

Р†РќРЎРЎР РЈРЉРІР†РЇ

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

[illegible]

P•P—PḥPḱPŷPḥPᶑ

(EZANTAL)

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

PrC-CHC $\ddagger$ P $^{\circ}$ CH $\ddagger$ P μ C $\ddagger$ PsPIP $\ddot{e}$ PSP $^{\circ}$: P $\gg$ P μ CH $\ddagger$ PeP $^{\circ}$ PSC-PrP $\ddot{e}$ P $\ddot{i}$ C-PS (lercanidipine);

1 C,P°P±P»PμC,PεP° PjC-Cl'C,PëC,CH 10 PjPi P°P±Ps 20 PjPi P»PμCηPεP°PSC-PrPëPiC-PSCr
PiC-PrCηPsC...P»PsCηPëPrCr (Cr PIPëPiP»CΠPrC- P»PμCηPεP°PSC-PrPëPiC-PSCr
PiC-PrCηPsC...P»PsCηPëPrCr PiPμPjC-PiC-PrCηP°C,Cr);

PrPsPiPsPjC-P¶PSC- C¶PµC±PsPIPëPSPë: P»P°PeC,PsP·P°, PjPsPSPsPiC-PrC¶P°C,;
C†PµP»C¶P»PsP·P° PjC-PeC¶PsPeC¶PëC¶C, P°P»C-C±PSP°; PSP°C,C¶C-C¶
PeC¶PsC¶PeP°C¶PjPµP»PsP·P°; PïPsPIC-PrPsPS; PïPsP»C-C¶PsC¶P±P°C,; PjP°PiPSC-C¶
C¶C,PµP°C¶P°C,; PiC-PïC¶PsPjPµP»PsP·P°; PïPsP»C-PµC,PëP»PµPSPiP»C-PePsP»CH 6000
(PjP°PeC¶PsPiPsP» 6000); C,PëC,P°PSC† PrC-PsPeC¶PëPr (P• 171); C,P°P»CHPe; P·P°P»C-P·P°
PsPeC¶PëPr P¶PsPIC,PëP№ (E 172).

P>C-PeP°C₇C₇CH₂PeP° C₇PsC₇PjP°. P₇P°P±P»P₇C₇PeP₇, PIPeC₇P₇C₇-P₇P»C-PIPePsPIP₇C₇ PsP±PsP»PsPSPePsC₇.

PhCfPSPsPIPS- C,,C-P·PëPePs-C...C-PjC-C±PSC- PIP»P°CfC,PëPIPsCfC,C-:
PrPIPsPsPiCfPeP»C- C,P°P±P»PµC,PePë PeCfCfPiP»PsC— C,,PsCfPjPë P¶PsPIC,PsPiPs
PePsP»CHbPsCfCf, PIPeCfPëC,C- PïP»C-PIPePsPIPsCf PsP±PsP»PsPSPePsCf.

P_αP°C₇P_jP°PePsC,P_μC₇P°P_iP_μPIC,PëC_‡PSP° PiC₇C₇P_iP°

P'P»PsPeP°C,PsC₇Pë PeP°P»CHC†C-C"PIPëC... PeP°PSP°P»C-PI. PŸP_μP»P_μPeC,PëPIPSC-P°PSC,P°PiPsPSC-C†C,Pë PeP°P»CHC†C-C† P· PīP_μC₇P_μPIP°P¶(PSPsC† PrC-C"C† PSP° C†C†PrPëPSPë. P_μPsC...C-PrPSC- PrPëPiC-PrC₇PsPiC-C₇PëPrPëPSC†. P>P_μC₇PeP°PSC-PrPëPiC-PS. P_μPsPr P_hPŸP† PŸ08PŸ P_h13.

PαP°CḡPjP°PεPsP»PsPiiC≠PSi PIP»P°CḡC,PëPIPSCḡC,C-

$$P \propto P^{\circ} C_T P_j P^{\circ} P_e P_s P_r P_{\ddot{e}} P_s P^{\circ} P_j C - P_e P^{\circ}$$

P>PμCḡPeP°PSC-PrPëPīC-PS бḡ“ C†Pμ P°PSC,P°PiPsPSC-CḡC, PeP°P»CHC†C-Cḡ
PrPëPiC-PrCḡPsPiC-CḡPëPrPëPSPsPIPsC— PiCḡCrPīPë. P†PSPiC-P±CḡC”
C,CḡP°PSCḡPjPμPjP±CḡP°PSPSPëP№ PiPsC,C-Pe PeP°P»CHC†C-Cḡ PICḡPμCḡPμPrPëPSCḡ
PeP°CḡPrC-PsPjC-PsC†PëC,C-PI C,P° PiP»P°PrPePsPjбḡ™ CḡP·PsPIPëC... PeP»C-C,PëPS
CḡCrPrPëPS. PḡPμC...P°PSC-P·Pj P°PSC,PëPiC-PīPμCḡC,PμPSP·PëPIPSPsC— PrC-C—
P»PμCḡPeP°PSC-PrPëPīC-PSCḡ P·CḡPjPsPIP»PμPSPëP№ PiCḡCḡPjPëPj
CḡPμP»P°PeCḡCḡCḡPëPj PIPiP»PëPIPsPj PSP° PiP»P°PrPeC- Pjбḡ™ CḡP·Pë CḡCrPrPëPS, P·P°
CḡP°C...CḡPSPsPe CḡPsPiPs P·PjPμPSCḡCḡC”C,CḡCḡCḡ P·P°PiP°P»CHPSPëP№
CḡCrPrPëPSPSPëP№ PiPμCḡPëC,,PμCḡPëCḡPSPëP№ PsPiC-Cḡ. PḡPμP·PIP°PḡP°CḡCḡPë PSP°
PePsCḡPsC,PePëP№ PiPμCḡC-PsPr PSP°PiC-PIPIPëPIPμPrPμPSPSCḡ P· PiP»P°P·PjPë PeCḡPsPIC-,
P»PμCḡPeP°PSC-PrPëPīC-PS CḡPëPSPëC,CH PiCḡPsP»PsPSPIPsPIP°PSCḡ
P°PSC,PëPiC-PiPμCḡC,PμPSP·PëPIPSḡ PrC-Cḡ P·P° CḡP°C...CḡPSPsPe PIPëCḡPsPePsPiPs
PePsPμC,,C-C†C-C”PSC,P° PjPμPjP±CḡP°PSPSPsPiPs CḡPsP·PiPsPrC-P»Cḡ. P—P°PIPrCḡPePë
PIPëCḡPsPeC-P№ CḡCrPrPëPSPSC-P№ CḡPμP»PμPeC,PëPIPSḡCḡC,C- PiCḡPμPiP°CḡP°C, PSPμ
CḡPëPSPëC,CH PSPμPiP°C,PëPIPSPsC— C-PSPsC,CḡPsPiPSPsC— PrC-C—. P”PsCḡC,CḡP°
P°CḡC,PμCḡC-P°P»CHPSP° PiC-PiPsC,PμPSP·C-Cḡ P· CḡPμC,,P»PμPeC,PsCḡPSPsCḡ
C,P°C...C-PeP°CḡPrC-C”Cḡ PIPëPSPëPeP°C” CḡC-PrPePs P·P°PIPrCḡPePë
PiPsCḡC,CḡPiPsPIPsPjCḡ CḡPsP·PIPëC,PeCḡ PIP°P·PsPrPëP»P°C,P°C†C-C— PiCḡPë
PiCḡPëP№PsPjC- P»PμCḡPeP°PSC-PrPëPīC-PSCḡ.

