP \gg C - PSP_{\mu}C\bar{H}PiC - C\bar{z}PSP\bar{e}C...$ $PjC\bar{r}C\bar{r}PeP^\circ C\bar{H}P\bar{e}PSPsPIP\bar{e}C...$ $C\bar{H}P_{\mu}C\bar{r}P_{\mu}PiC, PsC\bar{H}C - PI$ $(IC_{50} > 1000$ $PSP_{\bar{b}})$. $P"PIP^\circ$ $PsC\bar{r}PSPsPIPSP\bar{e}C...$ $P^\circ PeC, P\bar{e}PIPSP\bar{e}C...$ $PjP_{\mu}C, P^\circ P \pm PsP \gg C - C, P\bar{e}$ $\bar{b}\bar{H}$ " $PrP_{\mu}P \cdot PjP_{\mu}C, P\bar{e}P \gg - PeP^\circ C\bar{H}P\bar{e}PiC\bar{H}P^\circ P \cdot P\bar{e}PS$ C, P° $PrP\bar{e}PrP_{\mu}P \cdot PjP_{\mu}C, P\bar{e}P \gg - PeP^\circ C\bar{H}P\bar{e}PiC\bar{H}P^\circ P \cdot P\bar{e}PS$ $\bar{b}\bar{H}$ " $P \cdot PI\bar{b}\bar{H}^{\bar{r}m}$ $C\bar{r}P \cdot CrC\bar{H}C, CHC\bar{r}C\bar{r}$ $P \cdot$ $C\bar{H}P_{\mu}C\bar{r}P_{\mu}PiC, PsC\bar{H}P^\circ PjP\bar{e}$ C, P° $PjP^\circ C\bar{H}C, CH$ $PiC\bar{H}PsC_{\bar{r}}, C - P \gg CH$ $C_{\bar{r}}, CrPSPeC\bar{r}C - PsPSP^\circ P \gg CHPsPSC - P^\circ PeC, P\bar{e}PIPSPsC\bar{r}C, C - in vitro$, $P^\circ PSP^\circ P \gg PsPiC - C\bar{z}PSP\bar{e}PN_{\bar{e}}$ $PIP\bar{e}C...C - PrPSC - PN_{\bar{e}}$ $PrC - C\bar{H}C\bar{z}C - PN_{\bar{e}}$ $C\bar{H}P_{\mu}C\bar{z}PsPIP\bar{e}PSC -$.

$P_{\bar{r}}P^\circ C\bar{H}PjP^\circ PePsPrP\bar{e}PSP^\circ PjC - C\bar{z}PSC - P_{\mu}C_{\bar{r}}, P_{\mu}PeC, P\bar{e}$

$P"PsPeP \gg C - PSC - C\bar{z}PSC - PrPsC\bar{r}P \gg C - PrP \parallel P_{\mu}PSPSC\bar{r}$ $in vivo$ $PiPsPeP^\circ P \cdot CrC\bar{H}C, CH$, $C\%_{\bar{o}}Ps$ $PeP^\circ C\bar{H}P\bar{e}PiC\bar{H}P^\circ P \cdot P\bar{e}PS$ $P \cdot PI\bar{b}\bar{H}^{\bar{r}m}$ $C\bar{r}P \cdot CrC" C, CHC\bar{r}C\bar{r}$ $P \cdot D_3 - C\bar{H}P_{\mu}C\bar{r}P_{\mu}PiC, PsC\bar{H}P^\circ PjP\bar{e}$ $C, P^\circ PePsC\bar{H}$ $P \parallel PjC - C\bar{H}PsC\bar{H}$, $C\%_{\bar{o}}Ps$ $PN_{\bar{e}}$ $C - P \cdot D_2 - C\bar{H}P_{\mu}C\bar{r}P_{\mu}PiC, PsC\bar{H}P^\circ PjP\bar{e}$ Cr $C_{\bar{r}}, P^\circ C\bar{H}PjP^\circ PePsP \gg PsPiC - C\bar{z}PSPs$ $P_{\mu}C_{\bar{r}}, P_{\mu}PeC, P\bar{e}PIPSP\bar{e}C...$ $PrPsP \cdot P^\circ C...$ $P^\circ P\bar{e}C\bar{r}PIP \gg C\bar{r}P \gg PsC\bar{r}C\bar{r}$ $P \cdot P^\circ P \gg P_{\mu}P \parallel PSP_{\mu}$ $PIC - Pr$ $PrPsP \cdot P\bar{e}$ $P \cdot PI\bar{b}\bar{H}^{\bar{r}m}$ $C\bar{r}P \cdot CrPIP^\circ PSPSC\bar{r}$ $P \cdot D_3 - C, P^\circ$ $D_2 -$

PrPsC,,P°PjC-PSPsPIPëPjPë CḡPμC†PμPīC,PsCḡP°PjPë PiPsP»PsPIPSPsPiPs PjPsP·PeCḡ (P·
PīPμCḡPμPIP°P¶PSPëPj P·PIbḡ™CḡP·CḡPIP°PSPSCḡPj PI PrC-P»CḡPSPeP°C... P·
PīC-PrPIPëC%oPμPSPëPj CḡPëPSC,PμP·PsPj D3) Cḡ PīP°C†C-C”PSC,C-PI P·
CëPëP·PsC,,CḡPμPSC-C”Cḡ, CḡPeC- PīCḡPëPμPjP°P»Pë C,PμCḡP°PīPμPIC,PëC†PSC- PrPsP·Pë
PeP°CḡPëPīCḡP°P·PëPSCḡ PīCḡPsC,CḡPiPsPj 15 PrPSC-PI.

P·PīP»PëPI PeP°CḡPëPīCḡP°P·PëPSCḡ PSP° C-PSC,PμCḡPIP°P» QT P±CḡPI PsC†C-PSPμPSPëPμ
Cḡ PīP°C†C-C”PSC,C-PI C-P· CëPëP·PsC,,CḡPμPSC-C”Cḡ P°P±Ps CëPëP·PsP°C,,PμPeC,PëPIPSPëPj
CḡPsP·P»P°PrPsPj. P PμP·CḡP»CHbC,P°C,Pë C...PsP»C,PμCḡC-PICḡCHbPePsPiPs
PjPsPSC-C,PsCḡCḡPIP°PSPSCḡ B P·PḡP” B P±CḡP»Pë B P·C-P±CḡP°PSC- B Cḡ B 129
PīP°C†C-C”PSC,C-PI PīCḡPsC,CḡPiPsPj 12-PiPsPrPëPSPSPsPiPs PīPμCḡC-PsPrCḡ PSP°
PIPëC...C-PrPSPsPjCḡ CḡC-PIPSC- C,P° CḡC-PIPSPsPIP°P¶PSPsPjCḡ CḡC,P°PSC-. PJ CḡP°P·C-
PIPëPePsCḡPëCḡC,P°PSPSCḡ CḡCḡP±C,PμCḡP°PīPμPIC,PëC†PSPëC... PrPsP· (9 PjPi/PrPsP±Cḡ
P°P±Ps 18 PjPi/PrPsP±Cḡ) PSPμ P±CḡP»Ps PIC-PrPjC-C†PμPSPs PīPsPrPsPIP¶PμPSPSCḡ
C-PSC,PμCḡPIP°P»Cḡ QT. PJ P¶PsPrPSPsPiPs P· PīP°C†C-C”PSC,C-PI, CḡPeC-
P»C-PeCḡPIP°P»PëCḡCḡ PeP°CḡPëPīCḡP°P·PëPSPsPj, PSPμ PIC-PrPjC-C†P°P»PsCḡCḡ
PīPsPrPsPIP¶PμPSPSCḡ C-PSC,PμCḡPIP°P»Cḡ QT ≥ 60 PjCḡ PIC-Pr PIPëC...C-PrPSPsPiPs
CḡC-PIPSCḡ, C- P¶PsPrPμPS PīP°C†C-C”PSC, PSPμ PjP°PI PīPsPrPsPIP¶PμPSPSCḡ
C-PSC,PμCḡPIP°P»Cḡ QT > 500 PjCḡ PīC-Pr C†P°Cḡ PrPsCḡP»C-PrP¶PμPSPSCḡ.

PḡP»C-PSC-C†PSP° PμC,,PμPeC,PëPIPSC-CḡC,CHb C,P° P±PμP·PīPμPeP°

*P·C,,PμPeC,PëPIPSC-CḡC,CHb PīCḡPë PePsCḡPsC,PePsC†P°CḡPSPsPjCḡ
P·P°CḡC,PsCḡCḡPIP°PSPSC-*

P·C,,PμPeC,PëPIPSC-CḡC,CHb PeP°CḡPëPīCḡP°P·PëPSCḡ PI P»C-PeCḡPIP°PSPSC-
PiPsCḡC,CḡPsC— CëPëP·PsC,,CḡPμPSC-C— P±CḡP»P° PIPëPIC†PμPSP° PI C,CḡCHbPsC...
P±P°PiP°C,PsC†PμPSC,CḡPsPIPëC...B B B B PjC-P¶PSP°CḡPsPrPSPëC...B B
B CḡP°PSPrPsPjC-P·PsPIP°PSPëC...B B PīPsPrPIC-PμPSPsB B B CḡP»C-PīPëC...
B B B PīP»P°C†PμP±PsPePsPSC,CḡPsP»CHbPsPIP°PSPëC...B B B 6-C,PëP¶PSPμPIPëC...
PrPsCḡP»C-PrP¶PμPSPSCḡC..., PI CḡPePëC... P±CḡP°P»Pë CḡC†P°CḡC,CHb 1 754
PīP°C†C-C”PSC,Pë PIC-PePsPj PIC-Pr 18 PrPs 60 CḡPsPeC-PI. PJ PICḡC-C...
PrPsCḡP»C-PrP¶PμPSPSCḡC... PiPsCḡC,CḡPsC— CëPëP·PsC,,CḡPμPSC-C—
PīPμCḡPIPëPSPSPsCḡ PeC-PSC†PμPIPsCḡ C,PsC†PePsCḡ P±CḡP»P° P·PjC-PSP° PIC-Pr
PIPëC...C-PrPSPsPiPs CḡC-PIPSCḡ PrPs 6 C,PëP¶PSCḡ P·P° P·P°PiP°P»CHbPSPëPjPë
PīPsPeP°P·PSPëPeP°PjPë CëPeP°P»Pë PsC†C-PSPePë PīPsP·PëC,PëPIPSPëC... C,P°
PSPμPiP°C,PëPIPSPëC... CḡPëPjPīC,PsPjC-PI (PANSS), P° PIC,PsCḡPëPSPSPsCḡ
PeC-PSC†PμPIPsCḡ C,PsC†PePsCḡ bḡ” P·PjC-PSP° PIC-Pr PIPëC...C-PrPSPsPiPs CḡC-PIPSCḡ
PrPs 6 C,PëP¶PSCḡ P·P° PīPsPeP°P·PSPëPeP°PjPë CëPeP°P»Pë P·P°PiP°P»CHbPSPsPiPs
PeP»C-PSC-C†PSPsPiPs PICḡP°P¶PμPSPSCḡ PīCḡPs C,CḡP¶PeC-CḡC,CHb
P·P°C...PIPsCḡCHbPIP°PSPSCḡ (CGI-S). PJ PjC-P¶PSP°CḡPsPrPSPsPjCḡ
PīP»P°C†PμP±PsPePsPSC,CḡPsP»CHbPsPIP°PSPsPjCḡ PrPsCḡP»C-PrP¶PμPSPSC-, PI CḡPePsPjCḡ
PrP»Cḡ C†CḡC,P»PëPIPsCḡC,C- P°PSP°P»C-P·Cḡ PIPëPePsCḡPëCḡC,PsPICḡPIP°P»PëCḡCḡ
C,,C-PeCḡPsPIP°PSC- PrPsP·Pë 1,5 PjPi, 3,0 PjPi C,P° 4,5 PjPi PeP°CḡPëPīCḡP°P·PëPSCḡ C- 4,0
PjPi CḡPëCḡPīPμCḡPëPrPsPSCḡ, PICḡC- PrPsP·Pë PeP°CḡPëPīCḡP°P·PëPSCḡ C,P°
P°PeC,PëPIPSPëPμ PePsPSC,CḡPsP»CHb PīPsPeP°P·P°P»Pë CḡC,P°C,PëCḡC,PëC†PSPs
PrPsCḡC,PsPIC-CḡPSPμ PīPsPeCḡP°C%oPμPSPSCḡ CḡPe PīPμCḡPIPëPSPSPsC—, C,P°Pe C-
PIC,PsCḡPëPSPSPsC— PeC-PSC†PμPIPëC... C,PsC†PsPe Cḡ PīPsCḡC-PIPSCḡPSPSC- P·
PīP»P°C†PμP±Ps. PJ PjC-P¶PSP°CḡPsPrPSPsPjCḡ
PīP»P°C†PμP±PsPePsPSC,CḡPsP»CHbPsPIP°PSPsPjCḡ PrPsCḡP»C-PrP¶PμPSPSC-, PI CḡPePsPjCḡ
PrP»Cḡ C†CḡC,P»PëPIPsCḡC,C- P°PSP°P»C-P·Cḡ PIPëPePsCḡPëCḡC,PsPICḡPIP°P»PëCḡCḡ
C,,C-PeCḡPsPIP°PSC- PrPsP·Pë 3,0 PjPi C,P° 6,0 PjPi PeP°CḡPëPīCḡP°P·PëPSCḡ C- 10 PjPi

P°CTHPēPiC-PiCTHP°P·PsP»Cf, CŪPe PrPsP·Pē PeP°CTHPēPiCTHP°P·PēPSCf, C,P°Pe C-
P°PeC,PēPIPSPēPNē PePsPSC,CTHPsP»CH PiPsPeP°P·P°P»Pē CfC,P°C,PēCfC,PēC†PSPs
PrPsCfC,PsPIC-CTHPSPμ PiPsPeCTHP°C%oPμPSPSCŪ PiPμCTHPiPēPSPSPSc— C-
PIC,PsCTHPēPSPSPSc— PeC-PSC†PμPiPēC... C,PsC†PsPe Cf PiPsCTHC-PIPSCLIPSPSC- P·
PiP»P°C†PμP±Ps. PJ C,CTHPμC,CHPsPjCf PjC-P¶PSP°CTHPsPrPSPsPjCf
PiP»P°C†PμP±PsPePsPSC,CTHPsP»CHPsPiP°PSPsPjCf PrPsCfP»C-PrP¶PμPSPSC-, PI CŪPePsPjCf
PrP»CŪ C†CfC,P»PēPIPSCfC,C- P°PSP°P»C-P·Cf PiPēPePsCTHPēCfC,PsPICfPiP°P»PēCfCŪ
C,,C-PeCfPsPiP°PSC-/P°PrP°PiC,PsPiP°PSC- PrPsP·Pē 3,0BĤ"6,0 PjPi C,P° 6,0BĤ"9,0 PjPi
PeP°CTHPēPiCTHP°P·PēPSCf, PsP±PēPrPIC- PiCTHCfPiPē PeP°CTHPēPiCTHP°P·PēPSCf
PiPsPeP°P·P°P»Pē CfC,P°C,PēCfC,PēC†PSPs PrPsCfC,PsPIC-CTHPSPμ PiPsPeCTHP°C%oPμPSPSCŪ
CŪPe PiPμCTHPiPēPSPSPSc—, C,P°Pe C- PIC,PsCTHPēPSPSPSc— PeC-PSC†PμPiPēC... C,PsC†PsPe
Cf PiPsCTHC-PIPSCLIPSPSC- P· PiP»P°C†PμP±Ps.

P PμP·CfP»CHC,P°C,Pē PiPēP·PSP°C†PμPSPSCŪ PiPμCTHPiPēPSPSPSc— PeC-PSC†PμPiPSc—
C,PsC†PePē PμC,,PμPeC,PēPIPSPsCfC,C- PSP°PiPμPrPμPSC- PI C,P°P±P»PēC†C- 1 PSPēP¶C†Pμ.
P PμP·CfP»CHC,P°C,Pē PiPēP·PSP°C†PμPSPSCŪ PIC,PsCTHPēPSPSPSc— PeC-PSC†PμPiPSc—
C,PsC†PePē PμC,,PμPeC,PēPIPSPsCfC,C- (CGI) C,P° PrPsPrP°C,PePsPiPēC... PeC-PSC†PμPiPēC...
C,PsC†PsPe PiC-PrC,PIPμCTHPPrP¶CfPiP°P»Pē PiPμCTHPiPēPSPSCf PeC-PSC†PμPICf C,PsC†PeCf.

PŷP°P±P»PēC†CŪ 1.B B B B P—PjC-PSPē PIC-Pr PiPēC...C-PrPSPsPiPs CTHC-PIPSCLIP PrPs 6
C,PēP¶PSCŪ P·P° P·P°PiP°P»CHPSPēPj PiPsPeP°P·PSPēPePsPj PANSS Cf
PrPsCfP»C-PrP¶PμPSPSCŪC... P·P°PiPsCfC,CTHPμPSPSCŪ C€PēP·PsC,,CTHPμPSC-C— BĤ"
PiCTHCfPiP° ITT.

<i>P'PēC...C-PrPSPēPNē</i>	<i>P—PjC-PSP°</i>	<i>P C-P·PSPēC†CŪ PI</i>	<i>P -</i>
<i>CTHC-PIPμPSCĤ</i>	<i>PŷPμCTHPμPrPSC"</i>	<i>P»C-PeCžPiP°PSPSC-</i>	<i>P·PSP°C†PμPSPS</i>
<i>PŷPμCTHPμPrPSC"</i>	<i>P·PSP°C†PμPSPSCŪ</i>	<i>PI</i>	<i>CŪ</i>
<i>P·PSP°C†PμPSPSCŪ</i>	<i>LS (PŷPμ)</i>	<i>PiPsCTHC-PIPSCLIPSPS</i>	
<i>B± PŷP'</i>		<i>C- P· PiP»P°C†PμP±Ps</i>	
		<i>(95 % P"P†)</i>	

P—P°PiP°P»CHPSPēPNē PiPsPeP°P·PSPēPe PANSS (MMRM)

P"PsCfP»C-PrP¶PμPSPSCŪ RGH-MD-16 (n = 711)

PμP»P°C†PμP±Ps	97,3 B± 9,22	BĤ"13,29 (1,82)	BĤ"	BĤ"
PľP°CTHPēPiCTHP°P·Pē	97,1 B± 9,13	BĤ"21,27 (1,77)	BĤ"7,97 (BĤ"12,94, BĤ"3,01)	0,0017
PS 1,5 PjPi/PrPsP±Cf			BĤ"8,16 (BĤ"13,09, BĤ"3,22)	0,0013
PľP°CTHPēPiCTHP°P·Pē	97,2 B± 8,66	BĤ"21,45 (1,74)	BĤ"10,48 (BĤ"15,41, BĤ"5,55)	< 0,0001
PS 3 PjPi/PrPsP±Cf	96,7 B± 9,01	BĤ"23,77 (1,74)	BĤ"15,98 (BĤ"20,91, BĤ"11,04)	< 0,0001*
P PēCfPiPμCTHPēPrPsP98,1 B± 9,50		BĤ"29,27 (1,74)		
S 4 PjPi/PrPsP±Cf				

P"PsCfP»C-PrP¶PμPSPSCŪ RGH-MD-04 (n = 604)

PμP»P°C†PμP±Ps	96,5 B± 9,1	BĤ"14,3 (1,5)	BĤ"	BĤ"
PľP°CTHPēPiCTHP°P·Pē	96,1 B± 8,7	BĤ"20,2 (1,5)	BĤ"6,0 (BĤ"10,1, BĤ"1,9)	0,0044
PS 3 PjPi/PrPsP±Cf			BĤ"8,8 (BĤ"12,9, BĤ"4,7)	< 0,0001
PľP°CTHPēPiCTHP°P·Pē	95,7 B± 9,4	BĤ"23,0 (1,5)	BĤ"7,0 (BĤ"11,0, BĤ"2,9)	0,0008*
PS 6 PjPi/PrPsP±Cf	95,6 B± 9,0	BĤ"21,2 (1,4)		
PĥCTHPēPiC-PiCTHP°P·P	95,6 B± 9,0			
sP» 10 PjPi/PrPsP±Cf				

P"PsCfP»C-PrP¶PμPSPSCŪ RGH-MD-05 (n = 439)

PμP»P°C†PμP±Ps	96,6 B± 9,3	BĤ"16,0 (1,6)	BĤ"	BĤ"
PľP°CTHPēPiCTHP°P·Pē	96,3 B± 9,3	BĤ"22,8 (1,6)	BĤ"6,8 (BĤ"11,3, BĤ"2,4)	0,0029
PS				
3BĤ"6 PjPi/PrPsP±Cf				

РљР°СђРёРїСђР°Р·Рё				
PS	96,3 B± 9,0	вђ“25,9 (1,7)	вђ“9,9 (вђ“14,5, вђ“5,3)	< 0,0001
6вђ“9 РјРї/РrРsР±Сђ				

Р”Р† вђ“ РrРsРІС-СђСђРёР№ С-РSC,РµСђРІР°Р»; ІТТ вђ“ Р°РSP°Р»С-Р· Р·Р°Р
РїСђРёР·РSP°СђРµРSPёРј Р»С-РёСђРІР°РSPSCІРј; СђРµСђРІР°РSPSC” Р·РSP°СђРµРSPSCІ LS вђ“
СђРµСђРІР°РSPSC” РSP°Р№РјРµРSCёРёС... РёРІР°РrСђР°С,С-РІ; РANSS вђ“ СёРёР°Р»Р°Р°Р°
РsС†С-РSPёРё РїРsР·РёС,РёРІРSPёС... С,Р° РSPµРІР°С,РёРІРSPёС... СђРёРSPrСђРsРјС-РІ; РЎРµ
вђ“ СђС,Р°РSPrР°СђС,РSP° РїРsС...РёР±РёР°Р°; РЎР’ вђ“ СђС,Р°РSPrР°СђС,РSPµ
РІС-РrС...РёР»РµРSPSCІ; МMRM вђ“ РјРsРrРµР»Сђ Р·РјС-СёР°РSPёС... РµС,РµРёС,С-РІ
РrР»СІ Р±Р°Р°РІР°С,РsСђР°Р·РsРІРёС... РІРёРјС-СђСђРІР°РSPСђ.

* Рј РїРsСђС-РІРSCІРSPSC- Р· РїР»Р°СђРµРs.

*Р·С,РµРёС,РёРІРSC-СђС,Сђ РїСђРё РrРsРІРїРsС,СђРёРІР°Р»РsРјСђ
Р·Р°СђС,РsСђСђРІР°РSPSC-*

Р·С,РµРёС,РёРІРSC-СђС,Сђ РёР°СђРёРІР°Р»Р·Р·Р»Р» РrР»СІ РїС-РrС,СђРёРјР°РSPSCІ
Р°РSC,РёРїСђРёС...РsС,РёСђРSPsРїРs РµС,РµРёС,Сђ РrРsСђР»С-РrР»СђРІР°Р»Р°Р°Р°Р°Сђ Сђ
РrРsРІРїРsС,СђРёРІР°Р»Р»РsРјСђ Р РёР»С-РSC-СђРSPsРјСђ Р РrРsСђР»С-РrР»РµРSPSC- Р·
РјРsР»Р»Р»Р»Р»Р»Р»Р»Р»Р» Р СђС,Сђ Р»Р°РSPrРsРјС-Р·РsРІР°РSPsС— РІС-РrРјС-РSPё.
Р Р—Р°РІР°Р»Р»Р» Р 751 РїР°С†С-С”РSC, С-Р· РїРsСђС,СђРёРјРё СђРёРјРїС,РsРјР°РјРё
СёРёР·С,СђРµРSPSC— РsС,СђРёРјСђРІР°РІР° РёР°СђРёРІР°Р»Р·Р·Р»Р» Сђ РrРsР·С-
3вђ“9 РјРї/РrРsР±Сђ РІРїСђРsРrРsРІР» 20 С,РёР»РSPSC-РІ, Р· СІРёРёС... 337 РsС,СђРёРјР»Р»Р»Р»Р»Р»
РёР°СђРёРІР°Р»Р·Р·Р»Р» РІ РrС-Р°РІР°Р·РsРSC- РrРsР· РІС-Рr 3 РrРs 6 РјРї/РrРsР±Сђ.
РќР°РrР°Р»С- СђС,Р°Р±С-Р»С-Р·РsРІР°РSPSC- РїР°С†С-С”РSC,Рё Р±СђР»Р»Р»
СђР°РSPrРsРјС-Р·РsРІР°РSPSC- РїРsРrРІС-Р№РSPs СђР»С-РїРёРј РјРµС,РsРrРsРј РrР»СІ
РsС,СђРёРјР»Р»Р»Р»Р»Р» С,С-РёСђРsРІР°РSPёС... РrРsР· РёР°СђРёРІР°Р»Р·Р·Р»Р» РІС-Рr 3 РrРs
6 РјРї (n = 51) Р°Р±Рs РїР»Р°СђРµРs (n = 51) РїСђРsС,СІРїРsРј 72 С,РёР»РSPSC-РІ.
РµРµСђРІРёРSPSPёРј СђРµР·Р»С,Р°С,РsРј РrРsСђР»С-РrР»РµРSPSCІ Р±СђРІ СђР°Сђ
РrРs СђРsР·РІРёС,РёСђ СђРµСђРёРrРёРІСђ. Рј РёС-РSC†С- РrРsСђР»С-РrР»РµРSPSCІ 49,0 %
РїР°С†С-С”РSC,С-РІ, СІРёС- РsС,СђРёРјСђРІР°РІР°Р»Р»Р» РїР»Р°СђРµРs, РїСђРsС,Рё 21,6 %
РїР°С†С-С”РSC,С-РІ, СІРёС- РsС,СђРёРјСђРІР°РІР°Р»Р»Р» РёР°СђРёРІР°Р»Р·Р·Р»Р» РsР, РјР°Р»Р»Р»
СђРµСђРёРrРёРІ СђРёРјРїС,РsРјС-РІ СёРёР·С,СђРµРSPSC— РёС,Р»РµС,СђР°Сђ РrРs
СђРsР·РІРёС,РёСђ СђРµСђРёРrРёРІСђ (92 РїСђРsС,Рё 326 РrРSC-РІ РSP° РsСђРSPsРІС- 25-РїРs
РїРµСђРµРs,РёР»Сђ) Р±СђРІ Р·РSP°СђРSPs РrРsРІСёРёРј Сђ РїСђСђРїРё, СІРёР°Р°
РїСђРёР№РјР»Р»Р»Р»Р» РёР°СђРёРІР°Р»Р·Р·Р»Р» РsС,РSC-Р» Сђ РїСђСђРїРё, С%Рs
РsС,СђРёРјСђРІР°РІР°Р»Р» РїР»Р°СђРµРs (Сђ = 0,009).