PīPe C- PrP»Cḡ C-PSCḡPëC... P°CḡPëPjPμC,CḡPëCḡPSPëC... 1,4-
PrPëPiC-PrCḡPsPiC-CḡPëPrPëPSC-PI, P°PSC,PëPiC-PiPμCḡC,PμPSP·PëPIPSP°
P°PeC,PëPIPSḡ-CḡC,CH P»PμCḡPeP°PSC-PrPëPīC-PSCḡ PI PsCḡPSPsPIPSPsPjCḡ
PsP±CḡPjPsPIP»PμPSP° P№PsPiPs S-PμPSP°PSC,C-PsPjPμCḡPsPj.

PḡP»C-PSC-CḡPSP° PμC,,PμPeC,PëPIPSḡ-CḡC,CH C- P±PμP·PiPμPeP°
P·P°CḡC,PsCḡCḡPIP°PSPSCḡ P»PμCḡPeP°PSC-PrPëPīC-PSCḡ PI PrPsP·C- 10бḡ“20 PjPi 1 CḡP°P·
PSP° PrPsP±Cḡ P±CḡP»Pë PIPëPICḡPμPSC- PI PiPsPrPIC-P№PSPsPjCḡ CḡP»C-PiPsPjCḡ
PiP»P°C†PμP±Ps-PePsPSC,CḡPsP»CHPsPIP°PSPsPjCḡ PeP»C-PSC-CḡPSPsPjCḡ
PrPsCḡP»C-PrPḡPμPSPSC- (Cḡ CḡPePsPjCḡ 1200 PiP°C†C-C”PSC,C-PI PsC,CḡPëPjCḡPIP°P»Pë
P»PμCḡPeP°PSC-PrPëPīC-PS, 603 PiP°C†C-C”PSC,Pë PsC,CḡPëPjCḡPIP°P»Pë PiP»P°C†PμP±Ps)
C,P° PI P°PeC,PëPIPSPs PePsPSC,CḡPsP»CHPsPIP°PSPëC... C- PI
PSPμPePsPSC,CḡPsP»CHPsPIP°PSPëC... PrPsPIPiPsC,CḡPëPIP°P»PëC... PeP»C-PSC-CḡPSPëC...
PrPsCḡP»C-PrPḡPμPSPSCḡC... C-P· P·P°PiP°P»CHPSPsCḡ PeC-P»CHPeC-CḡC,Cḡ 3676
PiP°C†C-C”PSC,C-PI P· P°CḡC,PμCḡC-P°P»CHPSPsCḡ PiC-PiPμCḡC,PμPSP·C-C”Cḡ.

P”Ps P±C-P»CHCḡPsCḡC,C- PrPsCḡP»C-PrPḡPμPSCH P±CḡP»Pë P·P°P»CḡCḡPμPSC-
PiP°C†C-C”PSC,Pë P· PμCḡPμPSC†C-P°P»CHPSPsCḡ PiC-PiPμCḡC,PμPSP·C-C”Cḡ
P»PμPiPePsPiPs P°P±Ps PiPsPjC-CḡPSPsPiPs CḡC,CḡPiPμPSCḡ C,CḡPḡPePsCḡC,C-
(PIPeP»CḡCḡP°CḡCḡPë PiP°C†C-C”PSC,C-PI P»C-C,PSCHPsPiPs PIC-PeCḡ C,P° C...PIPsCḡPëC...
PSP° C†CḡPeCḡPsPIPëP№ PrC-P°P±PμC,), CḡPeC- PsC,CḡPëPjCḡPIP°P»Pë
P»PμCḡPeP°PSC-PrPëPīC-PS CḡPe PjPsPSPsC,PμCḡP°PiC-Cḡ P°P±Ps Cḡ
PePsPjP±C-PSP°C†C-C— P· C-PSPiC-P±C-C,PsCḡP°PjPë PḡPμPḡ, PrC-CḡCḡPμC,PëPeP°PjPë
P°P±Ps β-P±P»PsPeP°C,PsCḡP°PjPë.

P”PsPrP°C,PePsPIPs PrPs PeP»C-PSC-CḡPSPëC... PrPsCḡP»C-PrPḡPμPSCH,
PiCḡPsPIPμPrPμPSPëC... PrP»Cḡ PiC-PrC,PIPμCḡPrPḡPμPSPSCḡ C,PμCḡP°PiPμPIC,PëCḡPSPëC...
PiPsPeP°P·P°PSCH, PiPsPrP°P»CHCḡPμ PSPμPIPμP»PëPePμ
PSPμPePsPSC,CḡPsP»CHPsPIP°PSPμ, P°P»Pμ CḡP°PSPrPsPjC-P·PsPIP°PSPμ
PrPsCḡP»C-PrPḡPμPSPSCḡ PiCḡPμPiP°CḡP°C,Cḡ P·P° CḡCḡP°CḡC,Cḡ PiP°C†C-C”PSC,C-PI P·
C,CḡPḡPePsCḡ P°CḡC,PμCḡC-P°P»CHPSPsCḡ PiC-PiPμCḡC,PμPSP·C-C”Cḡ (CḡPμCḡPμPrPSC”
B± CḡC,P°PSPrP°CḡC,PSPμ PIC-PrC...PëP»PμPSPSCḡ CḡC-PIPSḡCḡ