*Р·С,РµРёС,РёРІРSC-СђС,Сђ РїСђРё РїРµСђРµРІР°РІР°Р»РSPs РSPµРІР°С,РёРІРSPёС...
СђРёРјРїС,РsРјР°Р° СёРёР·С,СђРµРSPSC—*

Р·С,РµРёС,РёРІРSC-СђС,Сђ РёР°СђРёРІР°Р»Р·Р·Р»Р» РІ Р»С-РёСђРІР°РSPSC-
РїРµСђРµРІР°Р»Р»Р» РSPµРІР°С,РёРІРSPёС... СђРёРмРїС,РsРјС-РІ СёРёР·С,СђРµРSPSC—
РrРsСђР»С-РrР»РµРSPSCІ РІ 26-С,РёР»РSPµРІРsРјСђ
Р±Р°Р°РІР°С,РsС†РµРSC,СђРsРІРsРјСђ РїРsРrРІС-Р№РSPs СђР»С-РїРsРјСђ РёР»С-РSC-СђРSPsРјСђ
РrРsСђР»С-РrР»РµРSPSC- Р· РёС,РёРІРSPёРј РёРsРSC,СђРsР»РµРµРј РљР°СђРёРІР°Р»Р·Р·Р»Р»Р»Р»Р»
(РrС-Р°РІР°Р·РsР РrРsР· 3вђ“4 РјРї, С†С-Р»СђРsРІР° РrРsР·Р° 4,5 РјРї)
РrРsСђР»С-РrР»РµРSPSCІ Сђ РїРsСђС-РІРSCІРSPSC- Р· СђРёСђРїРµРrРsРSPsРј
(РrС-Р°РІР°Р·РsР РrРsР· 3вђ“6 РјРї, С†С-Р»СђРsРІР° РrРsР·Р° 4 РјРї) Сђ РїР°СђС-С”РSC,С-РІ
С-Р· РїРµСђРёСђС,СђРІР°Р»Р»СђРSPёРј РїРµСђРµРІР°РІР°Р»Р»Р» РSPµРІР°С,РёРІРSPёРјР»Р»

PyP°P±P»PëC†CЦ 2.B B B B PцC-PrCГCрPjPsPe CГPμP·CрP»CHC,P°C,C-PI
PrPsCГP»C-PrP¶PμPSPSCЦ RGH-188-005

$P_{\mu}C-C\Gamma P \gg C\Gamma P \pm P^{\circ}P_iP^{\circ}C, PsPeC\Gamma P^{\circ}C, PSP\ddot{e}C \dots PrPsP \cdot P_jP^{\circ}PeC\Gamma P\ddot{e}P_jP^{\circ}P \gg CH\dot{P}SP^{\circ}$
 $PePsPSC\dot{+}P_{\mu}PSC, C\Gamma P^{\circ}C\dot{+}C-C\Gamma P \quad PeP^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PSC\dot{r} \quad C, P^{\circ} \quad P\ddot{N}ePsP_iPs \quad PsC\Gamma PSPsPIPSP\ddot{e}C \dots$
 $P^{\circ}PeC, P\ddot{e}PIPSP\ddot{e}C \dots P_jP_{\mu}C, P^{\circ}P \pm PsP \gg C-C, C-PI \quad C\dot{r} \quad B \quad P_iP \gg P^{\circ}P \cdot P_jC- \quad B \quad PeC\Gamma PsPIC-$
 $B \quad P \cdot P^{\circ}P \cdot PIP\ddot{e}C\dot{+}P^{\circ}P\ddot{N}e \quad B \quad PrPsC\Gamma C\Gamma P_iP^{\circ}C^{\prime\prime}C, CH\dot{C}\Gamma C\Gamma P \quad B \quad P_iC\Gamma P\ddot{e}P \pm P \gg P\ddot{e}P \cdot PSPs \quad B \quad C\dot{+}P_{\mu}C\Gamma P_{\mu}P \cdot$
 $B \quad 3B\dot{H}^{\circ}8 \quad P_iPsPrP\ddot{e}PS \quad P_iC-C\Gamma P \gg C\Gamma P \quad P \cdot P^{\circ}C\Gamma C, PsC\Gamma C\dot{r}PIP^{\circ}PSPSC\Gamma P.$

$P_{\mu}C\Gamma P\ddot{e}P\ddot{N}ePsP_j \quad PsPrPSC-C^{\prime\prime}C- \quad PrPsP \cdot P\ddot{e} \quad PeP^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PSC\dot{r} \quad 1,5 \quad P_jP_i \quad C\Gamma P^{\circ}P \cdot PsP_j \quad P \cdot$
 $P_iC\Gamma PsPrC\dot{r}PeC, P^{\circ}P_jP\ddot{e} \quad P \cdot \quad PIP\ddot{e}C\Gamma PsPeP\ddot{e}P_j \quad PIP_jC-C\Gamma C, PsP_j \quad P\parallel P\ddot{e}C\dot{H}C\dot{r} \quad (PIC-Pr \quad 900 \quad PrPs \quad 1 \quad 000$
 $PeP^{\circ}P \gg PsC\dot{H}C-P\ddot{N}e) \quad P \cdot PSP^{\circ}C\dot{+}PSPs \quad PSP_{\mu} \quad PIP_iP \gg P\ddot{e}PIP^{\circ}PI \quad PSP^{\circ} \quad P_iPsPeP^{\circ}P \cdot PSP\ddot{e}Pe \quad C_{max} \quad P^{\circ}P \pm Ps \quad AUC$
 $PeP^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PSC\dot{r} \quad (P \cdot PSP^{\circ}C\dot{+}P_{\mu}PSPSC\Gamma P \quad AUC_{0B\dot{H}^{\circ}\infty} \quad P_iC-PrPIP\ddot{e}C\%C\dot{r}PIP^{\circ}P \gg PsC\Gamma C\Gamma P \quad PSP^{\circ}$
 $12 \%, \quad P \cdot PSP^{\circ}C\dot{+}P_{\mu}PSPSC\Gamma P \quad C_{max} \quad P \cdot PSP\ddot{e}P\parallel C\dot{r}PIP^{\circ}P \gg PsC\Gamma C\Gamma P \quad PSP^{\circ} \quad 5 \%, \quad P_iC\dot{H}P\ddot{e} \quad P_iC\dot{H}P\ddot{e}P\ddot{N}ePsP_jC-$
 $P \gg C-PeP^{\circ}C\dot{H}C\Gamma CH\dot{P}PePsP_iPs \quad P \cdot P^{\circ}C\Gamma PsP \pm C\dot{r} \quad P_iC-C\Gamma P \gg C\Gamma P \quad C-P\parallel C- \quad PI \quad P_iPsC\dot{H}C-PIPSC\Gamma PSC- \quad P \cdot$
 $P_iC\dot{H}P\ddot{e}P\ddot{N}ePsP_jPsP_j \quad PSP^{\circ}C, C\%P_{\mu}). \quad P\dot{+}P\parallel P^{\circ} \quad C, P^{\circ}PePsP\parallel \quad P_jP^{\circ}P \gg P^{\circ} \quad P_jC-PSC-P_jP^{\circ}P \gg CH\dot{P}SP\ddot{e}P\ddot{N}e$
 $PIP_iP \gg P\ddot{e}PI \quad PSP^{\circ} \quad P_jP_{\mu}C, P^{\circ}P \pm PsP \gg C-C, P\ddot{e} \quad DCAR \quad C, P^{\circ} \quad DDCAR.$

$P_{\mu}P^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PS \quad P_jPsP\parallel PSP^{\circ} \quad P \cdot P^{\circ}C\Gamma C, PsC\Gamma PsPIC\dot{r}PIP^{\circ}C, P\ddot{e} \quad PSP_{\mu}P \cdot P^{\circ}P \gg P_{\mu}P\parallel PSPs \quad PIC-Pr$
 $P_iC\dot{H}P\ddot{e}P\ddot{N}ePsP_jC\dot{r} \quad C-P\parallel C-.$

P PsP · P_iPsPrC-P»

$P\acute{K}P^{\circ} \quad PsC\Gamma PSPsPIC- \quad P_iPsP_iC\dot{r}P \gg C\Gamma C\dot{+}C-P\ddot{N}ePSPsP_iPs \quad P^{\circ}PSP^{\circ}P \gg C-P \cdot C\dot{r}$
 $C,, P^{\circ}C\dot{H}P_jP^{\circ}PePsPeC-PSP_{\mu}C, P\ddot{e}PeP\ddot{e} \quad C\dot{r}C\Gamma PIPSP\ddot{e}P\ddot{N}e \quad PsP \pm B\dot{H}^{\circ}TM \quad C^{\prime\prime}P_j \quad C\dot{H}PsP \cdot P_iPsPrC-P \gg C\dot{r} \quad (P\dot{H}/P)$
 $C\Gamma PeP \gg P^{\circ}PrP^{\circ}PI \quad 916 \quad P \gg \quad PrP \gg C\Gamma P \quad PeP^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PSC\dot{r}, \quad 475 \quad P \gg \quad PrP \gg C\Gamma P \quad DCAR \quad C, P^{\circ} \quad 1 \quad 568 \quad P \gg$
 $PrP \gg C\Gamma P \quad DDCAR, \quad C\%Ps \quad PIPeP^{\circ}P \cdot C\dot{r}C^{\prime\prime} \quad PSP^{\circ} \quad P \cdot PSP^{\circ}C\dot{+}PSP\ddot{e}P\ddot{N}e \quad C\dot{H}PsP \cdot P_iPsPrC-P \gg$
 $PeP^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PSC\dot{r} \quad C, P^{\circ} \quad P\ddot{N}ePsP_iPs \quad PsC\Gamma PSPsPIPSP\ddot{e}C \dots \quad P^{\circ}PeC, P\ddot{e}PIPSP\ddot{e}C \dots$
 $P_jP_{\mu}C, P^{\circ}P \pm PsP \gg C-C, C-PI. \quad P_{\mu}P^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PS \quad C, P^{\circ} \quad P\ddot{N}ePsP_iPs \quad PsC\Gamma PSPsPIPSC-$
 $P^{\circ}PeC, P\ddot{e}PIPSC- \quad P_jP_{\mu}C, P^{\circ}P \pm PsP \gg C-C, P\ddot{e} \quad P^{\circ}PeC, P\ddot{e}PIPSPs \quad P \cdot P_iB\dot{H}^{\circ}TM \quad C\Gamma P \cdot C\dot{r}C\dot{H}C, CH\dot{C}\Gamma C\Gamma P$
 $(PeP^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PS \quad B\dot{H}^{\circ} \quad PSP^{\circ} \quad 96B\dot{H}^{\circ}97 \%, \quad DCAR \quad B\dot{H}^{\circ} \quad PSP^{\circ} \quad 94B\dot{H}^{\circ}97 \%, \quad DDCAR \quad B\dot{H}^{\circ} \quad PSP^{\circ}$
 $92B\dot{H}^{\circ}97 \%) \quad P \cdot \quad P \pm C-P \gg PeP^{\circ}P_jP\ddot{e} \quad P_iP \gg P^{\circ}P \cdot P_jP\ddot{e}.$

P'C-PsC, C_HP^oPSCΓC,, PsC_HP_jP^oC₊C-CΓP

$P_{\mu}P_{\mu}C, P^{\circ}P \pm PsP \gg C-P \cdot P_j \quad PeP^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PSC\dot{r} \quad PIPeP \gg CH\dot{C}\dot{+}P^{\circ}C^{\prime\prime}$
 $PrP_{\mu}P_jP_{\mu}C, P\ddot{e}P \gg CH\dot{P}IP^{\circ}PSPSC\Gamma P \quad (DCAR \quad C, P^{\circ} \quad DDCAR), \quad P_iC-PrC\dot{H}PsPeC\Gamma P\ddot{e}P \gg CH\dot{P}IP^{\circ}PSPSC\Gamma P$
 $(P_iC-PrC\dot{H}PsPeC\Gamma P\ddot{e}Pr \quad PeP^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PSC\dot{r}, \quad HCAR) \quad C, P^{\circ} \quad PePsP_jP \pm C-PSP^{\circ}C\dot{+}C-CH$
 $PrP_{\mu}P_jP_{\mu}C, P\ddot{e}P \gg CH\dot{P}IP^{\circ}PSPSC\Gamma P \quad C, P^{\circ} \quad P_iC-PrC\dot{H}PsPeC\Gamma P\ddot{e}P \gg CH\dot{P}IP^{\circ}PSPSC\Gamma P \quad (P_iC-PrC\dot{H}PsPeC\Gamma P\ddot{e}-$
 $PrP_{\mu}P \cdot P_jP_{\mu}C, P\ddot{e}P \gg -PeP^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PS, \quad HDCAR, \quad C, P^{\circ} \quad P_iC-PrC\dot{H}PsPeC\Gamma P\ddot{e}-$
 $PrP\ddot{e}PrP_{\mu}P \cdot P_jP_{\mu}C, P\ddot{e}P \gg -PeP^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PS, \quad HDDCAR). \quad P\acute{K}P^{\circ}PrP^{\circ}P \gg C- \quad P_jP_{\mu}C, P^{\circ}P \pm PsP \gg C-C, P\ddot{e}$
 $HCAR, \quad HDCAR \quad C, P^{\circ} \quad HDDCAR \quad P \pm C-PsC, C\dot{H}P^{\circ}PSC\Gamma C,, PsC\dot{H}P_jC\dot{r}C\dot{H}C, CH\dot{C}\Gamma C\Gamma P \quad C\dot{r}$
 $PIC-PrP_iPsPIC-PrPSC- \quad PePsPSB\dot{H}^{\circ}TM \quad CH\dot{P}IP^{\circ}C, P\ddot{e} \quad C\dot{r}C\dot{r}P \gg CH\dot{C}, P^{\circ}C, C\dot{r} \quad C, P^{\circ}$
 $P_iP \gg CH\dot{P}eC\dot{r}C\dot{H}PsPSC-PrC\dot{r}. \quad P^{\prime\prime}PsPrP^{\circ}C, PePsPIP\ddot{e}P\ddot{N}e \quad P_jP_{\mu}C, P^{\circ}P \pm PsP \gg C-C, \quad B\dot{H}^{\circ} \quad PeP\ddot{e}C\dot{r}P \gg PsC, P^{\circ}$
 $PrP_{\mu}C\dot{r}PrP\ddot{e}C \dots P \gg PsC\dot{H}PsC,, P_{\mu}PSC-P \gg C\dot{r}-P_iC-P_iP_{\mu}C\dot{H}P^{\circ}P \cdot P\ddot{e}PSC\dot{r}-PeP^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PSC\dot{r}$
 $(DDCPPCAR) \quad B\dot{H}^{\circ} \quad C\dot{r}C, PIPsC\dot{H}C\dot{H}C^{\prime\prime}C, CH\dot{C}\Gamma C\Gamma P \quad PI \quad P_iC\dot{H}PsC\dot{+}P_{\mu}C\dot{r}C-$
 $PrP_{\mu}P \cdot P^{\circ}P \gg PeC-P \gg C\dot{r}PIP^{\circ}PSPSC\Gamma P \quad C, P^{\circ} \quad PSP^{\circ}C\Gamma C, C\dot{r}PIPSPsP_iPs \quad PsPeP\ddot{e}C\dot{r}PSP_{\mu}PSPSC\Gamma P$
 $PeP^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PSC\dot{r}.$

$P_{\mu}P^{\circ}C\Gamma P\ddot{e}P_iC\Gamma P^{\circ}P \cdot P\ddot{e}PS \quad P_jP_{\mu}C, P^{\circ}P \pm PsP \gg C-P \cdot C\dot{r}C^{\prime\prime}C, CH\dot{C}\Gamma C\Gamma P \quad CYP3A4 \quad C, P^{\circ} \quad P_jP_{\mu}PSC\epsilon PsCH$
 $P_jC-C\dot{H}PsCH \quad CYP2D6 \quad PrPs \quad DCAR \quad C, P^{\circ} \quad HCAR. \quad P\acute{K}P^{\circ}PrP^{\circ}P \gg C- \quad DCAR$
 $P_jP_{\mu}C, P^{\circ}P \pm PsP \gg C-P \cdot C\dot{r}C^{\prime\prime}C, CH\dot{C}\Gamma C\Gamma P \quad CYP3A4 \quad C, P^{\circ} \quad P_jP_{\mu}PSC\epsilon PsCH \quad P_jC-C\dot{H}PsCH \quad CYP2D6 \quad PrPs$
 $DDCAR \quad C, P^{\circ} \quad HDCAR. \quad P_{\mu}PsC, C-P_j \quad DDCAR \quad P_jP_{\mu}C, P^{\circ}P \pm PsP \gg C-P \cdot C\dot{r}C^{\prime\prime}C, CH\dot{C}\Gamma C\Gamma P \quad P \cdot P^{\circ}$
 $PrPsP_iPsP_jPsP_iPsCH \quad CYP3A4 \quad PrPs \quad HDDCAR.$

$\text{P}_{4\text{C}}\text{C}-\text{C}\Gamma\text{P}\gg\text{C}\Pi\text{ P}\ddot{\text{P}}\text{S}\text{P}\text{IC}, \text{P}\text{sC}\overline{\text{C}}\text{P}\text{S}\text{P}\text{S}\text{P}\ddot{\text{P}}\text{P}\text{S P}\cdot\text{P}^{\circ}\text{C}\Gamma\text{C}, \text{P}\text{sC}\Gamma\text{C}\Gamma\text{P}\text{IP}^{\circ}\text{P}\text{S}\text{P}\text{S}\text{C}\Pi\text{ P}\text{e}\text{P}\text{s}\text{P}\text{S}\text{C}\ddagger\text{P}_{\mu}\text{P}\text{S}\text{C}, \text{C}\overline{\text{C}}\text{P}^{\circ}\text{C}\ddagger\text{C}-\text{C}\Pi$
 $\text{P}\text{eP}^{\circ}\text{C}\overline{\text{C}}\text{P}\ddot{\text{P}}\text{P}\ddot{\text{P}}\text{C}\overline{\text{C}}\text{P}^{\circ}\text{P}\cdot\text{P}\ddot{\text{P}}\text{P}\text{S}\text{C}\Gamma\text{ C}, \text{P}^{\circ}\text{ P}\text{N}_{\text{e}}\text{P}\text{s}\text{P}\text{P}\text{S PrP}\text{IP}\text{S}\text{C}\dots \text{P}\text{sC}\Gamma\text{P}\text{S}\text{P}\text{S}\text{P}\text{IP}\text{S}\text{P}\ddot{\text{P}}\text{C}\dots \text{P}^{\circ}\text{P}\text{eC}, \text{P}\ddot{\text{P}}\text{IP}\text{S}\text{P}\ddot{\text{P}}\text{C}\dots$
 $\text{PjP}_{\mu}\text{C}, \text{P}^{\circ}\text{P}\pm\text{P}\text{sP}\gg\text{C}-\text{C}, \text{C}-\text{PI}, \text{PrP}_{\mu}\text{P}\cdot\text{PjP}_{\mu}\text{C}, \text{P}\ddot{\text{P}}\text{P}\gg-\text{P}\text{eP}^{\circ}\text{C}\overline{\text{C}}\text{P}\ddot{\text{P}}\text{P}\ddot{\text{P}}\text{C}\overline{\text{C}}\text{P}^{\circ}\text{P}\cdot\text{P}\ddot{\text{P}}\text{P}\text{S}\text{C}\Gamma\text{ (DCAR) C}, \text{P}^{\circ}$
 $\text{PrP}\ddot{\text{P}}\text{PrP}_{\mu}\text{P}\cdot\text{PjP}_{\mu}\text{C}, \text{P}\ddot{\text{P}}\text{P}\gg-\text{P}\text{eP}^{\circ}\text{C}\overline{\text{C}}\text{P}\ddot{\text{P}}\text{P}\ddot{\text{P}}\text{C}\overline{\text{C}}\text{P}^{\circ}\text{P}\cdot\text{P}\ddot{\text{P}}\text{P}\text{S}\text{C}\Gamma\text{ (DDCAR), PI P}\ddot{\text{P}}\text{P}\gg\text{P}^{\circ}\text{P}\cdot\text{PjC}-$
 $\text{P}\ddot{\text{P}}\text{C}-\text{PrP}\text{IP}\ddot{\text{P}}\text{C}\%_{\text{O}}\text{C}\Gamma\text{C}^{\circ}\text{C}, \text{C}\text{H}\text{C}\Gamma\text{C}\Pi\text{ P}\ddot{\text{P}}\text{C}\overline{\text{C}}\text{P}\text{s}\text{P}\ddot{\text{P}}\text{P}\text{sC}\overline{\text{C}}\text{C}-\text{P}\text{N}_{\text{e}}\text{P}\text{S}\text{P}\text{s PrC}-\text{P}^{\circ}\text{P}\ddot{\text{P}}\text{P}^{\circ}\text{P}\cdot\text{P}\text{sP}\text{S}\text{C}\Gamma$
 $\text{C}, \text{P}_{\mu}\text{C}\overline{\text{C}}\text{P}^{\circ}\text{P}\ddot{\text{P}}\text{P}_{\mu}\text{P}\text{IC}, \text{P}\ddot{\text{P}}\text{C}\ddagger\text{P}\text{S}\text{P}\text{S}\text{C}-\text{PrP}\text{sP}\cdot\text{P}\ddot{\text{P}}\text{ PIC}-\text{Pr } 1,5 \text{ PjPi PrPs } 6 \text{ PjPi}.$

PhPeC₇H₇P_μPjC- PiC₇H₇PiP_ē PiP°C†C-C''PSC,C-PI

PμPsC₇H₇CzC€PμPSPSC† C,,CzPSPeC†C-C— PSPēC₇H₇PsPe

PμPsPiC†P»C†C†C-PM_ēPSPμ C,,P°C₇H₇PjP°PePsPeC-PSPμC,PēC†PSPμ PjPsPrPμP»C₇H₇PIP°PSPSC† PIPēPePsPSC†PIP°P»PsC†C† P· PIPēPePsC₇H₇PēC†C,P°PSPSC†Pj PrP°PSPēC... PIC-Pr PiP°C†C-C''PSC,C-PI, C†PeC- P±C†P»Pē P·P°P»C†C†PμPSC- PrPs PeP»C-PSC-C†PSPsPiPs PrPsC†P»C-PrP¶PμPSPSC† C†C,PSC†PsPIPS P»C-PeC†PIP°PSPSC† C€PēP·PsC,,C₇H₇PμPSC-C— PeP°C₇H₇PēPiC₇H₇P°P·PēPSPsPj, P· C₇H₇-P·PSPēPjPē PiPsPeP°P·PSPēPeP°PjPē C,,C†PSPeC†C-C— PSPēC₇H₇PsPe, PIPeP»C₇H₇C†PSPs P· PSPsC₇H₇PjP°P»C₇H₇PSPsC₇ C,,C†PSPeC†C-C₇ PSPēC₇H₇PsPe (PeP»C-C₇H₇PμPSC† PeC₇H₇PμP°C,PēPSC-PSC† (CrCl) ≥ 90 PjP»/C...PI) C- PiPsC₇H₇C€PμPSPSC†PjPē C,,C†PSPeC†C-C— PSPēC₇H₇PsPe P»PμPiPePsPiPs (CrCl 60B₇"89 PjP»/C...PI) C,P° C†PμC₇H₇PμPrPSC₇H₇PsPiPs C†C,C†PiPμPSC† C,C†P¶PePsC†C,C- (CrCl 30B₇"59 PjP»/C...PI). P₇H-C-P¶ PeP»C-C₇H₇PμPSC†PsPj PeP°C₇H₇PēPiC₇H₇P°P·PēPSC† PI PiP°P°P·PjC-C,P° PeP»C-C₇H₇PμPSC†PsPj PeC₇H₇PμP°C,PēPSC-PSC† PSPμ P±C†P»Ps P·PSP°PM_ēPrPμPSPs P¶PsPrPSPēC... C-C†C,PsC,PSPēC... P·PIB₇TM C†P·PeC-PI.

P''C-C† PeP°C₇H₇PēPiC₇H₇P°P·PēPSC† PSPμ PsC†C-PSC₇H₇PIP°P»P°C†C† C† PiP°C†C-C''PSC,C-PI C-P· C,C†P¶PePēPjPē (CrCl < 30 PjP»/C...PI) PiPsC₇H₇C€PμPSPSC†PjPē C,,C†PSPeC†C-C— PSPēC₇H₇PsPe (PrPēPI. C₇H₇PsP·PrC-P» B«PŸPiPsC†C-P± P·P°C†C,PSC†C†PIP°PSPSC† C,P° PrPsP·PēB»).

PμPsC₇H₇CzC€PμPSPSC† C,,CzPSPeC†C-C— PiPμC†C-PSPePē

P''PsC†P»C-PrP¶PμPSPSC† PI PrPIPsC... C†P°C†C,PēPSP°C... (PsPrPSPsPeC₇H₇P°C,PSP° PrPsP·P° PeP°C₇H₇PēPiC₇H₇P°P·PēPSC† 1 PjPi [C†P°C†C,PēPSP° P₇] C,P° PeP°C₇H₇PēPiC₇H₇P°P·PēPS C† PrPsP±PsPIC-PM_ē PrPsP·C- 0,5 PjPi PiC₇H₇PsC,C†PiPsPj 14 PrPSC-PI [C†P°C†C,PēPSP° P']) PiC₇H₇PsIPsPrPēP»PsC†C† P·P° C†C†P°C†C,C₇H₇ PiP°C†C-C''PSC,C-PI P· C₇H₇-P·PSPēPjPē C†C,C†PiPμPSC†PjPē PiPsC₇H₇C†C€PμPSC₇H₇ C,,C†PSPeC†C-C— PiPμC†C-PSPePē (PeP»P°C†Pē P₇ C,P° P' P·P° P₇SP°PM_ēP»PrPsPj B₇" PμB₇TM C₇H₇). PμC-C†P»C† P·P°C†C,PSC†C†PIP°PSPSC† PsPrPSPsPeC₇H₇P°C,PSPsC— PrPsP·Pē PeP°C₇H₇PēPiC₇H₇P°P·PēPSC† 1 PjPi P°P±Ps PiC₇H₇PēPM_ēPsPjC† 0,5 PjPi PeP°C₇H₇PēPiC₇H₇P°P·PēPSC† PiC₇H₇PsC,C†PiPsPj 14 PrPSC-PI PiP°C†C-C''PSC,Pē P· PiPsC₇H₇C€PμPSPSC†PjPē C,,C†PSPeC†C-C— PiPμC†C-PSPePē P»PμPiPePsPiPs C†Pē C†PμC₇H₇PμPrPSC₇H₇PsPiPs C†C,C†PiPμPSC† C† PiPsC₇H₇-PIPSC†PSPSC- B P· PiP°C†C-C''PSC,P°PjPē P· PSPsC₇H₇PjP°P»C₇H₇PSPsC₇ C,,C†PSPeC†C-C''C₇H₇ PiPμC†C-PSPePē PjP°P»Pē PiC₇H₇PēP±P»PēP·PSPs PSP° 25 % PIPēC%₀C† PePsPSC†PμPSC,C₇H₇P°C†C-C₇H₇ PeP°C₇H₇PēPiC₇H₇P°P·PēPSC† (C_{max} C,P° AUC) C,P° PiC₇H₇PēP±P»PēP·PSPs PSP° 45 % PSPēP¶C†C† PePsPSC†PμPSC,C₇H₇P°C†C-C₇H₇ PsC†PSPsPIPS PēC... P°PeC,PēPIPS PēC... PjPμC,P°P±PsP»C-C,C-PI, PrPμP·PjPμC,PēP»-PeP°C₇H₇PēPiC₇H₇P°P·PēPSC† C,P° PrPēPrPμP·PjPμC,PēP»-PeP°C₇H₇PēPiC₇H₇P°P·PēPSC†.

PμC-C†P»C† P±P°PiP°C,PSC₇H₇P°P·PsPIPsPiPs PiC₇H₇PēPM_ēPsPjC† PeP°C₇H₇PēPiC₇H₇P°P·PēPSC† PePsPSC†PμPSC,C₇H₇P°C†C-C† P·P°PiP°P»C₇H₇PSPsPiPs P°PeC,PēPIPS PiPs PePsPjPiPsPSPμPSC,P° (CAR DCAR DD CAR) (AUC C,P° C_{max}) C† PiPsC₇H₇-PIPSC†PSPSC- B P· PIC-PrPiPsPIC-PrPSPēPj PiPsPeP°P·PSPēPePsPj C†B PiP°C†C-C''PSC,C-PI P· PSPsC₇H₇PjP°P»C₇H₇PSPsC₇ C,,C†PSPeC†C-C''C₇H₇ PiPμC†C-PSPePē P·PSPēP·PēP»P°C†C† PSP° 21B₇"22 % C,P° 13B₇"15 % PIC-PrPiPsPIC-PrPSPs C† B PiP°C†C-C''PSC,C-PI P· PiPsC₇H₇C€PμPSPSC†Pj C,,C†PSPeC†C-C— PiPμC†C-PSPePē (PμPαPμ) P»PμPiPePsPiPs C†Pē C†PμC₇H₇PμPrPSC₇H₇PsPiPs C†C,C†PiPμPSC† C,C†P¶PePsC†C,C-, C†PeC%₀Ps PIC₇H₇P°C...PsPIC†PIP°P»PēC†C† PSPμP·PIB₇TM C†P·P°PSC- P·PIB₇TM C†P·P°PSC- PePsPSC†PμPSC,C₇H₇P°C†C-C—, B C,PSPrC- C†Pe PrP»C† PSPμP·PIB₇TM C†P·P°PSPsPiPs P·P°PiP°P»C₇H₇PSPsPiPs PePsPjPiPsPSPμPSC,P° P±C†P»Ps C₇H₇PsP·C₇H₇P°C...PsPIP°PSPs P·PSPēP¶PμPSPSC† PSP° 12B₇"13 % C,P° PiC-PrPIPēC%₀PμPSPSC† PSP° 20B₇"25 %

PIC-PrPⁱPsPIC-PrPSPs B Cr PⁱP^oC†C-C" PSC,C-PI P· P_uP_αP_u P»P_uP_iPePsPiPs C,P^o
C†P_uC†P_uPrPSC_HPsPiPs C†C,CrP_iP_uPSC† C,C†P_u¶PePsC†C,C-.

P" C-C† PeP^oC†P_iPePⁱC†P^oP·P_ePSC† PSP_u PsC†C-PSC_HPIP^oP»P^oC†C† Cr PⁱP^oC†C-C" PSC,C-PI C-P·
C,C†P_u¶PeP_eP_jP_e (PeP»P^oC† P_Y P·P^o P§P^oPN_eP»PrPsP_j B†" P_uB†TM C†) P_iPsC†C†C†P_uPSPSC†P_jP_e
C,,CrPSPeC†C-C— P_iP_uC†C-PSPeP_e (PrP_ePI. C†PsP·PrC-P» B«P_YP_iPsC†C-P±
P·P^oC†C,PsC†C†P^oPSPSC† C,P^o PrPsP·P_eB»).

P' C-Pe, C†C,P^oC,C_H C,P^o C†P^oC†P^o

P_j P_iPsP_iC†P»C†C†C-PN_ePSPsP_jC† C,,P^oC†P_jP^oPePsPeC-PSP_uC,P_eC†PSPsP_jC† P^oPSP^oP»C-P·C-
PSP_u P±C†P»Ps PeP»C-PSC-C†PSPs P·PSP^oC†P_eP_jP_eC... PIC-PrP_jC-PSPSPsC†C,P_uPN_e Cr
PⁱP^oC†P^oP_jP_uC,C†P^oC... C,,P^oC†P_jP^oPePsPeC-PSP_uC,P_ePeP_e (AUC C,P^o C_{max}B
PeP^oC†P_iPePⁱC†P^oP·P_ePSC† C,P^o PN_ePsPiPs PsC†PSPsPIPS_eC... P^oPeC,P_ePIPS_eC...
P_jP_uC,P^oP±PsP»C-C,C-PI) P·P^oP»P_uP¶PSPs PIC-Pr PIC-PeC†, C†C,P^oC,C- C,P^o C†P^oC†P_e
PⁱP^oC†C-C" PSC,C-PI. P_uC-Pr C†P^oC† P^oPSP^oP»C-P·C† B B PIP_ePIC†P^oP»P_e B PrP^oPSC-
B C_{oo}PsPrPs B 2 844 PⁱP^oC†C-C" PSC,C-PI C†C-P·PSP_eC... C†P^oC†, P· C†PeP_eC... 536
PⁱP^oC†C-C" PSC,C-PI P±C†P»P_e PIC-PePsP_j PIC-Pr 50 PrPs 65 C†PsPeC-PI, 933 PsC†PsP±P_e
P±C†P»P_e P¶C-PSPeP^oP_jP_e (PrP_ePI. C†PsP·PrC-P» B«P_YP_iPsC†C-P± P·P^oC†C,PsC†C†P^oPSPSC†
C,P^o PrPsP·P_eB»). P" P^oPSC- C_{oo}PsPrPs PⁱP^oC†C-C" PSC,C-PI P»C-C,PSC_HPsPiPs PIC-PeC†
(P_iPsPSP^oPr 65 C†PsPeC-PI) PsP±P_jP_uP¶P_uPSC-.

P_uP^oP»C-PSPSC†

P§P_uC†P_uP· C,P_u, C_{oo}Ps PeP^oC†P_iPePⁱC†P^oP·P_ePS PSP_u C" C†C†P±C†C,C†P^oC,PsP_j PrP»C†
CYP1A2, PSP_u PsC†C-PeC†C" C,C_HC†C† PIP_iP»P_ePIC† PⁱP^oP»C-PSPSC† PSP^o
C,,P^oC†P_jP^oPePsPeC-PSP_uC,P_ePeC† PeP^oC†P_iPePⁱC†P^oP·P_ePSC†.

P_uPsC,P_uPSC†C-PN_ePSP_ePN_e PIP_iP»P_ePI PeP^oC†P_iPePⁱC†P^oP·P_ePSC† PSP^o C-PSC†C-
P_iC†P_uPⁱP^oC†P^oC,P_e

P_uP^oC†P_iPePⁱC†P^oP·P_ePS C,P^o PN_ePsPiPs PsC†PSPsPIPS_eC- P^oPeC,P_ePIPS_eC- P_jP_uC,P^oP±PsP»C-C,P_e
PSP_u C-PSP_rC†PeC†P^oP»P_e P_uPSP·P_eP_jP_e CYP1A2, CYP2B6 C,P^o CYP3A4 C- PSP_u P±C†P»P_e
C-PSP_iC-P±C-C,PsC†P^oP_jP_e CYP1A2, CYP2A6, CYP2B6, CYP2C8, CYP2C9, CYP219, CYP2D6,
CYP2E1 C,P^o CYP3A4 *in vitro*. P_uP^oC†P_iPePⁱC†P^oP·P_ePS C,P^o PN_ePsPiPs PsC†PSPsPIPS_eC-
P^oPeC,P_ePIPS_eC- P_jP_uC,P^oP±PsP»C-C,P_e PSP_u C" C-PSP_iC-P±C-C,PsC†P^oP_jP_e
C,C†P^oPSC†P_iPsC†C,PSP_eC... P±C-P»PeC-PI OATP1B1, OATP1B3, BCRP,
PsC†P_iP^oPSC-C†PSPsPiPs PeP^oC,C-PsPSPSPsPiPs C,C†P^oPSC†P_iPsC†C,PSPsPiPs P±C-P»PeP^o 2
(OCT2) C,P^o PsC†P_iP^oPSC-C†PSP_eC... P^oPSC-PsPSPSP_eC... C,C†P^oPSC†P_iPsC†C,PSP_eC...
P±C-P»PeC-PI 1 C,P^o 3 (OAT1 C,P^o OAT3) *in vitro*. DCAR C,P^o DDCAR PSP_u P±C†P»P_e
C-PSP_iC-P±C-C,PsC†P^oP_jP_e C,C†P^oPSC†P_iPsC†C,PSPsPiPs P±C-P»PeP^o P-gp, C...PsC†P^o
PeP^oC†P_iPePⁱC†P^oP·P_ePS P±C†PI C-PSP_iC-P±C-C,PsC†PsP_j P-gp PI PeP_eC†P_uC†PSP_ePeC† (PrP_ePI.
C†PsP·PrC-P» B«P·P·P^oC" P_jPsPrC-C† P· C-PSC†P_eP_jP_e P»C-PeP^oC†C†C_HPeP_eP_jP_e
P·P^oC†PsP±P_jP_e C,P^o C-PSC†C- PIP_ePrP_e PIP·P^oC" P_jPsPrC-PN_eB»).

P" P^oPSC- PrPsPeP»C-PSC-C†PSP_eC... PrPsC†P»C-PrP¶P_uPSC_H P· P±P_uP·P_iP_uPeP_e

P_uP^oC†P_iPePⁱC†P^oP·P_ePS C†P_iC†P_eC†P_ePSC†PI PrPIP_sC†C,PsC†PsPSPSC_H PeP^oC,P^oC†P^oPeC,C†
C,P^o PIC,PsC†P_ePSPSC- P·P_jC-PSP_e C†C-C,PeC-PIPeP_e (PIC-PrC†P^oC†P^oPSPSC† C,P^o
PeC-C†C,PsPiPsPrC-P±PSC† PrP_uP_iP_uPSP_uC†C-C† C†C-C,PeC-PIPeP_e) Cr C†PsP±P^oPe.
P—P^oP_iP^oP»C_HPSP^o P_uPeC†P_iPsP·P_eC†C-C† (AUC) PeP^oC†P_iPePⁱC†P^oP·P_ePSC†,
B PrPsC†C†P_iPSC†C,P^o P_iC†P_e P·P^oC†C,PsC†C†P^oPSPSC- P_jP^oPeC†P_eP_jP^oP»C_HPSPsC—

PrPsP·Pë, C%Ps PSPµ PīCḡPëP·PIPsPrPëC,CH PrPs CḡPsP·PIPëC,PeCí CĪPIPSPëC...
PSPµP±P°P¶P°PSPëC... PµC,,PµPeC,C-PI (NOAEL) P·P±PsPeCí PsCḡPiP°PSC-PI P·PsCḡCí, Cí 4,2
CḡP°P·P° PīPµCḡPµPIPëC%CrC” PeP»C-PSC-C±PSCí AUC PīCḡPë PjP°PeCíPëPjP°P»CHPSC-PNë
CḡPµPePsPjPµPSPPrPsPIP°PSC-PNë PrPsP·C- PrP»CĪ P»CḡPrPëPSPë (PḡP P”P) 6 PjPi PSP°
PrPsP±Cí. P—P±C-P»CHCëPµPSPSCĪ C±P°CíC,PsC,Pë
PrPµPiPµPSPµCḡP°C†C-C—/P°C,CḡPsC,,C-C— CíC-C,PeC-PIPePë
CíPiPsCíC,PµCḡC-PiP°P»PsCíCĪ Cí C%CrCḡC-PI-P°P»CHP±C-PSPsCíC-PI Cí 2-
CḡC-C±PSPsPjCí PrPsCíP»C-PrP¶PµPSPSC- PSP° C,P»C- PeP»C-PSC-C±PSPs
P·PSP°C±CíC%PsC— PµPeCíPiPsP·PëC†C-C— PīCḡPµPiP°CḡP°C,Cí.

PαPsCíC,,PsP»C-PīC-PrPsP· CíPiPsCíC,PµCḡC-PiP°PICíCĪ Cí P»PµPiPµPSCĪC... C%CrCḡC-PI,
CíPsP±P°Pe C,P° PjPëCëPµPNë (P·P·P°PiP°P»PµPSPSCĪPj P°P±Ps P±PµP· PSCḡPsPiPs) C,P° Cí
PePsCḡC- PSP°PrPSPëCḡPePsPIPëC... P·P°P»PsP· CíPsP±P°Pe PSP° C,,PsPSC-
PeP»C-PSC-C±PSPs P·PSP°C±CíC%PëC... PµPeCíPiPsP·PëC†C-PNë PīCḡPµPiP°CḡP°C,Cí.
PŸPiPsCíC,PµCḡC-PiP°P»PsCíCĪ P·P°PiP°P»PµPSPSCĪ P»PµPiPµPSCḡ Cí CíPsP±P°Pe, CĪPeC-
PsC,CḡPëPjCíPIP°P»Pë PeP°CḡPëPīCḡP°P·PëPS PīCḡPsC,CĪPiPsPj 1 CḡPsPeCí, PīCḡPë
C†CHPsPjCí NOAEL C,P° PIC-PrPiPsPIC-PrPSC- PiPsPeP°P·PSPëPePë AUC
PīPµCḡPµPIPëC%CrPIP°P»Pë B C,P°PeC- Cí CḡP°P·C- P·P°CíC,PsCíCíPIP°PSPSCĪB PḡP P”P>
Cí 2,7 (CíP°PjC†C-) C,P° 1,7 (CíP°PjPePë) CḡP°P·P°. KóP°PīCḡPëPeC-PSC†C- 2-
PjC-CíCĪC±PSPsPiPs PīPµCḡC-PsPrCí P±PµP· PīCḡPëPNëPsPjCí PīCḡPµPiP°CḡP°C,Cí PīCḡPë
PµPeCíPiPsP·PëC†C-C—, CĪPeP° PI 4,2 CḡP°P·P° B PīPµCḡPµPIPëC%CrC” PeP»C-PSC-C±PSCí
PµPeCíPiPsP·PëC†C-CḡB PīCḡPë PḡP P”P>, P·P°PiP°P»PµPSPSCĪ PSPµ
CíPiPsCíC,PµCḡC-PiP°P»PsCíCĪ; PīCḡPsC,Pµ P·P°PiP°P»PµPSPSCĪ P·P±PµCḡC-PiP°P»PsCíCĪ
PīCḡPë P±C-P»CHCë PIPëCíPsPePëC... PrPsP·P°C....

P”C-PīPµCḡC,CḡPsC,,C-CĪ PePsCḡPë PSP°PrPSPëCḡPePsPIPëC... P·P°P»PsP·
CíPiPsCíC,PµCḡC-PiP°P»P°CíCĪ PSP° C,P»C- PePsPSC†PµPSC,CḡP°C†C-C—
PīCḡPµPiP°CḡP°C,Cí, C%Ps PI 4,1 CḡP°P·P° PīPµCḡPµPIPëC%CrC” B C,P°PeCí PīCḡPë
P·P°CíC,PsCíCíPIP°PSPSC- PḡP P”P>, Cí C%CrCḡC-PI (C,C-P»CHPePë Cí CíP°PjPsPe) C,P°
PīCḡPë PeP»C-PSC-C±PSPs P·PSP°C±CíC%PëC... P·P°PiP°P»CHPSPëC...
PePsPSC†PµPSC,CḡP°C†C-CĪC... PeP°CḡPëPīCḡP°P·PëPSCí PI PīP»P°P·PjC- Cí PjPëCëPµPNë. Pj
CíPsP±P°Pe CíPiPsCíC,PµCḡC-PiP°P»P°CíCĪ PsP±PsCḡPsC,PSP°
PiC-PīPµCḡC,CḡPsC,,C-CĪ/PiC-PīPµCḡPiP°P°P·C-CĪ C,P°
PIP°PeCíPsP»C-P·P°C†C-CĪ/PIPµP·PëPeCíP»CĪC†C-CĪ PePsCḡPë PSP°PrPSPëCḡPePsPIPëC...
P·P°P»PsP·, NOAEL Cí 4,2 CḡP°P·P° PīPµCḡPµPIPëC%CrPIP°P»P° B C,P°PeCí PīCḡPë PḡP P”P>.

Pj CíP°PjPsPe C%CrCḡC-PI PIC-PrP·PSP°C±P°P»PsCíCĪ P·PSPëP¶PµPSPSCĪ
PīP»PsPrCḡC±PsCíC,C- C,P° C-PSPPrPµPeCíCí P·P°PiP»C-PrPSPµPSPSCĪ PīCḡPë
PeP»C-PSC-C±PSPs P·PSP°C±CíC%PëC... PµPeCíPiPsP·PëC†C-CĪC... Cí
PīPµCḡPµCḡP°C...CíPSPeCí PSP° PjPi/Pj² PīP»PsC%C- PīPsPIPµCḡC...PSC- C,C-P»P°.
P’PīP»PëPICí PSP° C,,PµCḡC,PëP»CHPSC-CíC,CH CíP°PjC†C-PI PīCḡPë
PµPeCíPiPsP·PëC†C-CĪC..., C%Ps PīPµCḡPµPIPëC%CrCḡC,CH PeP»C-PSC-C±PSCí
PµPeCíPiPsP·PëC†C-Cḡ PīCḡPë PḡP P”P> Cí 4,3 CḡP°P·P°, PSPµ
CíPiPsCíC,PµCḡC-PiP°P»PsCíCĪ.

P’PIPµPrPµPSPSCĪ PeP°CḡPëPīCḡP°P·PëPSCí C%CrCḡP°Pj Cí PīPµCḡC-PsPr
PsCḡPiP°PSPsPiPµPSPµP·Cí B CíPīCḡPëC±PëPSCĪP»Ps PIP°PrPë CḡPsP·PIPëC,PeCí,
P·PSPëP¶PµPSPSCĪ PIPëP¶PëPIP°PSPSCĪ PīPsC,PsPjCíC,PIP° C,P° P·P°C,CḡPëPjPeCí
CḡPsP·PIPëC,PeCí PīCḡPë PµPeCíPiPsP·PëC†C-CĪC... PīCḡPµPiP°CḡP°C,Cí, PjPµPSCëPëC...,
PSC-P¶B PµPeCíPiPsP·PëC†C-CĪ Cí P»CḡPrPëPSPë PīCḡPë PḡP P”P> 6 PjPi PSP° PrPsP±Cí. Pj
PeCḡPsP»C-PI PeP°CḡPëPīCḡP°P·PëPS B C±PëPSPëPI C,PsPeCíPëC±PSCí PrC-Cḡ PSP°
PjP°C,PµCḡPëPSCíCHPePëPNë PsCḡPiP°PSC-P·Pj, P°P»Pµ PSPµ CíC,P°PSPsPIPëPI

PSPμP±PμP·PīPμPePē PrP»CΠ PīP»PsPrP° PīCḡPē PμPeCΓPīPsP·PēC†C-CΠC..., Cr 5,8 CḡP°P·P° PIPēC%oPēC... P·P° PeP»C-PSC-C‡PSCr PμPeCΓPīPsP·PēC†C-Cḡ PīCḡPē PḡP P°P».

PḡP»C-PSC-C‡PSPs P·PSP°C‡CrC%oC- PμPeCΓPīPsP·PēC†C-C— PeP°CḡPēPīCḡP°P·PēPSCr, CΠPePēPNē PIPsPrPēP»Pē PIP°PiC-C,PSPēPj CΓP°PjPeP°Pj C%oCrCḡC-PI Cr PīPμCḡC-PsPr PsCḡPīP°PSPsPiPμPSPμP·Cr, PīCḡPsC,CΠPiPsPj CrCΓC-C”C— PIP°PiC-C,PSPsCΓC,C- C,P° PīPμCḡC-PsPrCr PīCḡCrPrPSPsPiPs PIPēPiPsPrPsPICrPIP°PSPSCΠ,B P·PSPēP¶CΓPIP°P»Pē PīPsCΓC,PSP°C,P°P»CHPSPμ PIPēP¶PēPIP°PSPSCΠ, PjP°CΓCr C,C-P»P° PīCḡPē PSP°CḡPsPrP¶PμPSPSC- C,P° PjP°CΓCr C,C-P»P° PīC-CΓP»CΠ PIC-PrP»CrC‡PμPSPSCΠ PIC-Pr PīCḡCrPrPμPNē PīPsC,PsPjCΓC,PIP°B PīPμCḡC€PsPiPs PīPsPePsP»C-PSPSCΠ. PḡCḡC-Pj C,PsPiPs, P·P° PIC-PrCΓCrC,PSPsCΓC,C- C,PsPeCΓPēC‡PSPsC— PrC-C— PSP° PjP°C,PμCḡPēPSCΓCHPePēPNē PsCḡPīP°PSC-P·Pj CΓPīPsCΓC,PμCḡC-PiP°P»P°CΓCΠ P±P»C-PrC-CΓC,CH,B P·PSPēP¶PμPSPSCΠ C,PμPjPīPμCḡP°C,CrCḡPē C,C-P»P° C,P° P·P°C,CḡPēPjPeP° CḡPsP·PIPēC,PeCr Cr PīP»PsPrP° (PSPμPrPsCḡPsP·PIPēC,PsPe PSPēCḡPePsPIPēC... CΓPsCΓPsC‡PeC-PI C,P° P·PSPēP¶PμPSPSCΠ CΓP»CrC...PsPiPsPiPs CḡPμC,,P»PμPeCΓCr Cr CΓP°PjC†C-PI). PαPμCḡC,PēP»CHPSC-CΓC,CH B PīPsC,PsPjCΓC,PIP° B PīPμCḡC€PsPiPs PīPsPePsP»C-PSPSCΠ B PSPμ P·P°P·PSP°P»P° P·PjC-PS, PsPrPSP°Pe Cr C%oCrCḡC-PI PrCḡCrPiPsPiPs PīPsPePsP»C-PSPSCΠ PIC-PrP·PSP°C‡P°P»PēCΓCΠ P°PSP°P»PsPiC-C‡PSC- PeP»C-PSC-C‡PSC- PsP·PSP°PePē C,P° P·PSPēP¶PμPSPSCΠ PjP°CΓPē C,C-P»P°.

PḡP°CḡPēPīCḡP°P·PēPS C,P° PNēPsPiPs PjPμC,P°P±PsP»C-C,Pē PīCḡPsPSPēPeP°P»Pē PI PjPsP»PsPePs C%oCrCḡC-PI Cr PīPμCḡC-PsPr PīCḡCrPrPSPsPiPs PIPēPiPsPrPsPICrPIP°PSPSCΠ.

PḡP»C-PSC-C‡PSC- C...P°CḡP°PeC,PμCḡPēCΓC,PēPePē.

PμPsPeP°P·P°PSPSCΠ.

P°P»CΠ P»C-PeCrPIP°PSPSCΠ CēPēP·PsC,,CḡPμPSC-C— Cr PrPsCḡPsCΓP»PēC... PīP°C†C-C”PSC,C-PI.

PμCḡPsC,PēPīPsPeP°P·P°PSPSCΠ.

PμC-PrPIPēC%oPμPSP° C‡CrC,P»PēPIC-CΓC,CH PrPs PrC-CḡC‡PsC— CḡPμC‡PsPIPēPSPē P°P±Ps PrPs P±CΓPrCH-CΠPePsC— P· PrPsPīPsPjC-P¶PSPēC... CḡPμC‡PsPIPēPS.

PŸCrPīCrC,PSC” P·P°CΓC,PsCΓCrPIP°PSPSCΠ PīPsC,CrP¶PSPsPiPs C-PSPiC-P±C-C,PsCḡP° CYP3A4 (PrPēPI. CḡPsP·PrC-P» B«P’P·P°C”PjPsPrC-CΠ P· C-PSC€PēPjPē P»C-PeP°CḡCΓCHPePēPjPē P·P°CΓPsP±P°PjPē C,P° C-PSC€C- PIPēPrPē PIP·P°C”PjPsPrC-PNēB»).

PŸCrPīCrC,PSC” P·P°CΓC,PsCΓCrPIP°PSPSCΠ PīPsC,CrP¶PSPsPiPs P°P±Ps PīPsPjC-CḡPSPsPiPs C-PSPrCrPeC,PsCḡP° CYP3A4 (PrPēPI. CḡPsP·PrC-P» B«P’P·P°C”PjPsPrC-CΠ P· C-PSC€PēPjPē P»C-PeP°CḡCΓCHPePēPjPē P·P°CΓPsP±P°PjPē C,P° C-PSC€C- PIPēPrPē PIP·P°C”PjPsPrC-PNēB»).

P’P·P°C”PjPsPrC-CΠ P· C-PSC€PēPjPē P»C-PeP°CḡCΓCHPePēPjPē P·P°CΓPsP±P°PjPē C,P° C-PSC€C- PIPēPrPē PIP·P°C”PjPsPrC-PNē.