PrC-P°CfC,PsP»C-C†PSPsPiPs P°CfC,PμCfC-P°P»CHbPSPsPiPs C,PëCfPeCr 114,5 B± 3,7 PjPj CfC, CfC, PiCfPsPrPμPjPsPSCfC,CfCrPIP°P»Ps PSPsCfPjP°P»C-P·P°C†C-Cf P°CfC,PμCfC-P°P»CHbPSPsPiPs C,PëCfPeCr Cr 40 % C-P· 25 PïP°C†C-C"PSC,C-PI, CΠPeC-PiCfPëPNëPjP°P»Pë PiCfPμPïP°CfP°C, Cr PrPsP·C- B B B B B B B B 20 PjPi 1 CfP°P· PSP° PrPsP±Cr, C,P° Cr 56 % C-P· 25 PïP°C†C-C"PSC,C-PI, CΠPeC- PiCfPëPNëPjP°P»Pë P»PμCfPeP°PSC-PrPëPiC-PS Cr PrPsP·C- 10 PjPi 2 CfP°P·Pë PSP° PrPsP±Cr. Pj PïPsPrPIC-PNëPSPsPjCr CfP»C-PiPsPjCr CfP°PSPPrPsPjC-P·PsPIP°PSPsPjCr PïP»P°C†PμP±Ps- PePsPSC,CfPsP»CHbPsPIP°PSPsPjCr PrPsCfP»C-PrP¶PμPSPSC- Cr PïP°C†C-C"PSC,C-PI P· C-P·PsP»CHbPsPIP°PSPsCf CfPëCfC,PsP»C-C†PSPsCf PiC-PiPμCfC,PμPSP·C-C"Cf P»PμCfPeP°PSC-PrPëPiC-PS PμC,,PμPeC,PëPIPSPs P·PSPëP¶CfPIP°PI CfPëCfC,PsP»C-C†PSPëPNë P°CfC,PμCfC-P°P»CHbPSPëPNë C,PëCfPe C-P· CfPμCfPμPrPSCbPsPiPs PïPsC†P°C,PePsPIPsPiPs P·PSP°C†PμPSPSCΠ 172,6 B± 5,6 PjPj CfC, CfC, PrPs 140,2 B± 8,7 PjPj CfC, CfC,.

P"C-C,Pë

PμP»C-PSC-C†PSC- PrPsCfP»C-PrP¶PμPSPSCΠ Cr PrPëC,CΠC†C-PNë PïPsPiCrP»CΠC†C-C— PSPμ PiCfPsPIPsPrPëP»PëCfCΠ.

PαP°CfPjP°PePsPeC-PSPμC,PëPeP°

PbP±CfPsCfP±C†C-CΠ

P>PμCfPeP°PSC-PrPëPiC-PS PïPsPIPSC-CfC,Cf PICfPjPsPeC,CrC"C,CHbCfCΠ PiC-CfP»CΠ PiCfPëPNëPsPjCr PIPSCfC,CfC-C€PSCbPs Cr PrPsP·C- 10b¶"20 PjPi, PjP°PeCfPëPjP°P»CHbPSP° PePsPSC†PμPSC,CfP°C†C-CΠ Cr PïP»P°P·PjC- PeCfPsPIC- (C_{max}) 3,30 PSPi/PjP» B± 2,09 Cf. PI. C,P° 7,66 PSPi/PjP» B± 5,90 Cf. PI. PIC-PrPïPsPIC-PrPSPs CfPμC"CfC,CfCrC"C,CHbCfCΠ PiCfPëP±P»PëP·PSPs C†PμCfPμP· 1,5b¶"3 PiPsPrPëPSPë.

P"PIP° PμPSP°PSC,C-PsPjPμCfPë P»PμCfPeP°PSC-PrPëPiC-PSCr PrPμPjPsPSCfC,CfCrCfC,CHb PïPsPrC-P±PSPëPNë PiCfPsC,,C-P»CHb CfC-PIPSCΠ Cr PïP»P°P·PjC- PeCfPsPIC-: C†P°Cf PrPsCfCΠPiPSPμPSPSCΠ C_{max} C,P°PePëPNë CfP°PjPëPNë, C_{max} C,P° AUC (PïP»PsC%°P° PiC-Pr C,,P°CfPjP°PePsPeC-PSPμC,PëC†PSPsCf PeCfPëPIPsCf B«PePsPSC†PμPSC,CfP°C†C-CΠ- C†P°CfB») PI CfPμCfPμPrPSCbPsPjCr Cr 1,2 CfP°P·P° PIPëC%°C- PrP»CΠ S- PμPSP°PSC,C-PsPjPμCfP°, C- PïPμCfC-PsPr PSP°PiC-PIPIPëPIPμPrPμPSPSCΠ PrPIPsC... PμPSP°PSC,C-PsPjPμCfC-PI, PïPs CfCrC,C-, PsPrPSP°PePsPIPëPNë. P'P·P°C"PjPsPiPμCfPμC,PIPsCfPμPSPSCΠ PμPSP°PSC,C-PsPjPμCfC-PI in vivo PSPμ CfPiPsCfC,PμCfC-PiP°P»PsCfCf.

P'PSP°CfP»C-PrPsPe PIPëCfPsPePsPiPs PjPμC,P°P±PsP»C-P·PjCr PiCfPë PiPμCfPIPëPSPSPsPjCr PiCfPsC...PsPrP¶PμPSPSC- C†PμCfPμP· PïPμC†C-PSPeCr P°P±CfPsP»CfC,PSP° P±C-PsPrPsCfC,CrPïPSC-CfC,CHb P»PμCfPeP°PSC-PrPëPiC-PSCr, P·P°CfC,PsCfPsPIP°PSPsPiPs PïP°C†C-C"PSC,PsPj PiC-CfP»CΠ PIP¶PëPIP°PSPSCΠ C—P¶C-, CfC,P°PSPsPIPëC,CHb PiCfPëP±P»PëP·PSPs 10 %, C...PsC†P° PIPsPSP° P·PjPμPSC€CfPIP°P»P°CfCΠ PrPs 1/3 C†CHbPsPiPs P·PSP°C†PμPSPSCΠ, CΠPeC%°Ps PiCfPμPïP°CfP°C, P·P°CfC,PsCfPsPICrPIP°P»Pë P·PrPsCfPsPIC- PrPsP±CfPsPIPsP»CHbC†C- PSP°C,C%°Pμ. PïPeC%°Ps PiCfPμPïP°CfP°C, PiCfPëPNëPjP°C,Pë PSPμ PiC-P·PSC-C€Pμ PSC-P¶ C†PμCfPμP· 2 PiPsPrPëPSPë PiC-CfP»CΠ PIP¶PëPIP°PSPSCΠ PrCrP¶Pμ P¶PëCfPSPsC— C—P¶C-, PNëPsPiPs P±C-PsPrPsCfC,CrPïPSC-CfC,CHb P·P±C-P»CHbC€CfC"C,CHbCfCΠ Cr 4 CfP°P·Pë. PÿPsPjCr P»PμCfPeP°PSC-PrPëPiC-PS CfP»C-Pr PiCfPëPNëPjP°C,Pë PrPs PIP¶PëPIP°PSPSCΠ C—P¶C-.

P PsP·PiPsPrC-P»

P PsP·PiPsPrC-P» C-P· PiP»P°P·PjPë PeCḡPsPIC- Cr C,PeP°PSPëPSPë C,P° PsCḡPiP°PSPë C”
C€PiPëPrPePëPj C,P° PsP±C€PëCḡPSPëPj. PŸC,CrPiC-PSCH P·PiBḡ™ CḡP·CrPiP°PSPSCḡ
P»PμCḡPeP°PSC-PrPëPiC-PSCr P· P±C-P»PeP°PjPë CḡPëCḡPsPiP°C,PePë PeCḡPsPIC-
PiPμCḡPμPiPëC%CrC” 98 %. PhCḡPeC-P»CHPePë CḡC-PIPμPSCH P±C-P»PeP° Cr PiP»P°P·PjC-
PeCḡPsPIC- P·PjPμPSC€PμPSPëPnë Cr PiP°C†C-C”PSC,C-PI C-P· C,CḡP¶PePëPjPë
PiPsCḡCrC€PμPSPSCḡPjPë C,,CrPSPeC†C-C— PSPëCḡPsPe C,P° PiPμC±C-PSPePë, PIC-P»CHPSP°
C,,CḡP°PeC†C-Cḡ PiCḡPμPiP°CḡP°C,Cr PjPsP¶Pμ P·P±C-P»CHC€CrPiP°C,PëCḡCḡ.

P’C-PSc,CḡP°PSCḡC,,PsCḡPjP°C†C-Cḡ

P»PμCḡPeP°PSC-PrPëPiC-PS PμPeCḡC,PμPSCḡPëPiPSPs PjPμC,P°P±PsP»C-P·CrC”C,CHCḡCḡ
C-P·PsC,,PμCḡPjPμPSC,PsPj CYP3Pḡ4, PSPμP·PjC-PSPμPSPëPnë PiCḡPμPiP°CḡP°C, Cr CḡPμC±C-
C±Pë PeP°P»C- PSPμ PiPëCḡPiP»CḡC”C,CHCḡCḡ. P’C-PS PiPμCḡPμC,PIPsCḡCḡC”C,CHCḡCḡ
PiPμCḡPμPiP°P¶PSPs Cr PSPμP°PeC,PëPiPSC- PjPμC,P°P±PsP»C-C,Pë, PiCḡPëP±P»PëP·PSPs 50
% PiCḡPëPnëPSCḡC,PsC— PrPsP·Pë PiPëPiPsPrPëC,CHCḡCḡ C-P· CḡPμC±PμCḡ.

P·PeCḡPiPμCḡPëPjPμPSC,Pë *in vitro* P· P»CHPrCḡCHPePëPjPë PjC-PeCḡPsCḡPsPjP°PjPë
PiPμC±C-PSPePë CḡPIC-PrC±P°C,CH, C%Ps P»PμCḡPeP°PSC-PrPëPiC-PS PrPμC%Ps
PiCḡPëPiPSC-C±CrC” CYP3A4 C,P° CYP2D6 PiCḡPë PePsPSC†PμPSC,CḡP°C†C-CḡC...
PIC-PrPiPsPIC-PrPSPs Cr 160 C- 40 CḡP°P·C-PI PiPëC%PëC..., PSC-P¶ PnëPsPiPs C_{max} Cr
PiP»P°P·PjC- PeCḡPsPIC-, CḡPeP° PrPsCḡCḡPiP°C”C,CHCḡCḡ PiC-CḡP»Cḡ PiCḡPëPnëPsPjCr
PrPsP·Pë 20 PjPi. PḡCḡC-Pj C†CHPsPiPs, PrPsCḡP»C-PrP¶PμPSPSCḡ P»C-PeP°CḡCḡCHPePsC—
PiP·P°C”PjPsPrC-C— Cr P»CHPrPμPnë PiCḡPsPrPμPjPsPSCḡC,CḡCrPiP°P»Pë, C%Ps
P»PμCḡPeP°PSC-PrPëPiC-PS Cr PiP»P°P·PjC- PeCḡPsPIC- PSPμ P·PjC-PSCHC” CḡC-PIPμPSCH
PjC-PrP°P·PsP»P°PjCr, C,PëPiPsPiPsPiPs CḡCrP±CḡC,CḡP°C,Cr CYP3A4, P°P±Ps
PjPμC,PsPiCḡPsP»PsP»Cr, C,PëPiPsPiPsPiPs CḡCrP±CḡC,CḡP°C,Cr CYP2D6; C,P°PePëPj
C±PëPSPsPj, PiCḡPë P·P°CḡC,PsCḡCrPiP°PSPSC- P»PμCḡPeP°PSC-PrPëPiC-PSCr Cr
C,PμCḡP°PiPμPIC,PëC±PSPëC... PrPsP·P°C... PSPμ PsC±C-PeCrC”C,CHCḡCḡ
P±C-PsC,CḡP°PSCḡC,,PsCḡPjP°C†C-C— PiCḡPμPiP°CḡP°C,C-PI, C%Ps
PjPμC,P°P±PsP»C-P·CrCḡC,CHCḡCḡ CYP3A4 P°P±Ps CYP2D6.