PuPsC,PuPSC†C-PN,PSPëPN,PIP†P»PëPI C-PSC€PëC... P†C†PuP†P°C†P°C,C-PI PSP°
PeP°C†PëP†C†P°P·PëPS

$\text{P}\mu\text{C}, \text{P}^\circ\text{P}\pm\text{PsP}\gg\text{C}-\text{P}\cdot\text{Pj}$ $\text{PeP}^\circ\text{C}\mathfrak{H}\text{P}\ddot{\text{P}}\text{P}\text{C}\mathfrak{H}\text{P}^\circ\text{P}\cdot\text{P}\ddot{\text{P}}\text{PSCr}$ C, $\text{P}^\circ\text{P}\mathfrak{M}\text{PsPiPs}$ $\text{PsC}\acute{\text{I}}\text{PSPsPIPSP}\ddot{\text{C}}\dots$
 $\text{P}^\circ\text{PeC}, \text{P}\ddot{\text{P}}\text{PIPSP}\ddot{\text{C}}\dots$ $\text{PjP}\mu\text{C}, \text{P}^\circ\text{P}\pm\text{PsP}\gg\text{C}-\text{C}, \text{C}-\text{PI}$, $\text{PrP}\mu\text{P}\cdot\text{PjP}\mu\text{C}, \text{P}\ddot{\text{P}}\text{P}\gg\text{Cr}-\text{PeP}^\circ\text{C}\mathfrak{H}\text{P}\ddot{\text{P}}\text{P}\text{C}\mathfrak{H}\text{P}^\circ\text{P}\cdot\text{P}\ddot{\text{P}}\text{PSCr}$
(DCAR) C, $\text{P}^\circ\text{PrP}\ddot{\text{P}}\text{PrP}\mu\text{P}\cdot\text{PjP}\mu\text{C}, \text{P}\ddot{\text{P}}\text{P}\gg\text{Cr}-\text{PeP}^\circ\text{C}\mathfrak{H}\text{P}\ddot{\text{P}}\text{P}\text{C}\mathfrak{H}\text{P}^\circ\text{P}\cdot\text{P}\ddot{\text{P}}\text{PSCr}$ (DDCAR), $\text{PiP}\mu\text{C}\mathfrak{H}\text{P}\mu\text{PIP}^\circ\text{P}\mathfrak{I}\text{PSPs}$
 $\text{PsPiPsC}\acute{\text{I}}\text{P}\mu\text{C}\mathfrak{H}\text{P}\mu\text{PrPePsPICrC}''\text{C}, \text{C}\mathfrak{H}\text{C}\acute{\text{I}}\text{C}\mathfrak{I}$ CYP3A4 P· $\text{PjC}-\text{PSC}-\text{PjP}^\circ\text{P}\gg\text{C}\mathfrak{H}\text{PSP}\ddot{\text{P}}\text{Pj}$ $\text{PIPiP}\gg\text{P}\ddot{\text{P}}\text{IPsPj}$
CYP2D6.

P†PSPiC-P±C-C,PsCḤPë CYP3A4

РљРµС,PsPePsPSP°P·PsP», ГГРёР»СНPSPёPNё C-PSPiC-P±C-C,PsCђ CYP3A4,
 ГГРёCђPёC±PёPSCђC” PёC-PrPIPёC%oPµPSPSCђ PIPjC-ГГC,Gr P·P°PiP°P»СНPSPsPiPs
 PeP°CђPёРёPёCђP°P·PёPSCr (PeP°CђPёРёPёCђP°P·PёPSCr C,P° PNёPsPiPs P°PeC,PёPIPSPёC...
 PjPµC,P°P±PsP»C-C,C-PI) PI PёP»P°P·PjC- PI 2 CђP°P·Pё PёCђPsC,CђPiPsPj
 PSPµC,CђPёPIP°P»PsPiPs (4 PrPSC-) ГГGrPёCрC,PSCђPsPiPs P·P°ГГC,PsГГGrPIP°PSPSCђ,
 CђPeC%oPs PICђP°C...PsPICрCђC,СНГГCђ PSPµP·PIбђ™ CђP·P°PSC- P°P±Ps
 PSPµP·PIбђ™ CђP·P°PSC- P·PIбђ™ CђP·P°PSC- PePsPjPёPsPSPµPSC,Pё.

PSPμCΤῖPμP· PrPsPIPiPēPNē PiPμCΤῖC-PsPr PSP°PiC-PIPIPēPIPμPrPμPSPSCΠ P°PeC,PēPIPSPēC...
 PePsPjPiPsPSPμPSC,C-PI PeP°CΤῖPēPiCΤῖP°P·PēPSCr PiPsPrP°P»CHBCEPμ
 PiC-PrPIPēC%oPμPSPSCΠ PIPjC-CiC,Cr P·P°PiP°P»CHBPSPsPiPs PeP°CΤῖPēPiCΤῖP°P·PēPSCr PI
 PiP»P°P·PjC- PjPsP¶Pμ PsC‡C-PeCrPIP°C,PēCΓCΠ PiC-Pr C‡P°CΓ P±C-P»CHBCē
 C,CΤῖPēPIP°P»PsPiPs PePsPjP±C-PSPsPIP°PSPsPiPs PiCΤῖPēPNēPsPjCr. PŷPsPjCr
 PePsPjP±C-PSPsPIP°PSPēPNē PiCΤῖPēPNēPsPj PeP°CΤῖPēPiCΤῖP°P·PēPSCr P· PiPsC,CrP¶PSPēPjPē
 C-PSPiC-P±C-C,PsCΤῖP°PjPē CYP3A4 (PSP°PiCΤῖPēPeP»P°Pr, P±PsC‡PμPiCΤῖPμPIC-CΤῖ,
 PeP»P°CΤῖPēC,CΤῖPsPjC-C‡PēPS, PePsP±C-C‡PēCΓC,P°C,, C-PSPrPēPSP°PIC-CΤῖ,
 C-C,CΤῖP°PePsPSP°P·PsP», PePμC,PsPePsPSP°P·PsP», PSPμC,,P°P·PsPrPsPs,
 PSPμP»C,,C-PSP°PIC-CΤῖ, PiPsP·P°PePsPSP°P·PsP», CΤῖPēC,PsPSP°PIC-CΤῖ,
 CΓP°PePIC-PSP°PIC-CΤῖ, C,PμP»P°PiCΤῖPμPIC-CΤῖ, C,PμP»C-C,CΤῖPsPjC-C‡PēPS,
 PIPsCΤῖPēPePsPSP°P·PsP») PiCΤῖPsC,PēPiPsPeP°P·P°PSPēPNē (PrPēPI. CΤῖPsP·PrC-P»
 B«PμCΤῖPsC,PēPiPsPeP°P·P°PSPSCΠB»).

P·CḥPëC,CḥPSPjC-C†PëPS (500 PjPi PrPIC-C†C- PSP° PrPsP±C†), PïPsPjC-CḥPSPëPñ
C-PSPiC-P±C-C,PSCḥ CYP3A4, PIPëPeP»PëPeP°PI C† C†PµCḥPµPrPSCḥPsPjC† 1,4-
PeCḥP°C,PSPµ (PrC-P°PïP°P·PsPS 1,03Bḥ“2,32 CḥP°P·P°) P·P±C-P»CHC€PµPSPSCµ
PµPeC†PïPsP·PëC†C-C— P·P°PïP°P»CHPSPsPiPs PeP°CḥPëPïCḥP°P·PëPSC† PI PïP»P°P·PjC-
PeCḥPSPiC- PïC-C†P»Cµ 3 C,PëP¶PSC-PI PsPrPSPsC†P°C†PSPsPiPs P·P°C†C,PSC†C†PIP°PSPSCµ.
PŷP°PePëPj C†PëPSPsPj, PïCḥPsC,CµPiPsPj PïPµCḥC-PsPrC† PsPrPSPsC†P°C†PSPsPiPs
P·P°C†C,PSC†C†PIP°PSPSCµ PeP°CḥPëPïCḥP°P·PëPSC† P· PïPsPjC-CḥPSPëPj
C-PSPiC-P±C-C,PSCḥPsPj CYP3A4 (PSP°PïCḥPëPeP»P°Pr, PµCḥPëC,CḥPsPjC-C†PëPSPsPj,
C„P»C†PePsPSP°P·PsP»PsPj, PrPëP»C,C-P°P·PµPjPsPj, PIPµCḥP°PïP°PjC-P»PsPj)
CḥPµPePsPjPµPSP†C†C”C,CHC†Cµ PePsPSC,CḥPsP»CḥPIP°C,Pë C-PSPrPëPIC-PrC†P°P»CHPSC†
CḥPµP°PeC†C-Cḥ C,P° PïPµCḥPµPSPsC†PëPjC-C†C,CH C-, CµPeC‰Ps PSPµPsP±C...C-PrPSPs,
PrPsP·C† PeP°CḥPëPïCḥP°P·PëPSC† C†P»C-Pr (C,PëPjC†P°C†PsPIPs) P·PjPµPSC€PëC,Pë,
PICḥP°C...PsPIC†CḥC†Pë PïPsC,PµPSC†C-PñPSPsPiPµ P·P±C-P»CHC€PµPSPSCµ
PµPeC†PïPsP·PëC†C-C—. P’PSP°C†P»C-PrPsPe C,CḥPëPIP°P»PsPiPs PïPµCḥC-PsPrC†
PSP°PïC-PIPIPëPIPµPrPµPSPSCµ PeP°CḥPëPïCḥP°P·PëPSC† C,P° PñPsPiPs P°PeC,PëPIPSPëC...
PjPµC,P°P±PsP»C-C,C-PI PïPsC†P°C,PSPe, PïCḥPëPïPëPSPµPSPSCµ P»C-PeC†PIP°PSPSCµ
PïPsPjC-CḥPSPëPj C-PSPiC-P±C-C,PSCḥPsPj CYP3A4 P°P±Ps P·PjC-PSP° PrPsP·Pë
PïPsPIPSC-C†C,Cḥ PSPµ PIC-PrPsP±CḥP°P·PëC,CHC†Cµ PSP° CḥC-PIPCµC...

Pj PrPsCfP»C-PrP¶PµPSPSC- PIP·P°C”PjPsPrC— P»C-PeP°CĤCfCHbPePëC... P·P°CfPs±C-PI
28-PrPµPSPSPµ P»C-PeCfPIP°PSPSC¶ PeP°CĤPëPîCĤP°P·PëPSPsPj Cr PrPsP·C- 6 PjPi PSP°
PrPsP±Cf PSPµ PjP°P»Ps PeP»C-PSC-C†PSPs P·PSP°C†CrC%oPsPiPs PIPiP»PëPICf PSP°
C,,P°CĤPjP°PePsPeC-PSPµC,PëPeCf PîPµCĤPsCĤP°P»CHbPSPëC... PePsPSC,CĤP°C†PµPîC,PëPIC-PI
(PµC,PëPSC-P»PµCfC,CĤP°PrC-PsP»Cr C,P° P»PµPIPsPSPsCĤPiPµCfC,CĤPµP»Cr).

РѹС-Р·PSCЇ РrРёСГРёС-PSРµР·С-СЇ вѢ” С†Рµ СГРёPSPPrСѢPsPj, C%Ps
PіСѢPsСЇPIР»СЇС”C,СѢСГСЇ СѢРёС,PjC-С†PSPёPjPё PjPёPjPsPIC-P»СѢPSPёPjPё
СѢСГС...P°PjPё PіРµСѢРµPIР°P¶PSPs СЇР·РёРёР° С,P°/P°P±Ps PsP±P»РёС†С†СЇ С- PjPsP¶Рµ
СѢPsP·PIРёPIР°C,РёСГСЇ СГ РіР°С†С-С”PSC,C-PI, СЇРёС- P»C-РёСГСѢC,СѢСГСЇ
P°PSC,РёPіСГРёС...PsC,РёРёР°PjPё. PіРёC%Ps СГРёPjPіC,PsPjPё PіC-Р·PSCѢPsC—
РrРёСГРёС-PSРµР·С-С— Р·вѢ™ СЇPIР»СЇСѢC,СѢСГСЇ СГ РіР°С†С-С”PSC,P°, СЇРёРёPNё
P»C-РёСГС”C,СѢСГСЇ РёР°СѢРёPіСѢР°P·РёPSPsPj, PіPsC,СѢC-Р±PSPs СѢPsP·PіP»СЇPSCГС,Рё
PSРµPsP±C...C-PrPSC-СГС,СѢ PіСѢРёPіРёPSPµPSPSCЇ B P·P°СГС,PsСГСГPIР°PSPSCЇ
С†СѢPsPіPs PіСѢРµPіР°СѢР°C,СГ.

РГPIPsСѢPsP±P° РµР°СѢРёС-PSCГPsPSP°

PіРёC%Ps P°PSC,РёPіСГРёС...PsC,РёС†PSC- P·P°СГPsP±Pё PіСѢРёP·PSP°С†Р°СѢC,СѢСГСЇ
PіР°С†С-С”PSC,P°Pj C-Р· C...PIPsСѢPsP±PsСѢ РµР°СѢРёС-PSCГPsPSP°, C,Ps PjPsP¶P»РёPIРµ
P·P°PіPsСГС,СѢРµPSPSCЇ C„PsPSPsPіPsPіPs P·P°C...PIPsСѢСѢPIР°PSPSCЇ C,P°
PіPsPіC-СѢСРµPµPSPSCЇ PіРµСѢРµP±C-PіСГ C...PIPsСѢPsP±Pё РµР°СѢРёС-PSCГPsPSP°.
PŷPsPjСГ, PіСѢРёP·PSP°С†Р°СѢРё РёР°СѢРёPіСѢР°P·РёPS PіР°С†С-С”PSC,P°Pj C-Р·
C...PIPsСѢPsP±PsСѢ РµР°СѢРёС-PSCГPsPSP°, P»C-РёР°СѢC- PjP°СѢC,СѢ P·PIР°P¶СГPIР°C,Рё
СГРіC-PIPIC-PrPSPsСРµPµPSPSCЇ РёPsСѢРёСГС,СѢ/СѢРёP·РёРё.

РŷРёPjPіC,PsPjPё P· P±PsРёСГ PsC†PµPNё/ РёР°C,P°СѢР°РёC,P°

Pj PrPsРёP»C-PSC-С†PSPёC... PrPsСГР»C-PrP¶PµPSPSCЇC... РёР°СѢРёPіСѢР°P·РёPSCГ PI
СГPsP±P°Рё P±СГР»Ps PіPsPjC-С†PµPSPs PіPsPjСГC,PSC-PSPSCЇ РёСѢРёСРёC,P°P»РёРёР°/
РёР°C,P°СѢР°РёC,P° (PrРёPI. СѢPsP·PrC-P»Pё B«PµPsP±C-С†PSC- СѢРµP°РёС†С-С—B» C,P°
B«P”P°PSC- PrPsРёP»C-PSC-С†PSPёC... PrPsСГР»C-PrP¶PµPSCѢ P· P±PµP·PіРµРёPёB»).

РµСѢРёС†PёPSPSPs-PSP°СГР»C-PrРёPsPIРёPNё P·PівѢ™ СЇР·PsРё PjC-P¶ P·PjC-PSP°PjPё PI
РёСѢРёСРёC,P°P»РёРёСГ/ РёР°C,P°СѢР°РёC,PsСѢ, C%Ps СГРіPіPsСГС,РµСѢC-PіР°P»РёСГСЇ
PіC-Pr С†Р°СГ PrPsСГР»C-PrP¶PµPSCѢ СГ P»СѢPrPµPNё, C- P·P°СГС,PsСГСГPIР°PSPSCЇPj
РёР°СѢРёPіСѢР°P·РёPSCГ PSPµ P±СГPI СГСГС,P°PSPsPIР»PµPSPёPNё. РµСѢPsC,Рµ
PіР°С†С-С”PSC,P°Pj, СГ СЇРёРёС... PjPsP¶СГC,СѢ СѢPsP·PIРёPSCГC,РёСГСЇ СГРёPjPіC,PsPjPё,
C%Ps PіPsC,РµPSC†C-PNёPSPs PjP°СѢC,СѢ PIC-PrPSPsСРµPµPSPSCЇ PrPs РёР°C,P°СѢР°РёC,Рё,
PіPsC,СѢC-Р±PSPs PіPsСѢР°PrРёC,Рё PіСѢPsPNёC,Рё PsC„C,P°P»СѢPjPsP»PsPіC-С†PSPµ
PsP±СГС,РµP¶PµPSPSCЇ, P° PіPsC,C-Pj СГР»C-Pr PіPsPIC,PsСѢPSPs B СѢPsP·PіP»СЇPSCГC,Рё
PіРёC,P°PSPSCЇ PіСѢPs PіСѢPsPrPsPIР¶PµPSPSCЇ P»C-РёСГPIР°PSPSCЇ.

P—P»PsСЇРёС-СГPSPёPNё PSPµPNёСѢPsP»PµPіC,РёС†PSPёPNё СГРёPSPPrСѢPsPj
РµPsC,РµPSC†C-PNёPSPs P»PµC,P°P»СѢPSPёPNё РёPsPjPіP»PµРёСГ СГРёPjPіC,PsPjC-PI, C%Ps
PjP°C” PSP°P·PICГ P·P»PsСЇРёС-СГPSPёPNё PSPµPNёСѢPsP»PµPіC,РёС†PSPёPNё СГРёPSPPrСѢPsPj
(P—PќPŷ), PIC-PrP·PSP°С†Р°C”C,СѢСГСЇ PіСѢРё P»C-РёСГPIР°PSPSC—
P°PSC,РёPіСГРёС...PsC,РёРёР°PjPё. РљP»C-PSC-С†PSPёPjPё PіСѢPsСЇPIР°PjPё P—PќPŷ C”
PіC-PіРµСѢPіC-СѢРµРёСГС-СЇ, PjвѢ™ СЇР·PsPIР° СѢРёPіC-PrPSC-СГС,СѢ,
PіC-PrPIРёC%PµPSPёPNё СѢC-PIРµPSCѢ РёСѢРµP°C,РёPSC-PSC„PsСГС„PsРёC-PSP°P·Рё PI
СГРёСѢPsPIР°C,С†C-, P·PjC-PSPµPSPёPNё PіСГРёC...C-С†PSPёPNё СГС,P°PS C,P° PsP·PSP°РёРё
СѢPsP·P»P°PrСГ PIРµPіРµC,P°C,РёPIРSPsC— PSPµСѢPIPsPIPsC— СГРёСГС,РµPjPё
(PSPµСѢРµPіСГP»СЇСѢPSPёPNё PіСГP»СѢСГ P°P±Ps P°СѢC,РµСѢC-P°P»СѢPSPёPNё C,РёСГРё,
C,P°C...C-РёР°СѢPrC-СЇ, PrC-P°C„PsСѢРµP· C,P° СГРµСѢС†PµPIР° P°СѢРёC,PjC-СЇ).

P”PsPrP°C,РёPsPIC- PsP·PSP°РёРё PjPsP¶СГC,СѢ PIРёP»СѢС†P°C,Рё
PjC-PsPіP»PsP±C-PSCГСѢC-СѢ (СѢР°P±PrPsPjC-PsP»C-P·) C,P° PіPsСГС,СѢСГ PSPёСѢРёPsPICГ
PSPµPrPsСГС,P°C,PSC-СГС,СѢ. PіРёC%Ps СГ РіР°С†С-С”PSC,P° СѢPsP·PIРёPIР°СѢC,СѢСГСЇ
СГРёPjPіC,PsPjPё, C%Ps PIРёР°P·СГСѢC,СѢ PSP° P—PќPŷ, P°P±Ps СГ PSCѢPsPіPs
P·вѢ™ СЇPIР»СЇС”C,СѢСГСЇ PIРёСГPsРёР° C,РµPjPіРµСѢР°C,СГСѢР° P· PSPµPIC-PrPsPjPёC...

PiCḡPēC†PēPS P±PμP· PrPsPrP°C,PePsPIPēC... PeP»C-PSC-C†PSPēC... PiCḡPsCḡPIC-PI P—PḡPŸ,
P·P°C†C-,PsC†C†PIP°PSPSCḡ PeP°CḡPēPiCḡP°P·PēPSC† PjP°C” P±C†C,Pē PSPμPiP°PNḡPSPs
PiCḡPēPiPēPSPμPSPμ.

PŸC†PrPsPjPē C,P° PePsPSPIC†P»CHC†C-C—

PḡP°CḡPēPiCḡP°P·PēPS PiPsC,CḡC-P±PSPs PsP±PμCḡPμP¶PSPs PiCḡPēP·PSP°C†P°C,Pē
PiP°C†C-C”PSC,P°Pj, C† CḡPePēC... PI P°PSP°PjPSPμP·C- C” C†C†PrPsPjPē P°P±Ps
PiP°C,PsP»PsPiC-C†PSC- C†C,P°PSPē, C°Ps PiPsC,PμPSC†C-PNḡPSPs P·PSPēP¶C†CḡC,CH
C†C†PrPsPjPSPēPNḡ PiPsCḡC-Pi.

PμP°C†C-C”PSC,Pē P»C-C,PSCHPsPiPs PIC-PeC† P· PrPμPjPμPSC†C-C”Cḡ

P”C-Cḡ PeP°CḡPēPiCḡP°P·PēPSC† C† PiP°C†C-C”PSC,C-PI P»C-C,PSCHPsPiPs PIC-PeC† P·
PrPμPjPμPSC†C-C”Cḡ PSPμ P±C†P»P° PIPēPIC†PμPSP°, C,PsPjC† P»C-PeP°CḡC†CHPePēPNḡ
P·P°C†C-P± PSPμ CḡPμPePsPjPμPSP†C”C,CHC†Cḡ PrPs P·P°C†C,PsC†C†PIP°PSPSCḡ
C,P°PePēPj PiP°C†C-C”PSC,P°Pj C†PμCḡPμP· PiC-PrPIPēC°PμPSPēPNḡ CḡPēP·PēPe
P·P°PiP°P»CHPSPsC— P»PμC,P°P»CHPSPsC†C,C-.

P PēP·PēPe PIPēPSPēPePSPμPSPSCḡ PiPsCḡC†C€PμPSCH PjPsP·PePsPIPsPiPs
PeCḡPsPIPsPsP±C-PiC† (PμPḡPḡ)

Pj CḡP°PSP†PsPjC-P·PsPIP°PSPēC... PiP»P°C†PμP±PsPePsPSC,CḡPsP»CHPsPIP°PSPēC...
PeP»C-PSC-C†PSPēC... PrPsC†P»C-PrP¶PμPSPSCḡC... C† PiCḡC†PiC- PiP°C†C-C”PSC,C-PI C-P·
PrPμPjPμPSC†C-C”Cḡ, CḡPeC- P»C-PeC†PIP°P»PēC†Cḡ PrPμCḡPePēPjPē P°C,PēPiPsPIPēPjPē
P°PSC,PēPiC†PēC...PsC,PēPeP°PjPē, P±C†P»Ps PiPsPjC-C†PμPSPs, C°Ps CḡPēP·PēPe
PIPēPSPēPePSPμPSPSCḡ PμPḡPḡ P·P±C-P»CHC€PμPSPēPNḡ PiCḡPēP±P»PēP·PSPs PI C,CḡPē
CḡP°P·Pē. PḡPμC...P°PSC-P·Pj PiC-PrPIPēC°PμPSPSCḡ CḡPēP·PēPeC† PSPμPIC-PrPsPjPēPNḡ.
PμC-PrPIPēC°PμPSPēPNḡ CḡPēP·PēPe PμPḡPḡ PSPμ PjPsP¶PSP° PIPēPeP»CḡC†P°C,Pē PiCḡPē
PiCḡPēPNḡPsPjC- C-PSC€PēC... P°PSC,PēPiC†PēC...PsC,PēPeC-PI P°P±Ps PrP»Cḡ C-PSC€PēC...
PiCḡC†Pi PiP°C†C-C”PSC,C-PI. PḡP°CḡPēPiCḡP°P·PēPS PiPsC,CḡC-P±PSPs PsP±PμCḡPμP¶PSPs
PiCḡPēP·PSP°C†P°C,Pē PiP°C†C-C”PSC,P°Pj P· C†PēPSPSPēPeP°PjPē CḡPēP·PēPeC†
PIPēPSPēPePSPμPSPSCḡ C-PSC†C†P»CHC,C†.

P—P°C...PIPsCḡCHPIP°PSPSCḡ C†PμCḡC†PμPIPs-C†C†PrPēPSPSPsC— C†PēC†C,PμPjPē

P—PjC-PSPē P°CḡC,PμCḡC-P°P»CHPSPsPiPs C,PēC†PeCž

PḡP°CḡPēPiCḡP°P·PēPS PjPsP¶Pμ C†PiCḡPēC†PēPSCḡC,Pē CḡPe PsCḡC,PsC†C,P°C,PēC†PSC†
PiC-PiPsC,PμPSP·C-Cḡ, C,P°Pe C- PiC-PiPμCḡC,PμPSP·C-Cḡ (PrPēPI. CḡPsP·PrC-P»
B«PμPsP±C-C†PSC- CḡPμP°PeC†C-C—B»). PḡP°CḡPēPiCḡP°P·PēPS PiPsC,CḡC-P±PSPs
PsP±PμCḡPμP¶PSPs P·P°C†C,PsC†PsPIC†PIP°C,Pē PiP°C†C-C”PSC,P°Pj, C† CḡPePēC... PI
P°PSP°PjPSPμP·C- C” P·P°C...PIPsCḡCHPIP°PSPSCḡ C†PμCḡC†PμPIPs-C†C†PrPēPSPSPsC—
C†PēC†C,PμPjPē C- CḡPeC- C†C...PēP»CHPSC- PrPs P·PjC-PS P°CḡC,PμCḡC-P°P»CHPSPsPiPs
C,PēC†PeC†. PμPsC,CḡC-P±PSPs B PePsPSC,CḡPsP»CHPIP°C,Pē P°CḡC,PμCḡC-P°P»CHPSPēPNḡ
C,PēC†Pe.

P—PjC-PSPē PSP° PμP»PμPeC,CḡPsPeP°CḡPrC-PsPiCḡP°PjC- (P·PḡP”)

Pj PiP°C†C-C”PSC,C-PI, CḡPeC- PiCḡPēPNḡPjP°CḡC,CH P°PSC,PēPiC†PēC...PsC,PēPePē, PjPsP¶Pμ
CḡPsP·PIPēPIP°C,PēC†Cḡ PiPsPrPsPIP¶PμPSPSCḡ C-PSC,PμCḡPIP°P»C† QPŸ.

Pj PeP»C-PSC-C†PSPsPjC† PrPsC†P»C-PrP¶PμPSPSC-, CḡPePμ PiCḡPsPIPsPrPēP»Pē PrP»Cḡ

PsC†C-PSPePë PîPsPrPsPIP¶PµPSPSC‡ C-PSC,PµC‡PIP°P»C† QT, PîC‡Pë PîC‡PëPNëPsPjC-
 PeP°C‡PëPîC‡P°P·PëPSC† PSPµ P±C†P»Ps PîPsPjC-C†PµPSPs P¶PsPrPSPsPiPs
 PîPsPrPsPIP¶PµPSPSC‡ C-PSC,PµC‡PIP°P»C† QT C† PîPsC‡C-PIPSC‡PSPSC- P· PîP»P°C†PµP±Ps
 (PrPëPI. C‡PsP·PrC-P» B«PαP°C‡PjP°PePsPrPëPSP°PjC-PeP°B»). PJ PeP»C-PSC-C†PSPëC...
 PrPsC†P»C-PrP¶PµPSPSC‡C... PîPsPIC-PrPsPjP»C‡P»PsC†C‡ P»PëC€Pµ PîC‡Ps
 PrPµPeC-P»C‡PeP° PIPëPîP°PrPeC-PI PSPµP·PSP°C†PSPsPiPs PîPsPrPsPIP¶PµPSPSC‡
 C-PSC,PµC‡PIP°P»C† QT PîC‡Pë PîC‡PëPNëPsPjC- PeP°C‡PëPîC‡P°P·PëPSC† (PrPëPI.
 C‡PsP·PrC-P» B«PµPsP±C-C†PSC- C‡PµP°PeC†C-C—B»). PÿPsPjC† PeP°C‡PëPîC‡P°P·PëPS
 C,C‡PµP±P° PsP±PµC‡PµP¶PSPs P·P°C†C,PsC†PsPIC†PIP°C,Pë PîP°C†C-C”PSC,P°Pj, C†
 C‡PePëC... PI P°PSP°PjPSPµP·C- C” C†PµC‡C†PµPIPs-C†C†PrPëPSPSPµ
 P·P°C...PIPsC‡C‡PIP°PSPSC‡ P°P±Ps PîPsPrPsPIP¶PµPSPSC‡ C-PSC,PµC‡PIP°P»C† QT PI
 C†C-PjPµPNëPSPsPjC† P°PSP°PjPSPµP·C-, P° C,P°PePsP¶ PîP°C†C-C”PSC,P°Pj, C‡PeC-
 P»C-PeC†PIP°P»PëC†C‡ PîC‡PµPîP°C‡P°C,P°PjPë, C%oPs PjPsP¶C†C,C‡ PîPsPrPsPIP¶PëC,Pë
 C-PSC,PµC‡PIP°P» QT (PrPëPI. C‡PsP·PrC-P» B«PαP°C‡PjP°PePsPrPëPSP°PjC-PeP°B»).

P’PµPSPsP·PSP° C,C‡PsPjP±PsPµPjP±PsP»C-C‡ (P’PÿP·)

PµPsPIC-PrPsPjP»C‡P»PsC†C‡ PîC‡Ps PIPëPîP°PrPePë PIPëPSPëPePSPµPSPSC‡ P’PÿP· PîC‡Pë
 P·P°C†C,PsC†C†PIP°PSPSC- P°PSC,PëPîC†PëC...PsC,PëC†PSPëC... P·P°C†PsP±C-PI.
 P‡C†PeC-P»C‡PePë C† PîP°C†C-C”PSC,C-PI, C‡PeC- PîC‡PëPNëPjP°C‡C,C‡
 P°PSC,PëPîC†PëC...PsC,PëPePë, C†P°C†C,Ps PIPëPSPëPeP°C‡C,C‡ PSP°P±C†C,C-
 C†PëPSPSPëPePë C‡PëP·PëPeC† C‡PsP·PIPëC,PeC† P’PÿP·, PIC†C- PjPsP¶P»PëPIC-
 C†PëPSPSPëPePë PIPëPSPëPePSPµPSPSC‡ P’PÿP· C†P»C-Pr PIPëC‡PIPëC,Pë PîPµC‡PµPr C,P°
 PîC-Pr C†P°C† P»C-PeC†PIP°PSPSC‡ PeP°C‡PëPîC‡P°P·PëPSPsPj, P° C,P°PePsP¶ C,C‡PµP±P°
 PîC‡PsPIPsPrPëC,Pë C—C...PSC‡ PîC‡PsC,,C-P»P°PeC,PëPeC†.

P”C-PîPµC‡PiP»C-PePµPjC-C‡ C,P° C†C†PeC‡PsPIPëPNë PrC-P°P±PµC,

PJ PîP°C†C-C”PSC,C-PI C-P· C†C†PeC‡PsPIPëPj PrC-P°P±PµC,PsPj P°P±Ps C†
 PîP°C†C-C”PSC,C-PI C-P· C†PëPSPSPëPeP°PjPë C‡PëP·PëPeC† C‡PsP·PIPëC,PeC†
 C†C†PeC‡PsPIPsPiPs PrC-P°P±PµC,C† (PSP°PîC‡PëPeP»P°Pr, PsP¶PëC‡C-PSPSC‡,
 PSP°C‡PIPSC-C†C,C‡ PrC-P°P±PµC,C† PI C†C-PjPµPNëPSPsPjC† P°PSP°PjPSPµP·C-), C‡PeC-
 PîPsC†PëPSP°C‡C,C‡ P»C-PeC†PIP°PSPSC‡ P°C,PëPîPsPIPëPjPë
 P°PSC,PëPîC†PëC...PsC,PëPeP°PjPë, C†P»C-Pr PePsPSC,C‡PsP»C‡PIP°C,Pë C‡C-PIPSC-
 PiP»C‡PePsP·Pë PI C†PëC‡PsPIP°C,C†C-. PµC-Pr C†P°C† PeP»C-PSC-C†PSPëC...
 PrPsC†P»C-PrP¶PµPSC‡ PeP°C‡PëPîC‡P°P·PëPSC† PîPsPIC-PrPsPjP»C‡P»PsC†C‡ PîC‡Ps
 PîPsP±C-C†PSC- C‡PµP°PeC†C-C— C†C,PsC†PsPIPSPs C‡C-PIPSC-PI PiP»C‡PePsP·Pë (PrPëPI.
 C‡PsP·PrC-P» B«PαP°C‡PjP°PePsPrPëPSP°PjC-PeP°B»).

P—PjC-PSPë PjP°C†Pë C,C-P»P°

PµC‡Pë PîC‡PëPNëPsPjC- PeP°C‡PëPîC‡P°P·PëPSC† C†PîPsC†C,PµC‡C-PiP°P»PsC†C‡
 P·PSP°C†PSPµ P·P±C-P»C‡C€PµPSPSC‡ PjP°C†Pë C,C-P»P°. PµP°C†C-C”PSC,P°Pj C†P»C-Pr
 C‡PµPiC†P»C‡C‡PSPs PePsPSC,C‡PsP»C‡PIP°C,Pë PjP°C†C† C,C-P»P° (PrPëPI. C‡PsP·PrC-P»
 B«PµPsP±C-C†PSC- C‡PµP°PeC†C-C—B»).

P‡PrPSPsC†P°C†PSPµ P»C-PeC†PIP°PSPSC‡ PîPsPjC-C‡PSPëPjPë C-PSPiC-P±C-C,PsC‡P°PjPë
 CYP3A4

P‡PrPSPsC†P°C†PSPµ P·P°C†C,PsC†C†PIP°PSPSC‡ PeP°C‡PëPîC‡P°P·PëPSC† P·
 PîPsPjC-C‡PSPëPjPë C-PSPiC-P±C-C,PsC‡P°PjPë CYP3A4 PjPsP¶Pµ PîC‡PëP·PIPµC†C,Pë PrPs
 P·P±C-P»C‡C€PµPSPSC‡ P·P°PiP°P»C‡PSPsC— PµPeC†PîPsP·PëC†C-C—

PeP°CḡPēPīCḡP°P·PēPSCṛ. P PμPePsPjPμPSPṛCṛC" C,CHCṛCḷ PjPsPSC-C,PsCḡPēPSPi
C-PSPṛPēPIC-PrCṛP°P»CHBPSPsC— CḡPμP°PeC†C-C— C,P° PīPμCḡPμPSPsCṛPēPjPsCṛC,C- C,P°,
CḷPeC%Ps PSPμPsP±C...C-PrPSPs, PrPsP·Cṛ PeP°CḡPēPīCḡP°P·PēPSCṛ CṛP»C-Pr
(C,PēPjC†P°CṛPsPIPs) P·PjPμPSCēPēC,Pē P· CṛCḡP°C...CṛPIP°PSPSCḷPj
PīPsC,PμPSC†C-PNēPSPsPiPs P·P±C-P»CHCēPμPSPSCḷ PμPeCṛPīPsP·PēC†C-C— (PrPēPI.
CḡPsP·PrC-P» B«P'P·P°C" PjPsPrC-Cḷ P· C-PSCēPēPjPē P»C-PeP°CḡCṛCHPePēPjPē
P·P°CṛPsP±P°PjPē C,P° C-PSCēC- PIPēPrPē PIP·P°C" PjPsPrC-PNēB»).

P"PsPīPsPjC-PḡPSC- CḡPμC†PsPIPēPSPē

P PμP°PiC-P»P°, PeP°PīCṛCṛP»Pē C,PIPμCḡPrC- PīPs 3 PjPi, 4,5 PjPi C,P° 6 PjPi, PjC-CṛC,CḷC,CH
P±P°CḡPIPSPēPe CṛPīPμC†C-P°P»CHBPSPēPNē C†PμCḡPIPsPSPēPNē PḡPṽ (P• 129), C%Ps
PjPsPḡPμ CṛPīCḡPēC†PēPSCḷC,Pē P°P»PμCḡPiC-C†PSC- CḡPμP°PeC†C-C—.

*P—P°CṛC,PsCṛCṛPIP°PSPSCḷ Cṛ PīPμCḡC-PsPr PIP°PiC-C,PSPsCṛC,C- P°P±Ps
PiPsPrCṛPIP°PSPSCḷ PiCḡCṛPrCḡ.*

P-C-PSPePē CḡPμPīCḡPsPrCṛPeC,PēPIPSPsPiPs PIC-PeCṛ/ PḡC-PSPePē, CḷPeC-
PIPēPePsCḡPēCṛC,PsPICṛCḡC,CH PePsPSC,CḡP°C†PμPīC†C-Cḡ

P-C-PSPeP°Pj CḡPμPīCḡPsPrCṛPeC,PēPIPSPsPiPs PIC-PeCṛ PīPsC,CḡC-P±PSPs CṛPSPēPeP°C,Pē
PIP°PiC-C,PSPsCṛC,C- PiC-Pr C†P°Cṛ PīCḡPēPNēPsPjCṛ P»C-PeP°CḡCṛCHPePsPiPs
P·P°CṛPsP±Cṛ P PμP°PiC-P»P°. PμP°C†C-C" PSC,PePē CḡPμPīCḡPsPrCṛPeC,PēPIPSPsPiPs
PIC-PeCṛ PjP°CḡC,CH PIPēPePsCḡPēCṛC,PsPICṛPIP°C,Pē PIPēCṛPsPePsPμC,,PμPeC,PēPIPSC-
PjPμC,PsPrPē PePsPSC,CḡP°C†PμPīC†C-C— PiC-Pr C†P°Cṛ P»C-PeCṛPIP°PSPSCḷ C,P°
C%PsPSP°PNēPjPμPSCēPμ PīCḡPsC,CḷPiPsPj 10 C,PēPḡPSC-PI PiC-CṛP»Cḷ PīCḡPēPNēPsPjCṛ
PsCṛC,P°PSPSCHPsC— PrPsP·Pē PīCḡPμPīP°CḡP°C,Cṛ P PμP°PiC-P»P°.

P'P°PiC-C,PSC-CṛC,CH

P"°PSC- C%PsPrPs P·P°CṛC,PsCṛCṛPIP°PSPSCḷ PeP°CḡPēPīCḡP°P·PēPSCṛ PIP°PiC-C,PSPēPj
PḡC-PSPeP°Pj PIC-PrCṛCṛC,PSC- P°P±Ps PsP±PjPμPḡPμPSC-.

P"PsCṛP»C-PrPḡPμPSPSCḷ PSP° C,PIP°CḡPēPSP°C... PīPsPeP°P·P°P»Pē
CḡPμPīCḡPsPrCṛPeC,PēPIPSṛ C,PsPeCṛPēC†PSC-CṛC,CH, PIPeP»CḡC†PSPs P· PIP°PrP°PjPē
CḡPsP·PIPēC,PeCṛ Cṛ C%CṛCḡC-PI (PrPēPI. CḡPsP·PrC-P» B«P"°PSC-
PrPsPeP»C-PSC-C†PSPēC... PrPsCṛP»C-PrPḡPμPSCḡ P· P±PμP·PīPμPePēB»).

P—P°CṛC,PsCṛCṛPIP°PSPSCḷ PīCḡPμPīP°CḡP°C,Cṛ P PμP°PiC-P»P° PSPμ
CḡPμPePsPjPμPSPṛCṛC" C,CHCṛCḷ PīCḡPsC,CḷPiPsPj PIP°PiC-C,PSPsCṛC,C-, P° C,P°PePsPḡ
PḡC-PSPeP°Pj CḡPμPīCḡPsPrCṛPeC,PēPIPSPsPiPs PIC-PeCṛ, CḷPeC- PSPμ
PIPēPePsCḡPēCṛC,PsPICṛCḡC,CH PμC,,PμPeC,PēPIPSCṛ PePsPSC,CḡP°C†PμPīC†C-Cḡ.
PμC-CṛP»Cḷ PīCḡPēPīPēPSPμPSPSCḷ P»C-PeCṛPIP°PSPSCḷ PeP°CḡPēPīCḡP°P·PēPSPsPj
C†PμCḡPμP· PīPsPIC-P»CHBPSPμ PIPēPIPμPrPμPSPSCḷ P°PeC,PēPIPSPēC...
PePsPjPīPsPSPμPSC,C-PI PjPμC,PsPrPē PePsPSC,CḡP°C†PμPīC†C-C— PīPsC,CḡC-P±PSPs
P·P°CṛC,PsCṛPsPICṛPIP°C,Pē C%PsPSP°PNēPjPμPSCēPμ PīCḡPsC,CḷPiPsPj 10 C,PēPḡPSC-PI.

PkPsPIPsPSP°CḡPsPrPḡPμPSC-, PSP° CḷPePēC... PIPiP»PēPIP°P»Pē
P°PSC,PēPīCṛPēC...PsC,PēPePē (PIPeP»CḡC†PSPs P· PeP°CḡPēPīCḡP°P·PēPSPsPj) PīC-Pr C†P°Cṛ
C,CḡPμC,CHPsPiPs C,CḡPēPjPμCṛC,CḡCṛ PIP°PiC-C,PSPsCṛC,C-, P·PSP°C...PsPrCḷC,CHCṛCḷ
PI PiCḡCṛPīC- CḡPēP·PēPeCṛ CḡPsP·PIPēC,PeCṛ C,P°PePēC... PīPsP±C-C†PSPēC...

C₇H₁₂P^oPeC†C-PM₂, C₁₂He P₁₂PeC†C, C₇H₁₂P^oPiC-C₇H₁₂P^oPjC-PrPSC- C†PëPjPiC, PsPjPë C, P^o/P^oP±Ps
 C†PëPjPiC, PsPjPë PIC-PrPjC-PSPë, C₁₂HeC- PjPsP¶C†C, CH₂ P±C†C, Pë C₇H₁₂-P·PSPsC—
 C, C₁₂He¶PePsC†C, C- C, P^o C, C₇H₁₂PëPIP^oP»PsC†C, C-. P₁₂PsPIC-PrPsPjP»C₁₂He»PsC†C₁₂ PiC₇H₁₂Ps
 P·P±C†PrP¶P₁₂PSPSC₁₂, PiC-PiP₁₂C₇H₁₂, PsPSC-C₇H₁₂, PiC-PiPsC, PsPSC-C₇H₁₂, C, C₇H₁₂P₁₂PjPsC₇,
 C†PsPSP»PëPIC-C†C, CH₂, C₇H₁₂P₁₂C†PiC-C₇H₁₂P^oC, PsC₇H₁₂PSPëPM₂ PrPëC†C, C₇H₁₂P₁₂C†-C†PëPSP₁₂C₇H₁₂PsPj
 C, P^o PiPsC₇H₁₂C†C¶P₁₂PSPSC₁₂ PIP¶PëPIP^oPSPSC₁₂ C—P¶C-. P†C- C†C†PeP»P^oPrPSP₁₂PSPSC₁₂
 C₇H₁₂-P·PSPëP»PëC†C₁₂ P·P^o C, C₁₂He¶PeC-C†C, CH₂; C₁₂HeC%Ps C† PrP₁₂C₁₂HePëC...
 PIPëPiP^oPrPeP^oC... C†PëPjPiC, PsPjPë P·PSPëPeP^oP»Pë C†P^oPjC- C†PsP±PsC₇H₁₂, C, Ps PI
 C-PSCëPëC... PIPëPiP^oPrPeP^oC... PSPsPIPsPSP^oC₇H₁₂PsPrP¶P₁₂PSC- PiPsC, C₇H₁₂P₁₂P±C†PIP^oP»Pë
 P»C-PeC†PIP^oPSPSC₁₂ C† PIC-PrPrC-P»P₁₂PSPSC- C-PSC, P₁₂PSC†PëPIPSPsC— C, P₁₂C₇H₁₂P^oPiC-C—
 C, P^o C, C₇H₁₂PëPIP^oP»PsC— PiPsC†PiC-C, P^oP»C-P·P^oC†C-C—. P₇PsPjC† P·P^o C†C, P^oPSPsPj
 PSPsPIPsPSP^oC₇H₁₂PsPrP¶P₁₂PSPëC... PiPsC, C₇H₁₂-P±PSPs C₇H₁₂P₁₂C, P₁₂P»CH₂PSPs
 C†PiPsC†C, P₁₂C₇H₁₂-PiP^oC, Pë.

P"PsPrC†PIP^oPSPSC₁₂ PiC₇H₁₂C†PrPrC₇

P"PsC†C- PSP₁₂PIC-PrPsPjPs, C†Pë PIPëPrC-P»C₁₂HeC, CH₂C†C₁₂ PeP^oC₇H₁₂PëPiC₇H₁₂P^oPëPS C, P^o
 PM₂PsPiPs PsC†PSPsPIPsC- PjP₁₂C, P^oP±PsP»C-C, Pë P· PjPsP»PsPePsPj P»C₇H₁₂PrPëPSPë.
 P₇P^oC₇H₁₂PëPiC₇H₁₂P^oPëPS C, P^o PM₂PsPiPs PjP₁₂C, P^oP±PsP»C-C, Pë PIPëPrC-P»C₁₂HeC, CH₂C†C₁₂ P·
 PjPsP»PsPePsPj C%C†C₇H₁₂-PI PiC-Pr C†P^oC† P»P^oPeC, P^oC†C-C— (PrPëPI. C₇H₁₂PsP·PrC-P»
 B«P"°PSC- PrPsPeP»C-PSC-C†PSPëC... PrPsC†P»C-PrP¶P₁₂PSCH₂ P· P±P₁₂P·PiP₁₂PePëB»). P₁₂P₁₂
 PjPsP¶PSP^o PIPëPeP»C₇H₁₂C†P^oC, Pë PIPiP»PëPIC† PSP^o
 PSPsPIPsPSP^oC₇H₁₂PsPrP¶P₁₂PSPëC.../PSP₁₂PjPsPIP»C₁₂He, P₁₂C-Pr C†P^oC† P»C-PeC†PIP^oPSPSC₁₂
 PeP^oC₇H₁₂PëPiC₇H₁₂P^oPëPSPsPj PiPsC, C₇H₁₂-P±PSPs PiP₁₂C₇H₁₂PiP^oC, Pë PiPsPrC†PIP^oPSPSC₁₂
 PiC₇H₁₂C†PrPrC₇H₁₂.

P₁₂P₁₂C₇H₁₂, PëP»CH₂PSC-C†C, CH₂

P'PiP»PëPI PeP^oC₇H₁₂PëPiC₇H₁₂P^oPëPSC† PSP^o C, P₁₂C₇H₁₂, PëP»CH₂PSC-C†C, CH₂ P»C₇H₁₂PrPëPSPë PSP₁₂
 P±C†PI PIPëPIC†P₁₂PSPëPM₂. Pj PrPsC†P»C-PrP¶P₁₂PSPSC₁₂... PSP^o C%C†C₇H₁₂P^oC...
 C†PiPsC†C, P₁₂C₇H₁₂-PiP^oP»PsC†C₁₂ P·PSPëP¶P₁₂PSPSC₁₂ C, P₁₂C₇H₁₂, PëP»CH₂PSPsC†C, C- C, P^o
 PiPsPeP^oP·PSPëPeC-PI P·P^oPiP»C-PrPSP₁₂PSPSC₁₂ C† C†P^oPjPsPe (PrPëPI. C₇H₁₂PsP·PrC-P»
 B«P"°PSC- PrPsPeP»C-PSC-C†PSPëC... PrPsC†P»C-PrP¶P₁₂PSCH₂ P· P±P₁₂P·PiP₁₂PePëB»).

*P—PrP^oC, PSC-C†C, CH₂ PIPiP»PëPIP^oC, Pë PSP^o C¶PIPëPrPeC-C†C, CH₂ C₇H₁₂P₁₂P^oPeC†C-C— PiC₇H₁₂Pë
 PeP₁₂C₇H₁₂C†PIP^oPSPSC- P^oPIC, PsC, C₇H₁₂P^oPSC†PiPsC₇H₁₂, PsPj P^oP±PsB C-PSCëPëPjPë
 PjP₁₂C...P^oPSC-P·PjP^oPjPë.*

P₇P^oC₇H₁₂PëPiC₇H₁₂P^oPëPS PjP^oC" C†P»P^oP±PePëPM₂ P^oP±Ps PiPsPjC-C₇H₁₂PSPëPM₂ PIPiP»PëPI PSP^o
 P·PrP^oC, PSC-C†C, CH₂ PiP^oC†C-C" PSC, C-PI PeP₁₂C₇H₁₂C†PIP^oC, Pë P^oPIC, PsC, C₇H₁₂P^oPSC†PiPsC₇H₁₂, PsPj
 C, P^o PiC₇H₁₂P^oC†C₇H₁₂PIP^oC, Pë P· C-PSCëPëPjPë PjP₁₂C...P^oPSC-P·PjP^oPjPë. P₁₂P^oC†C-C" PSC, C-PI
 PiPsC, C₇H₁₂-P±PSPs P·P^oC†C, P₁₂C₇H₁₂PiC, Pë PIC-Pr C₇H₁₂PsP±PsC, Pë P· PSP₁₂P±P₁₂P·PiP₁₂C†PSPëPjPë
 PjP₁₂C...P^oPSC-P·PjP^oPjPë, PIPeP»C₇H₁₂C†PSPs P· P^oPIC, PsC, C₇H₁₂P^oPSC†PiPsC₇H₁₂, PsPj, PrPsPiPsPePë
 PIPsPSPë PSP₁₂ P±C†PrC†C, CH₂ C†C-P»PePsPj PIPiP₁₂PIPsP₁₂PSC-, C%Ps
 P»C-PeP^oC₇H₁₂C†CH₂PePëPM₂ P·P^oC†C-P± P P₁₂P^oPiC-P»P^o PSP₁₂ PjP^oC" PSP^o PSPëC...
 PSP₁₂PIP^oC, PëPIPSPsPiPs PIPiP»PëPIC†.

P₇YiPsC†C-P± P·P^oC†C, PsC†C¶PIP^oPSPSC₁₂ C, P^o PrPsP·Pë.

P"PsP·CfPIP°PSPSCl

P PμPePsPjPμPSPPrPsPIP°PSP° PīPsC±P°C,PePsPIP° PrPsP·P° PeP°CḡPēPīCḡP°P·PēPSCr
CfC,P°PSPsPIPēC,Ch 1,5 PjPi PsPrPēPS CḡP°P· PSP° PrPsP±Cf. P—PiPsPrPsPj PrPsP·Cf Cf
CḡP°P·C- PīPsC,CḡPμP±Pē PjPsP¶PSP° PīPsPIC-P»ChPSPs PīC-PrPIPēC‰CfPIP°C,Pē PīPs
1,5 PjPi PrPs PjP°PeCfPēPjP°P»ChPSPsC— PrPsP·Pē 6 PjPi/PrPsP±Cf. PkP°PNēPSPēP¶C±P°
PμC,,PμPeC,PēPIPSP° PrPsP·P° PjP°C" PīC-PrC,CḡPēPjCfPIP°C,PēCfCμ P·PiC-PrPSPs P·
PeP»C-PSC-C±PSPsCḡ PsC†C-PSPePsCḡ P»C-PeP°CḡCμ. P§PμCḡPμP· PrPsPIPīPēPNē
PīPμCḡC-PsPr PSP°PīC-PIPIPēPIPMPrPμPSPSCl PeP°CḡPēPīCḡP°P·PēPSCr C,P° PNēPsPiPs
P°PeC,PēPIPSPēC... PjPμC,P°P±PsP»C-C,C-PI P·PjC-PSPē PI PrPsP·CfPIP°PSPSC- PSPμ
P±CfPrCfC,Ch PīPsPIPSC-CfC,Cḡ PIC-PrPsP±CḡP°P¶P°C,PēCfCμ PI PīP»P°P·PjC-
PīCḡPsC,CμPiPsPj PeC-P»ChPePsC... C,PēP¶PSC-PI. P—P° CfC,P°PSPsPj PīP°C†C-C"PSC,P°
C,CḡPμP±P° CfPīPsCfC,PμCḡC-PiP°C,Pē CfC,PsCfPsPIPSs PīPsCμPIPē PīPsP±C-C±PSPēC...
CḡPμP°PeC†C-PNē C,P° PIC-PrPīPsPIC-PrC- PSP° P»C-PeCfPIP°PSPSCl PīCḡPsC,CμPiPsPj
PeC-P»ChPePsC... C,PēP¶PSC-PI PīC-CfP»Cμ PīPsC±P°C,PeCf PīCḡPēPNēPsPjCf
PeP°CḡPēPīCḡP°P·PēPSCr C,P° PīC-CfP»Cμ PePsP¶PSPsC— P·PjC-PSPē PrPsP·CfPIP°PSPSCl
(PrPēPI. CḡPsP·PrC-P» B«PαP°CḡPjP°PePsPeC-PSPμC,PēPeP°B».).