P’PëPiPμPrPμPSPSCḡ

P·P»C-PjC-PSP°C†C-Cḡ PiCḡPsC...PsPrPëC,CH PI PsCḡPSPsPiPSPsPjCr C€P»CḡC...PsPj
P±C-PsC,CḡP°PSCḡC,,PsCḡPjP°C†C-C—. PŸPμCḡPμPrPSC-Pnë C,PμCḡPjC-PSP°P»CHPSPëPnë
PiPμCḡC-PsPr PSP°PiC-PIPiPëPiPμPrPμPSPSCḡ CḡC,P°PSPsPiPëC,CH 8Bḡ“10 PiPsPrPëPS, P°
C,PμCḡP°PiPμPIC,PëC±PSP° PrC-Cḡ C,CḡPëPiP°C” 24 PiPsPrPëPSPë PiPSP°CḡP»C-PrPsPe
PiPëCḡPsPePsPiPs CḡC,CrPiPμPSCḡ P·PiBḡ™ CḡP·CrPiP°PSPSCḡ P»PμCḡPeP°PSC-PrPëPiC-PSCr
P· P»C-PiC-PrP°PjPë PeP»C-C,PëPSPSPëC... PjPμPjP±CḡP°PS. PμCḡPë PiPsPIC,PsCḡPSPsPjCr
P·P°CḡC,PsCḡCrPiP°PSPSC- PeCrPjCrP»CḡC†C-Cḡ PSPμ CḡPiPsCḡC,PμCḡC-PiP°P»P°CḡCH.

P»C-PSC-PnëPSC-CḡC,CH / PSPμP»C-PSC-PnëPSC-CḡC,CH

PμCḡPë PiPμCḡPsCḡP°P»CHPSPsPjCr P·P°CḡC,PsCḡCrPiP°PSPSC-
P»PμCḡPeP°PSC-PrPëPiC-PSCr PnëPsPiPs PePsPSC†PμPSC,CḡP°C†C-Cḡ Cr PiP»P°P·PjC-
PeCḡPsPIC- PSPμ C” PiCḡCḡPjPs PiCḡPsPiPsCḡC†C-PnëPSPsCḡ PrPs PiCḡPëPnëPSCḡC,PsC—
PrPsP·Pë (PSPμP»C-PSC-PnëPSP° PeC-PSPμC,PëPeP°). PμC-CḡP»Cḡ PiCḡPëPnëPsPjCr 10 PjPi, 20
PjPi C- 40 PjPi C_{max} Cr PiP»P°P·PjC- PeCḡPsPIC-, CḡPeC- CḡPiPsCḡC,PμCḡC-PiP°P»PëCḡCḡ,
PjP°P»Pë CḡPiC-PIPIC-PrPSPsC€PμPSPSCḡ 1:3:8, P° PiP»PsC%C- PiC-Pr PeCḡPëPiPëPjPë
P·P°P»PμP¶PSPsCḡC,C- PePsPSC†PμPSC,CḡP°C†C-C— Cr PiP»P°P·PjC- PeCḡPsPIC- PIC-Pr
C±P°CḡCr PjP°P»Pë CḡPiC-PIPIC-PrPSPsC€PμPSPSCḡ 1:4:18, C%Ps PiPeP°P·CrC” PSP°
PiPsCḡC,CrPiPsPiPμ PSP°CḡPëC±PμPSPSCḡ PjPμC,P°P±PsP»C-P·PjCr PiCḡPë PiPμCḡC€PsPjCr

- $\text{P}_{\mu}\text{C}-\text{PrPIP}\ddot{\text{C}}\%_{\text{P}}\text{P}_{\mu}\text{PSP}^{\circ}\text{C}\ddagger\text{C}\acute{\text{r}}\text{C},\text{P}\gg\text{P}\ddot{\text{e}}\text{PIC}-\text{C}\acute{\text{r}}\text{C},\text{CH}\text{PrPs}\text{P}\gg\text{P}_{\mu}\text{C}\ddot{\text{H}}\text{PeP}^{\circ}\text{PSC}-\text{PrP}\ddot{\text{e}}\text{P}\ddot{\text{i}}\text{C}-\text{PSC}\acute{\text{r}}\text{P}^{\circ}\text{P}\pm\text{PsPrPs}\text{P}\pm\text{C}\acute{\text{r}}\text{PrCH}\ddot{\text{e}}-\text{C}\ddot{\text{I}}\text{PePsPiPs}\text{PePsPjPiPsPSP}_{\mu}\text{PSC},\text{P}^{\circ}\text{P}\gg\text{C}-\text{PeP}^{\circ}\text{C}\ddot{\text{H}}\text{C}\acute{\text{r}}\text{CH}\text{PePsPiPs}\text{P}\cdot\text{P}^{\circ}\text{C}\acute{\text{r}}\text{Ps}\pm\text{C}\acute{\text{r}};$

- $\text{PsP}\pm\text{C}\acute{\text{r}}\text{C},\text{C}\ddot{\text{H}}\text{C}\acute{\text{r}}\text{PeC}\ddagger\text{C}-\text{C}\ddot{\text{I}}\text{PIP}\ddot{\text{e}}\text{PIC}-\text{PrPSPsPiPs}\text{PIC}-\text{PrPrC}-\text{P}\gg\text{C}\acute{\text{r}}\text{P}\gg\text{C}-\text{PIPsPiPs}\text{C}\mathbb{E}\text{P}\gg\text{C}\acute{\text{r}}\text{PSPsC}\ddagger\text{PeP}^{\circ};$

- $\text{PSP}_{\mu}\text{P}\gg\text{C}-\text{PePsPIP}^{\circ}\text{PSP}^{\circ}\text{P}\cdot\text{P}^{\circ}\text{C}\acute{\text{r}}\text{C},\text{C}-\text{PN}_{\mathbb{e}}\text{PSP}^{\circ}\text{C}\acute{\text{r}}\text{P}_{\mu}\text{C}\ddot{\text{H}}\text{C}\ddagger\text{P}_{\mu}\text{PIP}^{\circ}\text{PSP}_{\mu}\text{PrPsC}\acute{\text{r}}\text{C},\text{P}^{\circ}\text{C},\text{PSC}-\text{C}\acute{\text{r}}\text{C},\text{CH}\ddot{\text{e}};$