*PμPμCḡPμC...C-Pr P· C-PSCēPēC... P°PSC,PēPīCfPēC...PsC,PēPeC-PI PSP°
PeP°CḡPēPīCḡP°P·PēPS*

PμCḡPē PīPμCḡPμC...PsPrC- P· C-PSCēPēC... P°PSC,PēPīCfPēC...PsC,PēPeC-PI PSP°
PeP°CḡPēPīCḡP°P·PēPS PīPsC,CḡC-P±PSPs PICḡP°C...PsPICfPIP°C,Pē PīPsCfC,CfPīPsPIPM
PīPμCḡPμC...CḡPμCfPSPμ C,PēC,CḡCfPIP°PSPSCl P· PīPsCfC,CfPīPsPIPēPj
PīCḡPēPīPēPSPμPSPSClPj P·P°CfC,PsCfCfPIP°PSPSCl PīPsPīPμCḡPμPrPSCḡPsC—
C,PμCḡP°PīC-C— PīC-Pr C±P°Cf PīPsC±P°C,PeCf PīCḡPēPNēPsPjCf PeP°CḡPēPīCḡP°P·PēPSCr.

PμPμCḡPμC...C-Pr P· PeP°CḡPēPīCḡP°P·PēPSCē PSP° C-PSCēPēPNē P°PSC,PēPīCfPēC...PsC,PēPe

PμCḡPē PīPμCḡPμC...PsPrC- P· PeP°CḡPēPīCḡP°P·PēPSCr PSP° C-PSCēPēPNē
P°PSC,PēPīCfPēC...PsC,PēPe PīPsCfC,CfPīPsPIPsPiPs PīPμCḡPμC...CḡPμCfPSPsPiPs
C,PēC,CḡCfPIP°PSPSCl PSPμ PIC-PrP±CfPIP°C,PēPjPμC,ChCfCμ C,PsPjCf PīCḡPēPNēPsPj
PSPsPIPsPiPs P°PSC,PēPīCfPēC...PsC,PēPeP° PjPsP¶PSP° PīPsC±PēPSP°C,Pē P· PNēPsPiPs
PSP°PNēPSPēP¶C±PsC— PrPsP·Pē PīC-Pr C±P°Cf PīCḡPēPīPēPSPμPSPSCl
PIPēPePsCḡPēCfC,P°PSPSCl PeP°CḡPēPīCḡP°P·PēPSCr. PμPsC,CḡC-P±PSPs
PICḡP°C...PsPICfPIP°C,Pē, C‰Ps PePsPSC†PμPSC,CḡP°C†C-Cμ PeP°CḡPēPīCḡP°P·PēPSCr C,P°
PNēPsPiPs P°PeC,PēPIPSPēC... PjPμC,P°P±PsP»C-C,C-PI PI PīP»P°P·PjC- P·PjPμPSCēPēC,ChCfCμ
PSP° 50 % PīCḡPēP±P»PēP·PSPs PīCḡPsC,CμPiPsPj 1 C,PēP¶PSCl (PrPēPI. CḡPsP·PrC-P»
B«PαP°CḡPjP°PePsPeC-PSPμC,PēPeP°B»).

PμCḡPsPīCēCfPe PīCḡPēPNēPsPjCē C,P°P±P»PμC,PsPe

PμCḡPē PīCḡPsPīCfCfPeCf PīCḡPēPNēPsPjCf C,P°P±P»PμC,PePē PīP°C†C-C"PSC, PīPsPIPēPSPμPS
PSPμPiP°PNēPSPs PīCḡPēPNēPSClC,Pē PīCḡPsPīCfC‰PμPSCr C,P°P±P»PμC,PeCf. PhPrPSP°Pe
CμPeC‰Ps CfP¶Pμ PjP°PNēP¶Pμ PSP°CfC,P°PI C±P°Cf PīCḡPēPNēPsPjCf PSP°CfC,CfPīPSPsC—
C,P°P±P»PμC,PePē, PīCḡPsPīCfC‰PμPSCr C,P°P±P»PμC,PeCf PSPμ PIP°CḡC,Ps
PīCḡPēPNēPjP°C,Pē, P° CfP»C-Pr PīCḡPēPNēPSClC,Pē PSP°CfC,CfPīPSCr PrPsP·Cf P·PiC-PrPSPs
P·C- P·PIPēC±P°PNēPSPēPj PīCḡP°C,,C-PePsPj. PkPμ CḡPμPePsPjPμPSPPrCfC",ChCfCμ
PīCḡPēPNēPjP°C,Pē PīPsPrPIC-PNēPSCr PrPsP·Cf, C‰PsP± PePsPjPīPμPSCfCfPIP°C,Pē
PīCḡPsPīCfC‰PμPSCr PrPsP·Cf.

PhCfPsP±P»PēPIC- PīCḡCfPīPē PīP°C†C-C"PSC,C-PI

PμPsCḡCžC€PμPSPSCḡ C,,CžPSPeC†C-C— PSPëCḡPsPe

PμP°C†C-C"PS C,Pë P· PīPsCḡC†C€PμPSPSCḡPj C,,C†PSPeC†C-C— PSPëCḡPsPe P»PμPiPePsPiPs C†Pë C†PμCḡPμPrPSCḡPsPiPs C†C,C†PīPμPSCḡ C,CḡP¶PēPsC†C,C- (PeP»C-CḡPμPSC† PeCḡPμP°C,PëPSC-PSC† (CrCl) ≥ 30 PjP»/C...PI C,P° < 89 PjP»/C...PI) PSPμ PīPsC,CḡPμP±C†CḡC,Cḡ P¶PēPsPrPSPëC... PePsCḡPëPīC†PIP°PSCḡ PrPsP·Pë. P'PμP·PīPμC†PSC-C†C,Cḡ C,P° PμC,,PμPeC,PëPIPSC-C†C,Cḡ P·P°C†C,PSC†C†PIP°PSPSCḡ PeP°CḡPëPīCḡP°P·PëPSC† PrP»Cḡ PīP°C†C-C"PS C,C-PI C-P· C,CḡP¶PēPëPj PīPsCḡC†C€PμPSPSCḡPj C,,C†PSPeC†C-C— PSPëCḡPsPe (CrCl < 30 PjP»/C...PI) PSPμ PsC†C-PSCḡPIP°P»PëC†Cḡ. PμP°C†C-C"PS C,P°Pj C-P· C,CḡP¶PēPëPj PīPsCḡC†C€PμPSPSCḡPj C,,C†PSPeC†C-C— PSPëCḡPsPe PSPμ CḡPμPePsPjPμPSP†C†C,CḡC†Cḡ PīCḡPëPμPsPj PeP°CḡPëPīCḡP°P·PëPSC† (PrPëPI. CḡPsP·PrC-P» B«PαP°CḡPjP°PePsPeC-PSPμC,PëPeP°B»).

PμPsCḡCžC€PμPSPSCḡ C,,CžPSPeC†C-C— PīPμC†C-PSPePë

PμP°C†C-C"PS C,Pë P· PīPsCḡC†C€PμPSPSCḡ C,,C†PSPeC†C-C— PīPμC†C-PSPePë P»PμPiPePsPiPs P°P±Ps C†PμCḡPμPrPSCḡPsPiPs C†C,C†PīPμPSCḡ C,CḡP¶PēPsC†C,C- (5Bḡ"9 P·P° C€PeP°P»PsCḡ P§P°PNëP»PrP° Bḡ" PμBḡ™ Cḡ) PSPμ PīPsC,CḡPμP±C†CḡC,Cḡ P¶PēPsPrPSPëC... PePsCḡPëPīC†PIP°PSCḡ PrPsP·Pë. P'PμP·PīPμC†PSC-C†C,Cḡ C,P° PμC,,PμPeC,PëPIPSC-C†C,Cḡ P·P°C†C,PSC†C†PIP°PSPSCḡ PeP°CḡPëPīCḡP°P·PëPSC† B PrP»Cḡ PīP°C†C-C"PS C,C-PI C-P· C,CḡP¶PēPëPj PīPsCḡC†C€PμPSPSCḡPj C,,C†PSPeC†C-C— PīPμC†C-PSPePë (10Bḡ"15 P·P° C€PeP°P»PsCḡ P§P°PNëP»PrP° Bḡ" PμBḡ™ Cḡ) PSPμ PsC†C-PSCḡPIP°P»PëC†Cḡ. PμP°C†C-C"PS C,P°Pj C-P· C,CḡP¶PēPëPj PīPsCḡC†C€PμPSPSCḡPj C,,C†PSPeC†C-C— PīPμC†C-PSPePë PSPμ CḡPμPePsPjPμPSP†C†C,CḡC†Cḡ PīCḡPëPμPsPj PeP°CḡPëPīCḡP°P·PëPSC† (PrPëPI. CḡPsP·PrC-P» B«PαP°CḡPjP°PePsPeC-PSPμC,PëPeP°B»).

PμP°C†C-C"PS C,Pë P»C-C,PSCHPsPiPs PIC-PeCž

P"PsC†C,C†PīPSPëC... PrP°PSPëC... C†C,PSC†PsPIPSPs P»C-PeC†PIP°PSPSCḡ PeP°CḡPëPīCḡP°P·PëPSPsPj PīP°C†C-C"PS C,C-PI P»C-C,PSCHPsPiPs PIC-PeC† (≥ 65 CḡPsPeC-PI) PSPμPrPsC†C,P°C,PSCHPsB PrP»Cḡ PIPëCḡPIP»PμPSPSCḡ PIC-PrPjC-PSPSPC†C,PμPNë C—C...PSCHPsC— PIC-PrPīPsPIC-PrC- PSP° P»C-PeC†PIP°PSPSCḡ PIC-Pr PIC-PrPīPsPIC-PrC- PīP°C†C-C"PS C,C-PI PjPsP»PsPrC€PsPiPs PIC-PeC† (PrPëPI. CḡPsP·PrC-P» B«PαP°CḡPjP°PePsPeC-PSPμC,PëPeP°B»). PμC-PrP±C-Cḡ PrPsP·Pë PrP»Cḡ PīP°C†C-C"PS C,C-PI P»C-C,PSCHPsPiPs PIC-PeC† PjP°C" P±C†C,Pë P±C-P»CHC€ PsP±PμCḡPμP¶PSPëPj.

PŸPīPsC†C-P± P·P°C†C,PSC†C†PIP°PSPSCḡ

P»C-PeP°CḡC†CḡPePëPNë P·P°C†C-P± P PμP°PiC-P»P° P·P°C†C,PSC†PsPIC†PIP°C,Pë PīPμCḡPsCḡP°P»CHPSPs 1 CḡP°P· PSP° PrPsP±C† PI PsPrPëPS C- C,PSPNë C†P°PjPëPNë C†P°C† PSPμP·P°P»PμP¶PSPs PIC-Pr PIP¶PëPIP°PSPSCḡ C—P¶C-.

PμCḡPë P·P°C†C,PSC†C†PIP°PSPSC- PeP°CḡPëPīCḡP°P·PëPSC† C†P»C-Pr C†PSPëPeP°C,Pë PIP¶PëPIP°PSPSCḡ P°P»PePsPiPsP»Cḡ (PrPëPI. CḡPsP·PrC-P» B«P'P·P°C" PjPsPrC-Cḡ P· C-PSC€PëPjPë P»C-PeP°CḡC†CḡPePëPjPë P·P°C†PsP±P°PjPë C,P° C-PSC€C- PIPëPrPë PIP·P°C" PjPsPrC-PNëB»).

P"C-C,Pë.

P'PμP·PīPμPeC† C,P° PμC,,PμPeC,PëPIPSC-C†C,Cḡ P·P°C†C,PSC†C†PIP°PSPSCḡ

PeP°CḡPēPīCḡP°P·PēPSCř PrP»Cḡ PrC-C, PμPNē (PIC-PePsPj PrPs 18 CḡPsPeC-PI) PSPμ
PICřC, P°PSPsPIP»PμPSPs. PkPμPjP°C” PrP°PSPēC....

PμPμCḡPμPrPsP·CzPIP°PSPSCḡ.

PŸPēPjPīC, PsPjPē

PμPsPIC-PrPsPjP»CḡP»PsCřCḡ PīCḡPs PIPēPīP°PrPePsPIPμ PiPsCřC, CḡPμ
PīPμCḡPμPrPsP·CřPIP°PSPSCḡ (48 PjPi/PrPsP±Cř) PI PsPrPSPsPiPs PīP°C†C-C”PSC, P°. PJ
C†CHbPsPiPs PīP°C†C-C”PSC, P° P·Bḡ™ CḡPIPēP»PēCřCḡ PsCḡC, PsCřC, P°P· C, P°
CřPμPrP°C, PēPIPSPēPNē PμC,, PμPeC,. PŸC, P°PS PīP°C†C-C”PSC, P° PīPsPIPSC-CřC, Cḡ
PIC-PrPSPsPIPēPICřCḡ PI C, PsPNē PḡPμ PrPμPSCḡ.

P>C-PeCřPIP°PSPSCḡ PīCḡPē PīPμCḡPμPrPsP·CřPIP°PSPSC-

P>C-PeCřPIP°PSPSCḡ PīCḡPē PīPμCḡPμPrPsP·CřPIP°PSPSC- PjP°C” P±CřC, Pē
PīC-PrC, CḡPēPjCřPIP°P»CHbPSPμ, PIPeP»CḡC†PSPs P· PīC-PrC, CḡPēPjPePsCḡ PIC-P»CHbPSPsC—
PīCḡPsC...C-PrPSPsCřC, C- PrPēC...P°P»CHbPSPēC... CēP»CḡC...C-PI, PSP°CřPēC†PμPSPSCḡPj
PePēCřPSPμPj C, P° PIPμPSC, PēP»CḡC†C-C”Cḡ, P° C, P°PePsPḡ CřPēPjPīC, PsPjP°C, PēC†PSPμ.
PŸPiPsCřC, PμCḡPμPḡPμPSPSCḡ P·P° CřPμCḡC†PμPIPs-CřCřPrPēPSPSPsCḡ CřPēCřC, PμPjPsCḡ
PjP°C” CḡPsP·PiPsC†PēPSP°C, PēCřCḡ PSPμPīP°PNēPSPs C, P° PIPeP»CḡC†P°C, Pē
PμP»PμPeC, CḡPsPeP°CḡPrC-PsPiCḡP°C,, C-C†PSPēPNē PjPsPSC-C, PsCḡPēPSPi PrP»Cḡ
PIPēCḡPIP»PμPSPSCḡ PjPsPḡP»PēPIPēC... P°CḡPēC, PjC-PNē. PJ CḡP°P·C-
PIPēPSPēPePSPμPSPSCḡ C, CḡPḡPePēC... PμPeCřC, CḡP°PīC-CḡP°PjC-PrPSPēC...
CřPēPjPīC, PsPjC-PI CřP»C-Pr P·P°CřC, PsCřCřPIP°C, Pē P°PSC, PēC...PsP»C-PSPμCḡPiC-C†PSC-
PīCḡPμPīP°CḡP°C, Pē. PkCřPeC-P»CHbPePē PeP°CḡPēPīCḡP°P·PēPS CřPēP»CHbPSPs
P·PiBḡ™ CḡP·CřC”C, CHbCřCḡ P· P±C-P»PeP°PjPē PīP»P°P·PjPē, PiPμPjPsPrC-P°P»C-P· PSPμ
P±CřPrPμ B PμC,, PμPeC, PēPIPSPēPj PrP»Cḡ P>C-PeCřPIP°PSPSCḡ
PīPμCḡPμPrPsP·CřPIP°PSPSCḡ. PμPēP»CHbPSPēPNē PjPμPrPēC†PSPēPNē PSP°PiP»CḡPḡ C, P°
PjPsPSC-C, PsCḡPēPSPi PjP°CḡC, CHb C, CḡPēPIP°C, Pē PrPsC, Pē, PrPsPePē CřC, P°PS
PīP°C†C-C”PSC, P° PSPμ PIC-PrPSPsPIPēC, CHbCřCḡ.

PḡP°CḡPēPīCḡP°P·PēPS PSPμ PjP°C” CřPīPμC†PēC,, C-C†PSPsPiPs P°PSC, PēPrPsC, Cř.

PμPsP±C-C†PSC- CḡPμP°PeC†C-C—.

PμPsP±C-C†PSPēPjPē CḡPμP°PeC†C-CḡPjPē, PīCḡPs CḡPeC- PSP°PNēC†P°CřC, C-CēPμ
PīPsPIC-PrPsPjP»CḡP»PsCřCḡ PīCḡPē P·P°CřC, PsCřCřPIP°PSPSC- PeP°CḡPēPīCḡP°P·PēPSCř PI
PrC-P°PīP°P·PsPSC- PrPsP· (1,5Bḡ“6 PjPi), P±CřP»Pē P°PeP°C, PēP·C-Cḡ (19 %) C, P°
PīP°CḡPeC-PSCřPsPSC-P·Pj (17,5 %). P·C-P»CHbCēC-CřC, CHb CḡPμP°PeC†C-PNē P±CřP»Pē
P»PμPiPePsPiPs C†Pē CřPμCḡPμPrPSCḡPsPiPs CřC, CřPīPμPSCḡ C, CḡPḡPePsCřC, C-.

PŸP°P±P»PēC†PSPēPNē PīPμCḡPμP»C-Pe PīPsP±C-C†PSPēC... CḡPμP°PeC†C-PNē

PμPsP±C-C†PSC- P»C-PeP°CḡCřCHbPeC- CḡPμP°PeC†C-C— (PμP>P) PSP° PsCřPSPsPIC-
PsP±Bḡ™ C”PrPSP°PSPēC... PrP°PSPēC... P· PrPsCřP»C-PrPḡPμPSPSCḡ P>C-PeCřPIP°PSPSCḡ
CēPēP·PsC,, CḡPμPSC-C— PeP°CḡPēPīCḡP°P·PēPSPsPj PīCḡPμPrCřC, P°PIP»PμPSPs P·P°
CřPēCřC, PμPjP°PjPē PsCḡPiP°PSC-PI C- P· PIPēPePsCḡPēCřC, P°PSPSCḡPj
PIC-PrPiPsPIC-PrPSPsPiPs C, PμCḡPjC-PSP° Cř C, P°P±P»PēC†C- 3.

$P_{\mu}PsP \pm C - C \ddagger PSC - C \ddot{H}P_{\mu}P^{\circ}PeC \uparrow C - C - C \ddot{H}PsP \cdot \ddot{P}iPsPrC - P \gg P_{\mu}PSC - P \cdot P^{\circ}C \ddagger P^{\circ}C \uparrow C, PsC, PsC \ddot{H}$
 $PIP \ddot{e}PSP \ddot{e}PePSP_{\mu}PSPSC \uparrow C, P^{\circ}PeP \ddot{e}P \ddot{e} C \ddagger P \ddot{e}PSPsPj: PrC \acute{r}P \uparrow P_{\mu}C \ddagger P^{\circ}C \uparrow C, C - (\geq 1/10); C \ddagger P^{\circ}C \uparrow C, C -$
 $(PIC - Pr \geq 1/100 \text{ } PrPs < 1/10); PSP_{\mu}C \ddagger P^{\circ}C \uparrow C, C - (PIC - Pr \geq 1/1000 \text{ } PrPs < 1/100);$
 $\ddot{P}iPsPsPrP \ddot{e}PSPsPeC - (PIC - Pr \geq 1/10000 \text{ } PrPs < 1/1000); C \ddot{H}C - PrPeC - C \uparrow PSC - (< 1/1000);$
 $C \ddagger P^{\circ}C \uparrow C, PsC, P^{\circ}PSP_{\mu}PIC - PrPsPjP^{\circ} (PSP_{\mu}PjPsP \uparrow P \gg P \ddot{e}PIPs \text{ } PIP \ddot{e}P \cdot PSP^{\circ}C \ddagger P \ddot{e}C, P \ddot{e}PSP^{\circ}$
 $PsC \uparrow PSPsPIC - PSP^{\circ}C \uparrow PIPSP \ddot{e}C \dots PrP^{\circ}PSP \ddot{e}C \dots). PjPePsP \uparrow PSC - P \ddot{N}_{\ddot{e}}PiC \ddot{H}C \acute{r}P \ddot{e}iC - P \cdot P^{\circ}$
 $C \ddagger P^{\circ}C \uparrow C, PsC, PsC \ddot{H}PIP \ddot{e}PSP \ddot{e}PePSP_{\mu}PSPSC \uparrow P \ddot{e}PsP \pm C - C \ddagger PSC - C \ddot{H}P_{\mu}P^{\circ}PeC \uparrow C - C -$
 $P \ddot{e}iC \ddot{H}P_{\mu}PrC \uparrow C, P^{\circ}PIP \gg P_{\mu}PSC - PiP \ddot{e}PsC \ddot{H}C \uparrow PrPeC \acute{r}P \cdot PSP \ddot{e}P \uparrow P_{\mu}PSPSC \uparrow C - C \dots PSCH \ddot{H}PsC -$
 $C \uparrow P_{\mu}C \ddot{H}P \ddot{N}_{\ddot{e}}PsP \cdot PSPsC \uparrow C, C -.$

$P \ddot{y}P^{\circ}P \pm P \gg P \ddot{e}C \uparrow C \uparrow 3. P_{\mu}PsP \pm C - C \ddagger PSC - C \ddot{H}P_{\mu}P^{\circ}PeC \uparrow C - C - , C \% PsPIP \ddot{e}PSP \ddot{e}PeP^{\circ}CHC, CHC \acute{r}$
 $P \ddot{e}P^{\circ}C \uparrow C - C''PSC, C - PiC - P \cdot C \ddot{e}P \ddot{e}P \cdot PsC, C \ddot{H}P_{\mu}PSC - C''CH$

$P \ddot{V}PeC \uparrow C, P_{\mu}B \uparrow P \ddot{e}PsC \ddot{H}P \ddot{e}P^{\circ}PSC - PiP \cdot P \cdot P^{\circ}MedDRA$	$P^{\circ}C \uparrow P \uparrow P_{\mu}C \ddagger P^{\circ}C \uparrow C, C - (\geq 1/10)$	$PSP^{\circ}C \uparrow C, C - (PIC - Pr \geq 1/100 \text{ } PrPs < 1/10)$	$PkP_{\mu}C \ddagger P^{\circ}C \uparrow C, C - (PIC - Pr \geq 1/1000 \text{ } PrPs < 1/100)$	$P_{\mu}PsPsPrP \ddot{e}PSPsPeC - (PIC - Pr \geq 1/10000 \text{ } PrPsB \text{ } B \text{ } B \text{ } B \text{ } B \text{ } B \text{ } B < 1/1000)$	$PSP^{\circ}C \uparrow C, PsC, P^{\circ}PSP_{\mu}BPIC - PrPsPjP^{\circ}$
$P_{\mu}C \ddot{H}PsPiC, P^{\circ}P \gg C - PjC, P^{\circ}C, P \ddot{e}C \ddagger PSP^{\circ}C \uparrow P \ddot{e}C \uparrow C, P_{\mu}PjP^{\circ}$			$P_{\mu}PSP_{\mu}PjC - C \uparrow P \gg PsP \cdot P \ddot{e}PSPsC, C - P \gg C - C \uparrow$	$PkP_{\mu}P \ddot{N}_{\ddot{e}}C, C \ddot{H}PsPiP_{\mu}PSC - C \uparrow$	
$P \uparrow PjC \uparrow PSPSP^{\circ}C \uparrow P \ddot{e}C \uparrow C, P_{\mu}PjP^{\circ}$				$P_{\mu}C - PrPiP \ddot{e}C \% P_{\mu}PSP^{\circ}C \ddagger C \uparrow C, P \gg P \ddot{e}PIC - C \uparrow C, CH$	
$P \cdot PSPPrPsPeBC \ddot{H}P \ddot{e}PSPSP^{\circ}C \uparrow P \ddot{e}C \uparrow B - C, P_{\mu}PjP^{\circ}$			$P - PSP \ddot{e}P \uparrow P_{\mu}PSPSC \uparrow C \ddot{H}C - BPiPSC \uparrow C, P \ddot{e}C \ddot{H}P_{\mu}PsC, B - C \ddot{H}PsPiPSPsPiPsPiPsC \ddot{H}BPjPsPSC \acute{r}PiP \ddot{e}C \ddot{H}PsPIC -$	$P^{\circ}C - PiPsC, P \ddot{e}C \ddot{H}P_{\mu}PsP -$	
$P_{\mu}P \pm PjC - PS \ddot{H}P_{\mu}C \ddagger PsPiPePsC, P^{\circ}C \dots P^{\circ}C \ddot{H}C \uparrow C \uparrow P \ddot{e}PSPSC \uparrow$		$P^{\circ}P \ddot{e}C \uparrow P \gg C - PiC - PrP_{\mu}PjC - C \uparrow P - P \gg C - P \gg CHC \ddot{e}P_{\mu}PSPSC \uparrow PjP^{\circ}C \uparrow P \ddot{e}C, C, C - P \gg P^{\circ}P - PSP \ddot{e}P \uparrow P_{\mu}PSPSC \uparrow P^{\circ}PiP_{\mu}C, P \ddot{e}C, C \acute{r}P_{\mu}C - PrPiP \ddot{e}C \% P_{\mu}PSPSC \uparrow P^{\circ}PiP_{\mu}C, P \ddot{e}C, C \acute{r}$	$P_{\mu}PSPsC \ddot{H}PjP^{\circ}P \gg CHPSP \ddot{e}P \ddot{N}_{\ddot{e}}C \ddot{H}C - PiP_{\mu}PSC \ddot{H}PSP^{\circ}C, C \ddot{H}C - CHPiP \ddot{e}C \ddot{H}PsPIC - P \uparrow C \uparrow P \ddot{e}C \ddot{H}PsPiP \ddot{e}P \ddot{N}_{\ddot{e}}PrC - BP^{\circ}P \pm P_{\mu}C, P_{\mu}C - PrPiP \ddot{e}C \% P_{\mu}PSP \ddot{e}P \ddot{N}_{\ddot{e}}C \ddot{H}C - PiP_{\mu}PSC \ddot{H}PiP \gg CHPePsBP - P \ddot{e}PiP \ddot{e}C \ddot{H}PsPIC -$		
$P_{\mu}C \uparrow P \ddot{e}C \dots C - PeP^{\circ}$		$P \text{ } PsP \cdot P \gg P^{\circ}PrP \ddot{e}C \uparrow PSC \acute{r}P \ddot{y}C \ddot{H}P \ddot{e}PiPsP \uparrow PSC - C \uparrow C, CH$	$P \ddot{y}C \uparrow C - C \uparrow P \ddot{e}PrP^{\circ}P \gg CHPSP^{\circ}PiPsPiP_{\mu}PrC - PSP \ddot{e}P^{\circ}P^{\circ}P_{\mu}P \gg C - C \ddot{H}C - P \ddot{N}_{\ddot{e}}P^{\circ}P_{\mu}PiC \ddot{H}P_{\mu}C \uparrow C - C \uparrow P - PSP \ddot{e}P \uparrow P_{\mu}PSP_{\mu}P \gg C - P \ddot{e}PrPsP_{\mu}C - PrPiP \ddot{e}C \% P_{\mu}PSP_{\mu}P \gg C - P \ddot{e}C - PrPsP^{\circ}C \ddot{H}P_{\mu}PeC, P \ddot{e}P \gg CHPSP^{\circ}PrP \ddot{e}C \uparrow C, C \uparrow PSC - C \uparrow C$		
$PkP_{\mu}C \ddot{H}PiPsBPiP^{\circ}C \uparrow P \ddot{e}C \uparrow C, P_{\mu}PjP^{\circ}$	$P_{\mu}PeC, P \ddot{e}P - C - C \uparrow P^{\circ}P_{\mu}P^{\circ}C \ddot{H}PeC - PSC \uparrow PsBPSC - P \cdot PjP^{\circ}$	$P \ddot{y}P_{\mu}PrP^{\circ}C, P \ddot{e}PiPSP \ddot{e}P \ddot{N}_{\ddot{e}}P_{\mu}C, P_{\mu}PeC, P - P^{\circ}PiP^{\circ}PiPsC \ddot{H}PsBC \ddagger P_{\mu}PSPSC \uparrow P^{\circ}P \ddot{e}C \uparrow C, P \ddot{e}PSC - C \uparrow P^{\circ}PiPSC \ddot{e}C - P_{\mu}PeC \uparrow C, C \ddot{H}P^{\circ}B \ddot{e}iC - C \ddot{H}P^{\circ}PjC - PrPSC - C, P^{\circ}C \ddot{H}C \uparrow C \dots PsPIC - C \ddot{H}PsP \cdot P \gg P^{\circ}BPPrP \ddot{e}^{\circ}$	$P_{\mu}C - P \text{ } PSC \uparrow P \ddot{e}C \uparrow P \ddot{e}C - BPSP_{\mu}P - C - C \uparrow P^{\circ}P \ddot{e}C \uparrow C - PSP_{\mu}P - C - C \uparrow P^{\circ}P \ddot{e}P \cdot P_{\mu}C \uparrow C, P_{\mu}P - C - C \uparrow P - P^{\circ}PiP^{\circ}P \gg CHPjPsPiP^{\circ}BPSC - C \uparrow C, CH$	$P \ddot{y}C \uparrow P \ddot{e}PsBPjP \ddot{e}P_{\mu}PsPSPiC \uparrow P^{\circ}CHC \uparrow C - C - PjPjPSP_{\mu}P - C - C \uparrow PjC, P^{\circ}P^{\circ}C - C \uparrow$	$P - P \gg PsC \uparrow P \ddot{e}C - C \uparrow PSP \ddot{e}P \ddot{N}_{\ddot{e}}PSP_{\mu}P \ddot{N}_{\ddot{e}}C \ddot{H}PsP \gg P_{\mu}PiC, P \ddot{e}C \ddagger PSP \ddot{e}P \ddot{N}_{\ddot{e}}C \uparrow P \ddot{e}PSP \uparrow C \ddot{H}PsPj$
$P_{\mu}C \ddot{H}PiP^{\circ}PSP \ddot{e}P \cdot PsC \ddot{H}C \acute{r}$		$P \text{ } PsP \cdot PjP \ddot{e}C, P \ddot{e}P \ddot{N}_{\ddot{e}}P - C - C \ddot{H}$	$P_{\mu}C - PrPiP \ddot{e}C \% P_{\mu}PSP \ddot{e}P \ddot{N}_{\ddot{e}}PiPSC \uparrow C, C \ddot{H}C - C \ddot{e}PSC \ddot{H}PsBC \ddagger PSP \ddot{e}P \ddot{N}_{\ddot{e}}C, P \ddot{e}C \uparrow P \ddot{e}P_{\mu}PsC \ddot{H}C \ddot{e}P_{\mu}PSPSC \uparrow P^{\circ}PePsPiPsPrP^{\circ}C \uparrow C - C - P - PSP \ddot{e}P \uparrow P_{\mu}PSPSC \uparrow PiPsBC \uparrow C, C \ddot{H}PsC, P \ddot{e}P - PsC \ddot{H}C \acute{r}P_{\mu}PsPrC \ddot{H}P^{\circ}P - PSP_{\mu}PSPSC \uparrow PsPeP^{\circ}$	$P_{\mu}P^{\circ}C, P^{\circ}C \ddot{H}P^{\circ}PeC, P^{\circ}P_{\mu}PsC, PsC, PsP \pm C - C \uparrow$	
$P_{\mu}C \ddot{H}PiP^{\circ}PSP \ddot{e}C \uparrow P \gg C \uparrow C \dots C \acute{r}C, P^{\circ}C \ddot{H}C - PiBPSPsPiP^{\circ}PiP \ddot{e}P \ddot{y}P_{\mu}C \ddot{H}C \uparrow P_{\mu}$		$P \ddot{y}P^{\circ}C \dots C - P^{\circ}C \ddot{H}P \ddot{e}C, PjC - C \uparrow$	$P_{\mu}PsC \ddot{H}C \ddot{e}P_{\mu}PSPSC \uparrow C \uparrow P_{\mu}C \ddot{H}C \uparrow PjPiPSC - PiC \ddot{H}PsBPIC - PrPsPsC \uparrow C, C - P^{\circ}C \ddot{H}P^{\circ}PrC - P^{\circ}C \ddot{H}P \ddot{e}C, PjC - C \uparrow P_{\mu}PsPrPsPiP^{\circ}P_{\mu}PSPSC \uparrow C - PSC, P_{\mu}C \ddot{H}PiP^{\circ}P \gg C \acute{r}QT \text{ } PSP^{\circ}P_{\mu}P \gg P_{\mu}PeC, C \ddot{H}PsPeP^{\circ}BC \ddot{H}PrC - PsPiC \ddot{H}P^{\circ}PjC - P_{\mu}P^{\circ}C, PsP \gg PsPiC - C \ddagger PSP \ddot{e}P \ddot{N}_{\ddot{e}}P - C \uparrow P \ddot{e}P_{\mu}C \uparrow CH P \ddot{y} \text{ } PSP^{\circ}P_{\mu}P \gg P_{\mu}PeC, C \ddot{H}PsPeP^{\circ}C \ddot{H}BPPrC - PsPiC \ddot{H}P^{\circ}PjC -$		
$P \ddot{V}C \uparrow PrP \ddot{e}PSP \ddot{e}$		$P_{\mu}C \ddot{H}C, P_{\mu}C \ddot{H}C - P^{\circ}P \gg CHPSP^{\circ}PiC - PiP_{\mu}C \ddot{H}C, P_{\mu}PSP - C - C \uparrow$	$P_{\mu}C \ddot{H}C, P_{\mu}C \ddot{H}C - P^{\circ}P \gg CHPSP^{\circ}PiC - PiPsC, P_{\mu}PSP - C - C \uparrow$		
$P_{\mu}C \ddot{H}PiP^{\circ}PSP \ddot{e}PrP \ddot{e}C \dots P^{\circ}PSPSC \uparrow, PiC \ddot{H}C \uparrow PrPsPsC - PeP \gg C - BC, PeP \ddot{e}C, P^{\circ}C \uparrow P_{\mu}C \ddot{H}P_{\mu}BPPrPsC \uparrow C, C - PSPSC \uparrow$			$P^{\circ}P \ddot{e}P \ddot{e}P^{\circ}PiP \ddot{e}P^{\circ}$		
$PeP \gg C \uparrow PSPBP \ddot{e}PiPs - PeP \ddot{e}C \ddot{e}PePsPiP \ddot{e}P \ddot{N}_{\ddot{e}}C, C \ddot{H}P^{\circ}PeC,$		$P^{\circ}P \gg C \ddot{H}PiP^{\circ}PSPSC \uparrow PkC \uparrow PrPsC, P^{\circ}P - P^{\circ}PiPsC \ddot{H}$	$P^{\circ}P^{\circ}C \uparrow C, C \ddot{H}PsP_{\mu}P - PsBC, P^{\circ}PiP_{\mu}P^{\circ}P \gg CHPSP^{\circ}C \ddot{H}P_{\mu}C, P \gg CHPeC \uparrow PSP^{\circ}C \dots PiPsC \ddot{H}PsP \ddot{e}P^{\circ}$	$P^{\circ}P \ddot{e}C \uparrow C, P^{\circ}PiC - C \uparrow$	
$P_{\mu}PiPjC \uparrow C - PSP \ddot{e}P^{\circ}C, P^{\circ}P^{\circ}P^{\circ}PiPsC \ddagger PsPiP \ddot{e}B - PiC - PrPSC - C \ddot{e}P - C \uparrow B - C, P \ddot{e}P \ddot{e}C - C \ddot{H}P^{\circ}C - P^{\circ}PiC - PrC \ddot{e}P - C \ddot{H}B - PSC - C, P \ddot{e}P^{\circ}PSP \ddot{e}PSP \ddot{e}$		$P_{\mu}C - PrPiP \ddot{e}C \% P_{\mu}PSP \ddot{e}P \ddot{N}_{\ddot{e}}C \ddot{H}C - PiP_{\mu}PSC \ddot{H}PiP_{\mu}C \uparrow C - PSBP \ddot{e}PiP \ddot{e}C \dots C, P_{\mu}C \ddot{H}PjP_{\mu}BPSC, C - Pi$	$P_{\mu}C - PrPiP \ddot{e}C \% P_{\mu}PSP \ddot{e}P \ddot{N}_{\ddot{e}}C \ddot{H}C - PiP_{\mu}PSC \ddot{H}P \ddot{e}C \uparrow P_{\mu}C \uparrow CH P \ddot{y} \text{ } PSP^{\circ}P_{\mu}P \gg P_{\mu}PeC, C \ddot{H}PsPeP^{\circ}C \ddot{H}BPPrC - PsPiC \ddot{H}P^{\circ}PjC -$		$P \ddot{y}PsPeC \uparrow P \ddot{e}C \ddagger PSP \ddot{e}P \ddot{N}_{\ddot{e}}PiP_{\mu}PiP^{\circ}C, P \ddot{e}C,$
$P_{\mu}PiPsC \ddot{H}PSPs - C \ddot{H}C \uparrow C \dots PsPiP \ddot{e}P \ddot{N}_{\ddot{e}}P^{\circ}PiP^{\circ}C \ddot{H}P^{\circ}C, C, P^{\circ}C \uparrow PiPsP \gg C \uparrow C \ddagger PSP^{\circ}C, PeP^{\circ}PSP \ddot{e}PSP^{\circ}$		$P_{\mu}C - PrPiP \ddot{e}C \% P_{\mu}PSP \ddot{e}P \ddot{N}_{\ddot{e}}C \ddot{H}C - PiP_{\mu}PSC \ddot{H}PeC \ddot{H}P_{\mu}P^{\circ}BC, P \ddot{e}PSC - PSC, PsC \uparrow C, PsBP \ddot{e}C - PSP^{\circ}P \cdot P \ddot{e}PiP \ddot{e}C \ddot{H}PsPIC -$	$P \ddot{y}PiP_{\mu}C \ddot{H}P \ddot{e}P \ddot{N}_{\ddot{e}}P^{\circ}P \ddot{e}C \uparrow P \ddot{e}Pi$	$P \text{ } P^{\circ}P \pm PrPsPjC - PsP \gg C - P -$	
$PkP \ddot{e}C \ddot{H}PeP \ddot{e}C, P^{\circ}C \uparrow P_{\mu}C \ddagger PsPiP \ddot{e}BPIC - PrPSC - C \ddot{e}P \gg C \uparrow B - C \dots P \ddot{e}$			$P^{\circ}P \ddot{e}P - C \uparrow C \ddot{H}C - C \uparrow P_{\mu}PsBP \gg P^{\circ}PeC - C \uparrow C \ddot{H}C - C \uparrow$		
$P^{\circ}P^{\circ}PiC - C - PSC - C \uparrow C, CH, PiC - C \uparrow P \gg C \uparrow PjPsP \gg PsPiPsBPiP \ddot{e}P \ddot{N}_{\ddot{e}}PiP_{\mu}C \ddot{H}C - PsPrC, P^{\circ}PiP_{\mu}C \ddot{H}P \ddot{e}PSP^{\circ}C, P^{\circ}P \gg CHBPSC - C \uparrow C, P^{\circ}PSP \ddot{e}$					$PkP_{\mu}PsPSP^{\circ}C, P^{\circ}P \gg CHBPSP \ddot{e}P \ddot{N}_{\ddot{e}}P^{\circ}P \pm C \uparrow C, P \ddot{e}PSP_{\mu}BPSC, PSP \ddot{e}P \ddot{N}_{\ddot{e}}C \uparrow P \ddot{e}PSPPrBC \ddot{H}PsPj (PrP \ddot{e}Pi, C \ddot{H}PsP - PrC - P \gg B \gg P - P^{\circ}B - C \uparrow C, PsC \uparrow C \uparrow PjP^{\circ}PSPSC \uparrow C \uparrow PiP_{\mu}C \ddot{H}C - PsPrPiP_{\mu}PiC - C, BPSPsC \uparrow C, C - P^{\circ}P \pm PsPiPsBPPrC \uparrow PjP^{\circ}PSPSC \uparrow PiC \ddot{H}C \uparrow BPrPrC \ddot{H}B \gg)$
$PjC \uparrow P \ddot{e}P - P^{\circ}P \ddot{y}PSP_{\mu}PSPSC \uparrow P \cdot P^{\circ}PiP^{\circ}P \gg CHPSPsPiPsC \dots P^{\circ}C \ddot{H}P^{\circ}PeC, P_{\mu}C \ddot{H}C \acute{r}$		$P^{\circ}C, PsPjP^{\circ}$	$P \ddot{y}PiC \ddot{H}P^{\circ}PiP^{\circ}$		

¹ $P \text{ } PsP \cdot P \gg P^{\circ}PrP \ddot{e}C \uparrow PSC \acute{r}P \text{ } P \pm P_{\mu}P \cdot C \uparrow P \ddot{e}PSPSC \uparrow, P^{\circ}PSPsPjP^{\circ}P \gg CHPSC - C \uparrow PSP \ddot{e}, PePsC \ddot{e}PjP^{\circ}C \ddot{H}P \ddot{e},$

P PsP·PIPëC,PsPe PeP°C,P°CĤP°PeC,Pë CÍPíPsCÍC,PμCĤC-PiP°PICÍCĬ PíC-Pr C±P°CÍ
 PrPsPeP»C-PSC-C±PSPëC... PrPsCÍP»C-PrP¶PμPSCĤ PeP°CĤPëPíCĤP°P·PëPSCr (PrPëPI.
 CĤPsP·PrC-P» B«P”P°PSC- PrPsPeP»C-PSC-C±PSPëC... PrPsCÍP»C-PrP¶PμPSCĤ P·
 P±PμP·PíPμPePëB»). PÿPsPjCÍ P·P° C„PsCĤPjCÍPIP°PSPSCĬPj PeP°C,P°CĤP°PeC,Pë
 CÍPíPsCÍC,PμCĤC-PiP°P»Pë P·P° PrPsPíPsPjPsPiPsCĤ PsPiP»CĬPrC-PI C-P·
 C%oC-P»PëPSPSPSCĤ P»P°PjPíPsCĤ, P° PíP°C†C-C”PSC,Pë, CĬPeC- PIP¶Pμ PjP°P»Pë
 PeP°C,P°CĤP°PeC,CÍ, P±CÍP»Pë PIPëPeP»CĤC±PμPSC- P· PrPsCÍP»C-PrP¶PμPSPSCĬ. PμC-Pr
 C±P°CÍ PeP»C-PSC-C±PSPsC— PíCĤPsPiCĤP°PjPë PrPsCÍP»C-PrP¶PμPSPSCĬ PrC-C—
 PeP°CĤPëPíCĤP°P·PëPSCr CÍ PíP°C†C-C”PSC,C-PI P· C€PëP·PsC„CĤPμPSC-C”CĤ
 PíPsPIC-PrPsPjP»CĬP»PsCÍCĬ PíCĤPs PeC-P»CĤPeP° PIPëPíP°PrPeC-PI PeP°C,P°CĤP°PeC,Pë,
 C%oPs C...P°CĤP°PeC,PμCĤPëP·CÍPIP°P»P°CÍCĬ PSPμP·PSP°C±PSPëPj PíPsPjCÍC,PSC-PSPSCĬPj
 PeCĤPëC€C,P°P»PëPeP° P±PμP· PíPsCĤCÍC€PμPSPSCĬ P·PsCĤCÍ (13/3192; 0,4 %). P”PμCĬPeC-
 P· C†PëC... PíP°C†C-C”PSC,C-PI PjP°P»Pë C±PëPSPSPëPePë CĤPëP·PëPeCÍ.
 PkP°PNëC±P°CÍC,C-C€Pμ PíPsPIC-PrPsPjP»CĬP»PsCÍCĬ PíCĤPs C,P°PePëPNë PíPsP±C-C±PSPëPNë
 PμC„PμPeC, P· P±PsPeCÍ PsCĤPiP°PSC-PI P·PsCĤCÍ, CĬPe PSPμC±C-C,PeC-CÍC,CĤ P·PsCĤCÍ

(PīP»P°C†PμP±Ps: 1/683, 0,1 %; PeP°CḡPēPīCḡP°P·PēPS: 22/2048, 1,1 %).

P•PeCfC,CḡP°PīC-CḡP°PjC-PrPSC- CfPēPjPīC,PsPjPē (P•PμPŸ)

Pj PePsCḡPsC,PePsC,CḡPēPIP°P»PēC... PrPsCfP»C-PrP¶PμPSPSC¶C... C†P°CfC,PsC,P° PIPēPSPēPePSPμPSPSC¶ P•PμPŸ CfC,P°PSPsPIPēP»P° 27 %, 11,5 %, 30,7 % C,P° 15,1 % PīP°C†C-C°PSC,C-PI, C¶PeC- P»C-PeCfPIP°P»PēCfC¶ PIC-PrPīPsPIC-PrPSPs PeP°CḡPēPīCḡP°P·PēPSPsPj, PīP»P°C†PμP±Ps, CḡPēCfPīPμCḡPēPrPsPSPsPj C,P° P°CḡPēPīC-PīCḡP°P·PsP»PsPj. PḡPeP°C,PēP·C-C¶ CfPīPsCfC,PμCḡC-PiP°P»P°CfC¶ PI 13,6 %, 5,1 %, 9,3 % C,P° 9,9 % PīP°C†C-C°PSC,C-PI, C¶PeC- P»C-PeCfPIP°P»PēCfC¶ PIC-PrPīPsPIC-PrPSPs PeP°CḡPēPīCḡP°P·PēPSPsPj, PīP»P°C†PμP±Ps, CḡPēCfPīPμCḡPēPrPsPSPsPj C,P° P°CḡPēPīC-PīCḡP°P·PsP»PsPj. PμP°CḡPeC-PSCfPsPSC-P·Pj CfPīPsCfC,PμCḡC-PiP°PICfC¶ PI 13,6 %, 5,7 %, 22,1 % C,P° 5,3 % PīP°C†C-C°PSC,C-PI, C¶PeC- P»C-PeCfPIP°P»PēCfC¶ PIC-PrPīPsPIC-PrPSPs PeP°CḡPēPīCḡP°P·PēPSPsPj, PīP»P°C†PμP±Ps, CḡPēCfPīPμCḡPēPrPsPSPsPj C,P° P°CḡPēPīC-PīCḡP°P·PsP»PsPj. P°PēCfC,PsPSC-Cḡ CfPīPsCfC,PμCḡC-PiP°P»Pē PI 1,8 %, 0,2 %, 3,6 % C,P° 0,7 % PīP°C†C-C°PSC,C-PI, C¶PeC- P»C-PeCfPIP°P»PēCfC¶ PIC-PrPīPsPIC-PrPSPs PeP°CḡPēPīCḡP°P·PēPSPsPj, PīP»P°C†PμP±Ps, CḡPēCfPīPμCḡPēPrPsPSPsPj C,P° P°CḡPēPīC-PīCḡP°P·PsP»PsPj.

Pj PīP»P°C†PμP±PsPePsPSC,CḡPsP»CḡPsPIP°PSC-PNē C†P°CfC,PēPSC- PrPsCfP»C-PrP¶PμPSPSC¶ PrPsPIPīPsC,CḡPēPIP°P»PsC— PīC-PrC,CḡPēPjPePē PμC,,PμPeC,Cr C†P°CfC,PsC,P° P•PμPŸ CfC,P°PSPsPIPēP»P° 13,7 % Cr PīCḡCfPīC-, C¶PeP° PīCḡPēPNēPjP°P»P° PeP°CḡPēPīCḡP°P·PēPS, Cr PīPsCḡC-PIPSC¶PSPSC- P· 3,0 % Cr PīCḡCfPīC-, C¶PeP° PsC,CḡPēPjCfPIP°P»P° PīP»P°C†PμP±Ps. PḡPeP°C,PēP·C-C¶ CfPīPsCfC,PμCḡC-PiP°P»P°CfC¶ PI 3,9 % PīP°C†C-C°PSC,C-PI, C¶PeC- P»C-PeCfPIP°P»PēCfC¶ PeP°CḡPēPīCḡP°P·PēPSPsPj, PīCḡPsC,Pē 2,0 % PI PīCḡCfPīC-, C¶PeP° PīCḡPēPNēPjP°P»P° PīP»P°C†PμP±Ps. PμP°CḡPeC-PSCfPsPSC-P·Pj CfPīPsCfC,PμCḡC-PiP°PICfC¶ PI 7,8 % C,P° 1,0 % Cr PīCḡCfPīP°C..., C%Ps PIC-PrPīPsPIC-PrPSPs PīCḡPēPNēPjP°P»Pē PeP°CḡPēPīCḡP°P·PēPS C,P° PīP»P°C†PμP±Ps.

Pj PrPsCfP»C-PrP¶PμPSPSC- PSPμPīP°C,PēPIPSPēC... CfPēPjPīC,PsPjC-PI PīCḡPs P•PμPŸ PīPsPIC-PrPsPjP»C¶P»PsCfC¶ PI 14,3 % Cr PīCḡCfPīC-, C%Ps PīCḡPēPNēPjP°P»P° PeP°CḡPēPīCḡP°P·PēPS, C,P° PI 11,7 % Cr PīCḡCfPīC-, C%Ps P»C-PeCfPIP°P»P°CfC¶ CḡPēCfPīPμCḡPēPrPsPSPsPj. PḡPeP°C,PēP·C-C¶ CfPīPsCfC,PμCḡC-PiP°P»P°CfC¶ PI 10,0 % PīP°C†C-C°PSC,C-PI, C¶PeC- P»C-PeCfPIP°P»PēCfC¶ PeP°CḡPēPīCḡP°P·PēPSPsPj, C,P° PI 5,2 % Cr PīCḡCfPīC-, C%Ps PīCḡPēPNēPjP°P»P° CḡPēCfPīPμCḡPēPrPsPS. PμP°CḡPeC-PSCfPsPSC-P·Pj CfPīPsCfC,PμCḡC-PiP°PICfC¶ PI 5,2 % C,P° 7,4 % Cr PīCḡCfPīP°C..., C%Ps PIC-PrPīPsPIC-PrPSPs PīCḡPēPNēPjP°P»Pē PeP°CḡPēPīCḡP°P·PēPS C,P° CḡPēCfPīPμCḡPēPrPsPS. P·C-P»CḡCfC-CfC,Cḡ PIPēPīP°PrPeC-PI P•PμPŸ P±CfP»Pē P»PμPiPePsPiPs C†Pē CfPμCḡPμPrPSCḡPsPiPs CfC,CrPīPμPSC¶ C,C¶P¶PePsCfC,C- C,P° PePsPSC,CḡPsP»CḡPIP°P»PēCfC¶ P·PIPēC†P°PNēPSPēPjPē P»C-PeP°CḡCfCḡPePēPjPē P·P°CfPs±P°PjPē PrP»C¶ P»C-PeCfPIP°PSPSC¶ P•PμPŸ. P C-PIPμPSCḡ PīCḡPēPīPēPSPμPSPSC¶ P»C-PeCfPIP°PSPSC¶ C†PμCḡPμP· PIPēPīP°PrPePē P•PμPŸ, C%Ps PīPsPIbḡ™ C¶P·P°PSC- P· PμP»P , P±CfPI PSPēP·CḡPePēPj.

P'PμPSPsP·PSP° C,CḡPsPjP±PsPμPjP±PsP»C-C¶ (P'PŸP•)

P'PēPīP°PrPePē P'PŸP•, PIPeP»CḡC†PSPs P· PIPēPīP°PrPeP°PjPē P»PμPiPμPSPμPIPSC— PμPjP±PsP»C— C,P° C,CḡPsPjP±PsP·Cr PīP»PēP±PsPePēC... PIPμPS, CfPīPsCfC,PμCḡC-PiP°P»PēCfC¶ PīCḡPē PīCḡPēPNēPsPjC- P°PSC,PēPīCfPēC...PsC,PēPeC-PI. PḡP°CfC,PsC,P° PSPμPIC-PrPsPjP°.