- $\text{PSP}_{\mu}\text{C}\acute{\text{r}}\text{C},\text{P}^{\circ}\text{P}\pm\text{C}-\text{P}\gg\text{CH}\text{PSP}^{\circ}\text{C}\acute{\text{r}}\text{C},\text{P}_{\mu}\text{PSPsPeP}^{\circ}\text{C}\ddot{\text{H}}\text{PrC}-\text{C}\ddot{\text{I}}\text{C}\ddagger\text{P}\ddot{\text{e}}\text{PSP}_{\mu}\text{C}\%_{\text{Ps}}\text{PrP}^{\circ}\text{PIPSC}-\text{PN}_{\mathbb{e}}(\text{PiC}\ddot{\text{H}}\text{PsC},\text{C}\ddot{\text{I}}\text{PiPsPj}\text{ }1\text{ } \text{PjC}-\text{C}\acute{\text{r}}\text{C}\ddot{\text{I}}\text{C}\ddagger\text{C}\ddot{\text{I}})\text{C}-\text{PSC},,\text{P}^{\circ}\text{C}\ddot{\text{H}}\text{PeC},\text{PjC}-\text{PsPeP}^{\circ}\text{C}\ddot{\text{H}}\text{PrP}^{\circ};$

- $\text{C},\text{C}\ddot{\text{I}}\text{P}\P\text{PeP}^{\circ}\text{PiP}_{\mu}\text{C}\ddagger\text{C}-\text{PSPePsPIP}^{\circ}\text{PSP}_{\mu}\text{PrPsC}\acute{\text{r}}\text{C},\text{P}^{\circ}\text{C},\text{PSC}-\text{C}\acute{\text{r}}\text{C},\text{CH}\ddot{\text{e}};$

- $\text{C},\text{C}\ddot{\text{I}}\text{P}\P\text{PeP}^{\circ}\text{PSP}\ddot{\text{e}}\text{C}\ddot{\text{H}}\text{PePsPIP}^{\circ}\text{PSP}_{\mu}\text{PrPsC}\acute{\text{r}}\text{C},\text{P}^{\circ}\text{C},\text{PSC}-\text{C}\acute{\text{r}}\text{C},\text{CH}\ddot{\text{e}}(\text{PeP}\gg\text{C}-\text{C}\ddot{\text{H}}\text{P}_{\mu}\text{PSC}\acute{\text{r}}\text{PeC}\ddot{\text{H}}\text{P}_{\mu}\text{P}^{\circ}\text{C},\text{P}\ddot{\text{e}}\text{PSC}-\text{PSC}\acute{\text{r}}<30\text{ } \text{PjP}\gg/\text{C}\dots\text{PI}),\text{PIPeP}\gg\text{C}\ddot{\text{H}}\text{C}\ddagger\text{P}^{\circ}\text{C}\ddot{\text{H}}\text{C}\ddagger\text{P}\ddot{\text{e}}\text{PiP}^{\circ}\text{C}\ddagger\text{C}-\text{C}''\text{PSC},\text{C}-\text{PI}\text{PSP}^{\circ}\text{PiPuPiPsPrC}-\text{P}^{\circ}\text{P}\gg\text{C}-\text{P}\cdot\text{C}-;$

- PsPrPSPsC $\ddagger$ P $^{\circ}$ Cl $\ddagger$ PSP μ P $\cdot$ P $^{\circ}$ Cl $\ddagger$ C,PsCl $\ddagger$ CrPIP $^{\circ}$ PSPSC $\ddagger$ P $\cdot$ P $\ddot{\text{I}}$ PsC,CrP $\ddagger$ PSP $\ddot{\text{P}}$ PjP $\ddot{\text{e}}$
C-PSPiC-P $\pm$ C-C,PsC $\ddot{\text{H}}$ P $^{\circ}$ PjP $\ddot{\text{e}}$ CYP3P $\ddot{\text{H}}$ 4, C $\ddagger$ P $\ddot{\text{e}}$ PeP $\gg$ PsCl $\ddagger$ PiPsC $\ddot{\text{H}}$ P $\ddot{\text{e}}$ PSPsPj, PiC $\ddot{\text{H}}$ P μ PN $\ddot{\text{e}}$ PiC $\ddot{\text{H}}$ CrC,PsPj
C $\ddagger$ P $\ddot{\text{e}}$ PiC $\ddot{\text{H}}$ P μ PN $\ddot{\text{e}}$ PiC $\ddot{\text{H}}$ CrC,PsPIP $\ddot{\text{e}}$ Pj Cl $\ddagger$ PsPePsPj.

***P $\cdot$ P $^{\circ}$ C $\ddot{\text{H}}$ PjPsPrC-C $\ddagger$ P $\cdot$ C-PSC $\ddot{\text{E}}$ P $\ddot{\text{e}}$ PjP $\ddot{\text{e}}$ P $\gg$ C-PeP $^{\circ}$ C $\ddot{\text{H}}$ Cl $\ddagger$ CHPeP $\ddot{\text{e}}$ PjP $\ddot{\text{e}}$ P $\cdot$ P $^{\circ}$ Cl $\ddagger$ PsP $\pm$ P $^{\circ}$ PjP $\ddot{\text{e}}$ C,P $^{\circ}$
C-PSC $\ddot{\text{E}}$ C- PIP $\ddot{\text{e}}$ PrP $\ddot{\text{e}}$ PIP $\cdot$ P $^{\circ}$ C $\ddot{\text{H}}$ PjPsPrC-PN $\ddot{\text{e}}$***

P μ C $\ddot{\text{H}}$ PsC,P $\ddot{\text{e}}$ PiPsPeP $^{\circ}$ P $\cdot$ P $^{\circ}$ PSPs PsPrPSPsC $\ddagger$ P $^{\circ}$ Cl $\ddagger$ PSP μ P $\cdot$ P $^{\circ}$ Cl $\ddagger$ C,PsCl $\ddagger$ C $\ddot{\text{e}}$ PIP $^{\circ}$ PSPSC $\ddagger$