$P_{\mu}C-PrPIP\ddot{e}C\%_{\circ}P_{\mu}PSP\ddot{e}PN_{\circ}C\overline{H}C-PIP_{\mu}PSC\overline{H}PiP_{\mu}C\ddot{+}C-PSPePsPIP\ddot{e}C...C,C\overline{H}P^{\circ}PSC\acute{I}P^{\circ}PjC-PSP^{\circ}P.$

$P_{\mu}C\overline{H}P\ddot{e}P\gg C-PeC\acute{r}PIP^{\circ}PSPSC-P^{\circ}PSC,P\ddot{e}PiC\acute{I}P\ddot{e}C...PsC,P\ddot{e}PeP^{\circ}PjP\ddot{e}C\ddot{+}P^{\circ}C\acute{I}C,Ps$
 $C\acute{I}PiPsC\acute{I}C,P_{\mu}C\overline{H}C-PiP^{\circ}C''C,C\overline{H}C\acute{I}C\overline{C}PiC-PrPIP\ddot{e}C\%_{\circ}P_{\mu}PSPSC\overline{C}C\overline{H}C-PIPSC\overline{C}$
 $PiP_{\mu}C\ddot{+}C-PSPePsPIP\ddot{e}C...C,C\overline{H}P^{\circ}PSC\acute{I}P^{\circ}PjC-PSP^{\circ}P.$
 $(P^{\circ}P\gg P^{\circ}PSC-PSP^{\circ}PjC-PSPsC,C\overline{H}P^{\circ}PSC\acute{I}C,,P_{\mu}C\overline{H}P^{\circ}P\cdot P\ddot{e}(P\overline{h}P\cdot P\ddot{y}),$
 $P^{\circ}C\acute{I}PiP^{\circ}C\overline{H}C,P^{\circ}C,P^{\circ}PjC-PSPsC,C\overline{H}P^{\circ}PSC\acute{I}C,,P_{\mu}C\overline{H}P^{\circ}P\cdot P\ddot{e}(P\overline{h}P\ddot{y}P\ddot{y})).PjPeP\gg C-PSC-C\ddot{+}PSP\ddot{e}C...$
 $PrPsC\acute{I}P\gg C-PrP\overline{P}_{\mu}PSPSC\overline{C}C...PeP^{\circ}C\overline{H}P\ddot{e}PiC\overline{H}P^{\circ}P\cdot P\ddot{e}PSC\acute{r}PiC-PrPIP\ddot{e}C\%_{\circ}P_{\mu}PSPSC\overline{C}$
 $C\overline{H}C-PIPSC\overline{C}P\overline{h}P\cdot P\ddot{y},P\overline{h}P\ddot{y}P\ddot{y}PIP\ddot{e}PSP\ddot{e}PeP^{\circ}P\gg PsPi2,2\%,1,6\%C,P^{\circ}0,4\%PiP^{\circ}C\ddot{+}C-C''PSC,C-Pi,$
 $C\overline{C}PeC-PIC-PrPiPsPIC-PrPSPsP\gg C-PeC\acute{r}PIP^{\circ}P\gg P\ddot{e}C\acute{I}C\overline{C}PeP^{\circ}C\overline{H}P\ddot{e}PiC\overline{H}P^{\circ}P\cdot P\ddot{e}PSPsPj,$
 $C\overline{H}P\ddot{e}C\acute{I}PiP_{\mu}C\overline{H}P\ddot{e}PrPsPSPsPjC,P^{\circ}PiP\gg P^{\circ}C\ddot{+}P_{\mu}P\pm Ps.P-PsPrP_{\mu}PS P\cdot PiP^{\circ}C\ddot{+}C-C''PSC,C-Pi,C\overline{C}PeC-$
 $P\gg C-PeC\acute{r}PIP^{\circ}P\gg P\ddot{e}C\acute{I}C\overline{C}PeP^{\circ}C\overline{H}P\ddot{e}PiC\overline{H}P^{\circ}P\cdot P\ddot{e}PSPsPj,PSP_{\mu}PjP^{\circ}PiP\overline{P}_{\mu}PsPrPSP\ddot{e}C...$
 $C\acute{r}C\overline{H}P^{\circ}P\overline{P}_{\mu}PSC\overline{H}PiP_{\mu}C\ddot{+}C-PSPeP\ddot{e}.$

$P-PjC-PSP\ddot{e}B PjP^{\circ}C\acute{I}P\ddot{e}C,C-P\gg P^{\circ}PiP^{\circ}C\ddot{+}C-C''PSC,C-Pi$

$PjPePsC\overline{H}PsC,PePsC,C\overline{H}P\ddot{e}PIP^{\circ}P\gg P\ddot{e}C...PrPsC\acute{I}P\gg C-PrP\overline{P}_{\mu}PSPSC\overline{C}C...PIC-PrPjC-C\ddot{+}P^{\circ}P\gg PsC\acute{I}C\overline{C}$
 $C,C\overline{H}PsC...P\ddot{e}P\pm C-P\gg C\overline{H}C\overline{C}P_{\mu}P\cdot P^{\circ}C\acute{I}P_{\mu}C\overline{H}P_{\mu}PrPSC''P\cdot P\pm C-P\gg C\overline{H}C\overline{C}P_{\mu}PSPSC\overline{C}B PjP^{\circ}C\acute{I}P\ddot{e}$
 $C,C-P\gg P^{\circ}PiP^{\circ}C\ddot{+}C-C''PSC,C-PiPiC\overline{H}C\acute{r}PiC-,C\%_{\circ}PsP\gg C-PeC\acute{r}PIP^{\circ}P\gg P^{\circ}C\acute{I}C\overline{C}$
 $PeP^{\circ}C\overline{H}P\ddot{e}PiC\overline{H}P^{\circ}P\cdot P\ddot{e}PSPsPj,C\acute{r}PiPsC\overline{H}C-PIPSC\overline{C}PSPSC-C\cdot PiC\overline{H}C\acute{r}PiPsC\overline{H},C\%_{\circ}Ps$
 $PiC\overline{H}P\ddot{e}PN_{\circ}PjP^{\circ}P\gg P^{\circ}PiP\gg P^{\circ}C\ddot{+}P_{\mu}P\pm Ps:1PePiC,P^{\circ}0,3PePiPIC-PrPiPsPIC-PrPSPs.Pj$
 $PrPsPiPiPsC,C\overline{H}P\ddot{e}PIP^{\circ}P\gg P\ddot{e}C...PrPsC\acute{I}P\gg C-PrP\overline{P}_{\mu}PSPSC\overline{C}C...PiC-PrC,C\overline{H}P\ddot{e}PjP^{\circ}PSPSC\overline{C}$
 $P_{\mu}C,,P_{\mu}PeC,C\acute{r}PSP_{\mu}C\acute{I}PiPsC\acute{I}C,P_{\mu}C\overline{H}C-PiP^{\circ}P\gg PsC\acute{I}C\overline{C}PeP\gg C-PSC-C\ddot{+}PSPsP\cdot PSP^{\circ}C\ddot{+}C\acute{r}C\%_{\circ}PsC-$
 $C\overline{H}C-P\cdot PSP\ddot{e}C\ddot{+}C-PiP\cdot PjC-PSC-PjP^{\circ}C\acute{I}P\ddot{e}C,C-P\gg P^{\circ}PIC-PrPIP\ddot{e}C...C-PrPSPsPiPsC\overline{H}C-PIPSC\overline{C}$
 $PrPsPeC-PSC\ddot{+}C\overline{C}P\gg C-PeC\acute{r}PIP^{\circ}PSPSC\overline{C}(1,1PePiPiC\overline{H}P\ddot{e}P\cdot P^{\circ}C\acute{I}C,PsC\acute{I}C\acute{r}PIP^{\circ}PSPSC-$
 $PeP^{\circ}C\overline{H}P\ddot{e}PiC\overline{H}P^{\circ}P\cdot P\ddot{e}PSC\acute{r}C,P^{\circ}B0,9PePiB\overline{H}''PiP\gg P^{\circ}C\ddot{+}P_{\mu}P\pm Ps).B\acute{K}P^{\circ}B PIC-PrPeC\overline{H}P\ddot{e}C,C-PN_{\circ}$
 $B C\acute{I}C,P^{\circ}PrC-C-B PrPsC\acute{I}P\gg C-PrP\overline{P}_{\mu}PSPSC\overline{C}PiC\overline{H}PsC,C\overline{C}PiPsPjB20C,P\ddot{e}P\overline{P}_{\mu}PSC-Pi$
 $P\gg C-PeC\acute{r}PIP^{\circ}PSPSC\overline{C}PeP^{\circ}C\overline{H}P\ddot{e}PiC\overline{H}P^{\circ}P\cdot P\ddot{e}PSPsPjC\acute{r}9,0\%PiP^{\circ}C\ddot{+}C-C''PSC,C-Pi$
 $C\overline{H}PsP\cdot PIP\ddot{e}PSC\acute{r}P\gg PsC\acute{I}C\overline{C}PiPsC,P_{\mu}PSC\ddot{+}C-PN_{\circ}PSPsPeP\gg C-PSC-C\ddot{+}PSPsP\cdot PSP^{\circ}C\ddot{+}C\acute{r}C\%_{\circ}P_{\mu}$
 $(P_{\mu}P\overline{L}P-)P\cdot P\pm C-P\gg C\overline{H}C\overline{C}P_{\mu}PSPSC\overline{C}PjP^{\circ}C\acute{I}P\ddot{e}C,C-P\gg P^{\circ}(PIP\ddot{e}P\cdot PSP^{\circ}C\ddot{+}C''C,C\overline{H}C\acute{I}C\overline{C}C\overline{C}Pe$
 $P\cdot P\pm C-P\gg C\overline{H}C\overline{C}P_{\mu}PSPSC\overline{C}PSP^{\circ}\geq 7\%),P^{\circ}PiC-PrC\ddot{+}P^{\circ}C\acute{I}PiPsPrPIC-PN_{\circ}PSPsC\acute{I}P\gg C-PiPsC-$
 $C\acute{I}C,P^{\circ}PrC-C-C\ddot{+}P_{\mu}C\overline{H}P_{\mu}P\cdot 20C,P\ddot{e}P\overline{P}_{\mu}PSC-PiPiC-C\acute{I}P\gg C\overline{C}PIC-PrPeC\overline{H}P\ddot{e}C,PsPiPs$
 $P\gg C-PeC\acute{r}PIP^{\circ}PSPSC\overline{C}PeP^{\circ}C\overline{H}P\ddot{e}PiC\overline{H}P^{\circ}P\cdot P\ddot{e}PSPsPjP_{\mu}P\overline{L}P-P\cdot P\pm C-P\gg C\overline{H}C\overline{C}P_{\mu}PSPSC\overline{C}PjP^{\circ}C\acute{I}P\ddot{e}$
 $C,C-P\gg P^{\circ}PjP^{\circ}P\gg P\ddot{e}9,8\%PiP^{\circ}C\ddot{+}C-C''PSC,C-Pi,C\overline{C}PeC-PiC\overline{H}PsPrPsPIP\overline{P}_{\mu}C\acute{I}PIP^{\circ}P\gg P\ddot{e}$
 $P\gg C-PeC\acute{r}PIP^{\circ}PSPSC\overline{C}PeP^{\circ}C\overline{H}P\ddot{e}PiC\overline{H}P^{\circ}P\cdot P\ddot{e}PSPsPj,PiC\overline{H}PsC,P\ddot{e}7,1\%PiP^{\circ}C\ddot{+}C-C''PSC,C-Pi,$
 $C\overline{C}PeC-P\pm C\acute{r}P\gg P\ddot{e}C\overline{H}P^{\circ}PSPPrPsPjC-P\cdot PsPIP^{\circ}PSC-PrP\gg C\overline{C}PiC\overline{H}P\ddot{e}PN_{\circ}PsPjC\acute{r}PiP\gg P^{\circ}C\ddot{+}P_{\mu}P\pm Ps.Pj$
 $PrPsC\acute{I}P\gg C-PrP\overline{P}_{\mu}PSPSC-PSP_{\mu}PiP^{\circ}C,P\ddot{e}PiPSP\ddot{e}C...C\acute{I}P\ddot{e}PjPiC,PsPjC-PiC\acute{I}P_{\mu}C\overline{H}P_{\mu}PrPSC\overline{C}$
 $P\cdot PjC-PSP^{\circ}PjP^{\circ}C\acute{I}P\ddot{e}C,C-P\gg P^{\circ}C\acute{I}C,P^{\circ}PSPsPiP\ddot{e}P\gg P^{\circ}B\overline{H}''0,3PePiC\acute{r}PiC\overline{H}C\acute{r}PiC-PiC\overline{H}P\ddot{e}PN_{\circ}PsPjC\acute{r}$
 $PeP^{\circ}C\overline{H}P\ddot{e}PiC\overline{H}P^{\circ}P\cdot P\ddot{e}PSC\acute{r}C,P^{\circ}0,6PePiC\acute{r}PiC\overline{H}C\acute{r}PiC-PiC\overline{H}P\ddot{e}PN_{\circ}PsPjC\acute{r}$
 $C\overline{H}P\ddot{e}C\acute{I}PiP_{\mu}C\overline{H}P\ddot{e}PrPsPSC\acute{r},P^{\circ}P_{\mu}P\overline{L}P-P\cdot P\pm C-P\gg C\overline{H}C\overline{C}P_{\mu}PSPSC\overline{C}B PjP^{\circ}C\acute{I}P\ddot{e}C,C-P\gg P^{\circ}$
 $C\acute{I}PiPsC\acute{I}C,P_{\mu}C\overline{H}C-PiP^{\circ}P\gg PsC\acute{I}C\overline{C}C\acute{r}6\%PiP^{\circ}C\ddot{+}C-C''PSC,C-PiP\cdot PiC\overline{H}C\acute{r}PiP\ddot{e},C\%_{\circ}Ps$
 $PiC\overline{H}P\ddot{e}PN_{\circ}PjP^{\circ}P\gg P\ddot{e}PeP^{\circ}C\overline{H}P\ddot{e}PiC\overline{H}P^{\circ}P\cdot P\ddot{e}PS,C,P^{\circ}C\acute{r}7,4\%PiP^{\circ}C\ddot{+}C-C''PSC,C-Pi,C\overline{C}PeC-$
 $P\gg C-PeC\acute{r}PIP^{\circ}P\gg P\ddot{e}C\acute{I}C\overline{C}C\overline{H}P\ddot{e}C\acute{I}PiP_{\mu}C\overline{H}P\ddot{e}PrPsPSPsPj.$

$P_{\mu}PsPrPsPIP\overline{P}_{\mu}PSPSC\overline{C}C-PSC,P_{\mu}C\overline{H}PIP^{\circ}P\gg C\acute{r}QT$

$PjPeP\gg C-PSC-C\ddot{+}PSPsPjC\acute{r}PrPsC\acute{I}P\gg C-PrP\overline{P}_{\mu}PSPSC-PrP\gg C\overline{C}PsC\ddot{+}C-PSPeP\ddot{e}$
 $PiPsPrPsPIP\overline{P}_{\mu}PSPSC\overline{C}C-PSC,P_{\mu}C\overline{H}PIP^{\circ}P\gg C\acute{r}QTPiC\overline{H}P\ddot{e}PiC\overline{H}P\ddot{e}PN_{\circ}PsPjC-$
 $PeP^{\circ}C\overline{H}P\ddot{e}PiC\overline{H}P^{\circ}P\cdot P\ddot{e}PSC\acute{r}PSP_{\mu}P\pm C\acute{r}P\gg PsPiPsPjC-C\ddot{+}P_{\mu}PSPsP\overline{P}_{\mu}PsPrPSPsPiPs$
 $PiPsPrPsPIP\overline{P}_{\mu}PSPSC\overline{C}C-PSC,P_{\mu}C\overline{H}PIP^{\circ}P\gg C\acute{r}QT C\acute{r}PiPsC\overline{H}C-PIPSC\overline{C}PSPSC-P\cdot PiP\gg P^{\circ}C\ddot{+}P_{\mu}P\pm Ps$
 $(PrP\ddot{e}Pi.C\overline{H}PsP\cdot PrC-P\gg B\ll P\alpha P^{\circ}C\overline{H}PjP^{\circ}PePsPrP\ddot{e}PSP^{\circ}PjC-PeP^{\circ}B\gg).P'C-PSC\overline{C}P\ddot{e}C...$
 $PeP\gg C-PSC-C\ddot{+}PSP\ddot{e}C...PrPsC\acute{I}P\gg C-PrP\overline{P}_{\mu}PSPSC\overline{C}C...PiPsPIC-PrPsPjP\gg C\overline{C}P\gg PsC\acute{I}C\overline{H}P\gg P\ddot{e}C\overline{C}P_{\mu}$

ПісПс ПрПмПеС-Р»СНПеР° ПІПєПіР°ПрПеС-ПІ ПСПмР·ПСП°С±ПСПсПіПс ПіПсПрПсПІП¶ПмПСПСЦІ
 С-ПSC,РмСНПІР°Р»Ср QT ПіСНПє ПіСНПєПмПсПсС- ПеР°СНПєПіСНП°Р·ПєПSCр. РмС-Пр С±Р°СІ
 ПрПсПіПіПсС,СНПєПІР°Р»ПсПіПс ПІС-ПрПеСНПєС,ПсПіПс ПіРмСНС-ПсПрСр Р»С-ПеСрПІР°ПСПСЦІ 3
 ПіР°С±С-С"ПSC,Пє (0,4 %) ПјР°Р»Пє СІПеПсСНПєПіПсПІР°ПСПєПм П·Р° С„ПсСНПсСрР»ПсСН
 Р'Р°Р·РмС,С,Р° С-ПSC,РмСНПІР°Р» QT > 500 ПјСІ, Р° ПсПрПєПс С-Р· ПСПєС... ПјР°ПІ
 С-ПSC,РмСНПІР°Р» QT Р· ПеПсСНПєПіСрПІР°ПСПСЦІПј Р±СНПєПрРмСНПєС±С-СІ > 500 ПјСІ.
 РмПсПрПсПІП¶ПмПСПСЦІ ПСП° > 60 ПјСІ СІПеПсСНПєПіПсПІР°ПСПсПіПс Р·Р° С„ПсСНПсСрР»ПсСН
 Р'Р°Р·РмС,С,Р° С-ПSC,РмСНПІР°Р»Ср QT ПІС-Пр ПІПєС...С-ПрПСПсПіПс СНС-ПІПСЦІ
 СІПіПсСІС,РмСНС-ПіР°Р»ПсСІСІ Ср 7 ПіР°С±С-С"ПSC,С-ПІ (1 %), Р° Ср 2 ПіР°С±С-С"ПSC,С-ПІ
 (0,3 %) вН" Р· ПеПсСНПєПіСрПІР°ПСПСЦІПј Р±СНПєПрРмСНПєС±С-СІ. Пј
 ПрПсПіПіПсС,СНПєПІР°Р»ПсПсСІР»С-ПрП¶ПмПСПSC- ПрР»СІ ПіС-ПрС,СНПєПјР°ПСПСЦІ
 РмС„РмПеС,Ср ПіС-Пр С±Р°СІ ПІС-ПрПеСНПєС,ПсС— СІС,Р°ПрС-С— ПіПсПрПсПІП¶ПмПСПСЦІ
 СІПеПсСНПєПіПсПІР°ПСПсПіПс Р·Р° С„ПсСНПсСрР»ПсСН Р'Р°Р·РмС,С,Р° С-ПSC,РмСНПІР°Р»Ср QT
 ПСП° > 60 ПјСІ ПІС-Пр ПІПєС...С-ПрПСПсПіПс СНС-ПІПСЦІ СІПіПсСІС,РмСНС-ПіР°Р»ПсСІСІ Ср
 12 ПіР°С±С-С"ПSC,С-ПІ (1,6 %), Р° С-ПSC,РмСНПІР°Р»Ср QT Р· ПеПсСНПєПіСрПІР°ПСПСЦІПј
 Р±СНПєПрРмСНПєС±С-СІ вН" Ср 4 ПіР°С±С-С"ПSC,С-ПІ (0,5 %). РмС-Пр С±Р°СІ
 ПіПсПрПІС-ПмПСПс СІР»С-ПіПсПіПс ПіРмСНС-ПсПрСр Р»С-ПеСрПІР°ПСПСЦІ
 ПіПсПрПсПІП¶ПмПСПСЦІ СІПеПсСНПєПіПсПІР°ПСПсПіПс Р·Р° С„ПсСНПсСрР»ПсСН Р'Р°Р·РмС,С,Р°
 С-ПSC,РмСНПІР°Р»Ср QT ПСП° > 60 ПјСІ ПІС-Пр ПІПєС...С-ПрПСПсПіПс СНС-ПІПСЦІ
 СІПіПсСІС,РмСНС-ПіР°Р»ПсСІСІ Ср 3 ПіР°С±С-С"ПSC,С-ПІ (3,1 %), СІПеС-
 Р»С-ПеСрПІР°Р»ПєСІСІ ПеР°СНПєПіСНП°Р·ПєПСПсПј, С,Р° Ср 2 ПіР°С±С-С"ПSC,С-ПІ (2 %),
 СІПеС- ПіСНПєПмПјР°Р»Пє ПіР»Р°С±РмР±Пс.

РмПсПІС-ПрПсПјР»РмПСПСЦІ ПіСНПс ПіС-ПрПсР·СНСНПІР°ПSC- ПіПсР±С-С±ПSC-
 СНРмР°ПеС±С-С—

РмПсПІС-ПрПсПјР»РмПСПСЦІ ПіСНПс ПіПсР±С-С±ПSC- СНРмР°ПеС±С-С— ПіС-СІР»СІ
 СНРмС"СІС,СНП°С±С-С— Р»С-ПеР°СНСІСНПеПсПіПс Р·Р°СІПсР±Ср ПјР°С" ПІР°П¶Р»ПєПІРм
 Р·ПСП°С±РмПСПСЦІ. Р|Рм ПрР°С" Р·ПсПіСр ПіСНПсПіПсПрПєС,Пє ПјПсПSC-С,ПсСНПєПСПі
 СІПіС-ПІПІС-ПрПСПсСєРмПСПСЦІ ПеПсСНПєСІС,СН/СНПєР·ПєПе ПіСНПє
 Р·Р°СІС,ПсСІСрПІР°ПСПSC- С±СНПсПіПс Р»С-ПеР°СНСІСНПеПсПіПс Р·Р°СІПсР±Ср.
 РмРмПрРєС±ПСПєПј С,Р° С„Р°СНПјР°С±РмПІС,ПєС±ПСПєПј ПіСНП°С±С-ПІПСПєР°Рј, Р°
 С,Р°ПеПсП¶ ПіР°С±С-С"ПSC,Р°Пј Р°Р±Пс С—С...ПSC-Пј Р·Р°ПеПсПСПСПєПј
 ПіСНПмПрСІС,Р°ПІПСПєР°Рј СІР»С-Пр ПіПсПІС-ПрПсПјР»СІС,Пє ПіСНПс СрСІС-
 ПІПєПіР°ПрПеПє ПіС-ПрПсР·СНСНПІР°ПСПєС... ПіПсР±С-С±ПСПєС... СНРмР°ПеС±С-Пм С,Р°
 ПІС-ПрСІСрС,ПСПсСІС,С- РмС„РмПеС,ПєПІПСПсСІС,С- Р»С-ПеР°СНСІСНПеПсПіПс Р·Р°СІПсР±Ср
 С±РмСНПмР· Р°ПІС,ПсПјР°С,ПєР·ПсПІР°ПSCр С-ПSC„ПсСНПсР°С±С-ПмПSCр СІПєСІС,РмПсСр Р·
 С„Р°СНПјР°ПеПсПСП°ПіР»СІПрСр Р·Р° ПіПсСІПєР»Р°ПСПСЦІПј: <https://aisf.dec.gov.ua>.

РјРмСНПсС-Пс ПіСНПєПрР°С,ПСПсСІС,С-.

5 СНПсПеС-ПІ.

ПјПсПІПє Р·Р±РмСНС-ПіР°ПСПСЦІ.

Р—Р±РмСНС-ПіР°С,Пє ПІ ПсСНПєПіС-ПСП°Р»СНПSC-Пм СрПіР°ПеПсПІС±С- ПрР»СІ
 Р·Р°С...ПєСІС,Ср ПІС-Пр ПІПіР»ПєПІСр СІПІС-С,Р»Р°.

P>C-PeP°CḡCḡCHPePëPNḡ P·P°CḡC-P± PSPµ PIPëPjP°PiP°C” CḡPḡPµC†C-P°P»CHPSPëC...
C,PµPjPḡPµCḡP°C,CḡCḡPSPëC... CḡPjPsPI P·P±PµCḡC-PiP°PSPSCḡ.

P—P±PµCḡC-PiP°C,Pë PI PSPµPrPsCḡC,CḡPḡPSPsPjCḡ PrP»Cḡ PrC-C,PµPNḡ PjC-CḡC†C-.

PJPḡP°PePsPIPeP°.

PµPs 1,5 PjPi P°P±Ps PḡPs 3 PjPi: PḡPs 7 PeP°PiCḡCḡP» Cḡ P±P»C-CḡC,PµCḡC-; PḡPs 1 P°P±Ps 4
P±P»C-CḡC,PµCḡPë PI PeP°CḡC,PSPSPSC-PNḡ CḡPḡP°PePsPIC†C-.

PµPs 4,5 PjPi P°P±Ps PḡPs 6 PjPi: PḡPs 7 PeP°PiCḡCḡP» Cḡ P±P»C-CḡC,PµCḡC-; PḡPs 4
P±P»C-CḡC,PµCḡPë PI PeP°CḡC,PSPSPSC-PNḡ CḡPḡP°PePsPIC†C-.

PḡP°C,PµPiPsCḡC-Cḡ PIC-PrPḡCḡCḡPeCḡ.

P—P° CḡPµC†PµPiC,PSPj.

P’PëCḡPsP±PSPëPe.

P’PhPḡ B«P“PµPrPµPsPS P C-C...C,PµCḡB», PJPiPsCḡC%PëPSP°.

**PḡC-CḡC†PµP·PSP°C...PsPrPḡPµPSPSCḡ PIPëCḡPsP±PSPëPeP° C,P° P°PrCḡPµCḡP°
PjC-CḡC†Cḡ PḡCḡPsPIP°PrPḡPµPSPSCḡ PNḡPsPiPs PrC-CḡP»CHPSPsCḡC,C-.**

H-1103, P’CḡPrP°PiPµC€C,, PICḡP». P”PµPjCḡPµC—, 19-21, PJPiPsCḡC%PëPSP°.