P $\ddagger$ PSPiC-P $\pm$ C-C,PsC $\ddot{\text{H}}$ P $\ddot{\text{e}}$ CYP3A4

P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PS PjP μ C,P $^{\circ}$ P $\pm$ PsP $\gg$ C-P $\cdot$ CrC $\ddot{\text{H}}$ C,CHC $\ddagger$ Cl $\ddagger$ P $\ddot{\text{I}}$ C-Pr PrC-C $\ddot{\text{H}}$
C $\ddot{\text{H}}$,P μ C $\ddot{\text{H}}$ PjP μ PSC,Cr CYP3A4, C,PsPjCr C-PSPiC-P $\pm$ C-C,PsC $\ddot{\text{H}}$ P $\ddot{\text{e}}$ C,P $^{\circ}$ C-PSPCr $\ddagger$ PeC,PsC $\ddot{\text{H}}$ P $\ddot{\text{e}}$
C $\ddagger$ CHPsPiPs C $\ddot{\text{H}}$,P μ C $\ddot{\text{H}}$ PjP μ PSC,Cr, Cl $\ddagger$ PeC- P $\ddot{\text{I}}$ C $\ddot{\text{H}}$ P $\ddot{\text{e}}$ PN $\ddot{\text{e}}$ PjP $^{\circ}$ CHC,CH PsPrPSPsC $\ddagger$ P $^{\circ}$ Cl $\ddagger$ PSPs P $\cdot$
P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PSPsPj, PjPsP $\ddagger$ CrC,CH PIP $\ddot{\text{e}}$ P $\gg$ P $\ddot{\text{e}}$ PIP $^{\circ}$ C,P $\ddot{\text{e}}$ PSP $^{\circ}$ PjP μ C,P $^{\circ}$ P $\pm$ PsP $\gg$ C-P $\cdot$ Pj
C,P $^{\circ}$ PIP $\ddot{\text{e}}$ PIP μ PrP μ PSPSC $\ddagger$ P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PSC $\ddagger$. P $\ddot{\text{H}}$ PsCl $\ddagger$ P $\gg$ C-PrP $\ddagger$ P μ PSPSC $\ddagger$
PIP $\cdot$ P $^{\circ}$ C $\ddot{\text{H}}$ PjPsPrC-C— P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PSC $\ddagger$ P $\cdot$ P $\ddot{\text{I}}$ PsC,CrP $\ddagger$ PSP $\ddot{\text{P}}$ Pj
C-PSPiC-P $\pm$ C-C,PsC $\ddot{\text{H}}$ PsPj CYP3A4 PeP μ C,PsPePsPSP $^{\circ}$ P $\cdot$ PsP $\gg$ PsPj
P $\ddot{\text{I}}$ C $\ddot{\text{H}}$ PsPrP μ PjPsPSC $\ddagger$ C,C $\ddot{\text{H}}$ CrPIP $^{\circ}$ P $\gg$ P $\ddot{\text{e}}$ P $\cdot$ PSP $^{\circ}$ C $\ddagger$ PSP μ P $\ddot{\text{I}}$ C-PrPIP $\ddot{\text{e}}$ C $\%_{\text{P}}$ P μ PSPSC $\ddagger$ C $\ddot{\text{H}}$ C-PIPSC $\ddagger$
P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PSC $\ddagger$ Cr P $\ddot{\text{I}}$ P $\gg$ P $^{\circ}$ P $\cdot$ PjC- PeC $\ddot{\text{H}}$ PsPIC- (15-PeC $\ddot{\text{H}}$ P $^{\circ}$ C,PSP μ
P $\ddot{\text{I}}$ C-PrPIP $\ddot{\text{e}}$ C $\%_{\text{P}}$ P μ PSPSC $\ddagger$ P $\ddot{\text{I}}$ PsPeP $^{\circ}$ P $\cdot$ PSP $\ddot{\text{e}}$ PeP $^{\circ}$ AUC C,P $^{\circ}$ 8-PeC $\ddot{\text{H}}$ P $^{\circ}$ C,PSP μ
P $\ddot{\text{I}}$ C-PrPIP $\ddot{\text{e}}$ C $\%_{\text{P}}$ P μ PSPSC $\ddagger$ C max P μ CrC,PsPjP μ C $\ddot{\text{H}}$ P $^{\circ}$ S-P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PSC $\ddagger$).

P $\ddot{\text{Y}}$ P $\gg$ C-Pr CrPSP $\ddot{\text{e}}$ PeP $^{\circ}$ C,P $\ddot{\text{e}}$ PsPrPSPsC $\ddagger$ P $^{\circ}$ Cl $\ddagger$ PSPsPiPs P $\cdot$ P $^{\circ}$ Cl $\ddagger$ C,PsCl $\ddagger$ CrPIP $^{\circ}$ PSPSC $\ddagger$
P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PSC $\ddagger$ P $\cdot$ C-PSPiC-P $\pm$ C-C,PsC $\ddot{\text{H}}$ P $^{\circ}$ PjP $\ddot{\text{e}}$ CYP3A4 (P $\cdot$
PeP μ C,PsPePsPSP $^{\circ}$ P $\cdot$ PsP $\gg$ PsPj, C-C,C $\ddot{\text{H}}$ P $^{\circ}$ PePsPSP $^{\circ}$ P $\cdot$ PsP $\gg$ PsPj, C $\ddot{\text{H}}$ P $\ddot{\text{e}}$ C,PsPSP $^{\circ}$ PIC-C $\ddot{\text{H}}$ PsPj,
P μ C $\ddot{\text{H}}$ P $\ddot{\text{e}}$ C,C $\ddot{\text{H}}$ PsPjC-C $\ddagger$ P $\ddot{\text{e}}$ PSPsPj, C,C $\ddot{\text{H}}$ PsP $\gg$ P μ P $^{\circ}$ PSPrPsPjC-C $\ddagger$ P $\ddot{\text{e}}$ PSPsPj,
PeP $\gg$ P $^{\circ}$ C $\ddot{\text{H}}$ P $\ddot{\text{e}}$ C,C $\ddot{\text{H}}$ PsPjC-C $\ddagger$ P $\ddot{\text{e}}$ PSPsPj).

P $\ddot{\text{I}}$ P $\ddot{\text{e}}$ PeP $\gg$ PsCl $\ddagger$ PiPsC $\ddot{\text{H}}$ P $\ddot{\text{e}}$ PS

P μ C $\ddot{\text{H}}$ P $\ddot{\text{e}}$ PsPrPSPsC $\ddagger$ P $^{\circ}$ Cl $\ddagger$ PSPsPjCr P $\cdot$ P $^{\circ}$ Cl $\ddagger$ C,PsCl $\ddagger$ CrPIP $^{\circ}$ PSPSC- P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PSC $\ddagger$
C- C $\ddagger$ P $\ddot{\text{e}}$ PeP $\gg$ PsCl $\ddagger$ PiPsC $\ddot{\text{H}}$ P $\ddot{\text{e}}$ PSC $\ddagger$ P $\ddot{\text{I}}$ C-PrPIP $\ddot{\text{e}}$ C $\%_{\text{Cr}}$ C $\ddot{\text{H}}$ C,CHC $\ddagger$ Cl $\ddagger$ C $\ddot{\text{H}}$ C-PIP μ PSCH PsP $\pm$ PsC...
C $\ddot{\text{H}}$ P μ C $\ddagger$ PsPIP $\ddot{\text{e}}$ PS Cr P $\ddot{\text{I}}$ P $\gg$ P $^{\circ}$ P $\cdot$ PjC- PeC $\ddot{\text{H}}$ PsPIC-. P $\ddot{\text{H}}$ PsCl $\ddagger$ P $\gg$ C-PrP $\ddagger$ P μ PSPSC $\ddagger$ P $\cdot$ P $^{\circ}$ CrC $\ddagger$ P $^{\circ}$ Cl $\ddagger$ C,CH
PjPsP $\gg$ PsPrP $\ddot{\text{e}}$ C... P $\cdot$ PrPsC $\ddot{\text{H}}$ PsPIP $\ddot{\text{e}}$ C... PrPsP $\pm$ C $\ddot{\text{H}}$ PsPIP $\ddot{\text{e}}$ P $\gg$ CHC $\ddagger$ C-PI P $\ddot{\text{I}}$ PsPeP $^{\circ}$ P $\cdot$ P $\gg$ Ps, C $\%_{\text{Ps}}$
P $\cdot$ P $^{\circ}$ Cl $\ddagger$ C,PsCl $\ddagger$ CrPIP $^{\circ}$ PSPSC $\ddagger$ C $\ddagger$ P $\ddot{\text{e}}$ PeP $\gg$ PsCl $\ddagger$ PiPsC $\ddot{\text{H}}$ P $\ddot{\text{e}}$ PSC $\ddagger$ C $\ddagger$ P μ C $\ddot{\text{H}}$ P μ P $\cdot$ 3 PiPsPrP $\ddot{\text{e}}$ PSP $\ddot{\text{e}}$
P $\ddot{\text{I}}$ C-Cl $\ddagger$ P $\gg$ C $\ddagger$ P $\ddot{\text{I}}$ C $\ddot{\text{H}}$ P $\ddot{\text{e}}$ PN $\ddot{\text{e}}$ PsPjCr P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PSC $\ddagger$ PSP μ P $\cdot$ PjC-PSCHPIP $^{\circ}$ P $\gg$ Ps
C $\ddot{\text{H}}$ C-PIP μ PSCH P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PSC $\ddagger$ Cr P $\ddot{\text{I}}$ P $\gg$ P $^{\circ}$ P $\cdot$ PjC- PeC $\ddot{\text{H}}$ PsPIC-, PIPsPrPSPsC $\ddagger$ P $^{\circ}$ Cl $\ddagger$
P $\ddot{\text{I}}$ PsPeP $^{\circ}$ P $\cdot$ PSP $\ddot{\text{e}}$ Pe AUC C $\ddagger$ P $\ddot{\text{e}}$ PeP $\gg$ PsCl $\ddagger$ PiPsC $\ddot{\text{H}}$ P $\ddot{\text{e}}$ PSC $\ddagger$ P $\cdot$ P $\pm$ C-P $\gg$ CHC $\ddot{\text{E}}$ CrPIP $^{\circ}$ PIC $\ddagger$ Cl $\ddagger$ PSP $^{\circ}$ 27 %.
PhPrPSP $^{\circ}$ Pe PsPrPSPsC $\ddagger$ P $^{\circ}$ Cl $\ddagger$ PSP μ P $\cdot$ P $^{\circ}$ Cl $\ddagger$ C,PsCl $\ddagger$ CrPIP $^{\circ}$ PSPSC $\ddagger$ P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PSC $\ddagger$
C- C $\ddagger$ P $\ddot{\text{e}}$ PeP $\gg$ PsCl $\ddagger$ PiPsC $\ddot{\text{H}}$ P $\ddot{\text{e}}$ PSC $\ddagger$ P $\ddot{\text{I}}$ C $\ddot{\text{H}}$ P $\ddot{\text{e}}$ P $\cdot$ PIPsPrP $\ddot{\text{e}}$ C,CH PrPs 3-PeC $\ddot{\text{H}}$ P $^{\circ}$ C,PSPsPiPs
P $\cdot$ P $\pm$ C-P $\gg$ CHC $\ddot{\text{E}}$ P μ PSPSC $\ddagger$ C $\ddot{\text{H}}$ C-PIPSC $\ddagger$ P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PSC $\ddagger$ Cr P $\ddot{\text{I}}$ P $\gg$ P $^{\circ}$ P $\cdot$ PjC-
PeC $\ddot{\text{H}}$ PsPIC- C,P $^{\circ}$ P $\cdot$ P $\pm$ C-P $\gg$ CHC $\ddot{\text{E}}$ P μ PSPSC $\ddagger$ P $\ddot{\text{I}}$ PsPeP $^{\circ}$ P $\cdot$ PSP $\ddot{\text{e}}$ PeP $^{\circ}$ AUC
C $\ddagger$ P $\ddot{\text{e}}$ PeP $\gg$ PsCl $\ddagger$ PiPsC $\ddot{\text{H}}$ P $\ddot{\text{e}}$ PSC $\ddagger$ PSP $^{\circ}$ 21 %.

P $\ddot{\text{I}}$ P $\ddot{\text{e}}$ PeP $\gg$ PsCl $\ddagger$ PiPsC $\ddot{\text{H}}$ P $\ddot{\text{e}}$ PS C,P $^{\circ}$ P $\gg$ P μ C $\ddot{\text{H}}$ PeP $^{\circ}$ PSC-PrP $\ddot{\text{e}}$ PiC-PS PSP μ Cl $\ddagger$ P $\gg$ C-Pr
P $\cdot$ P $^{\circ}$ Cl $\ddagger$ C,PsCl $\ddagger$ PsPIC $\ddagger$ PIP $^{\circ}$ C,P $\ddot{\text{e}}$ C $\ddot{\text{H}}$ P $^{\circ}$ P $\cdot$ PsPj.

P $\ddot{\text{H}}$ C $\ddot{\text{H}}$ P μ PN $\ddot{\text{e}}$ PiC $\ddot{\text{H}}$ CrC, P $^{\circ}$ P $\pm$ Ps PiC $\ddot{\text{H}}$ P μ PN $\ddot{\text{e}}$ PiC $\ddot{\text{H}}$ CrC,PsPIP $\ddot{\text{e}}$ PN $\ddot{\text{e}}$ Cl $\ddagger$ C-Pe

$P\ddot{I}P\epsilon$ C- $PrP \gg C\dot{I} C-PSC\epsilon P\ddot{e}C \dots PrP\ddot{e}P\dot{I}C-PrC\dot{H}PsP\dot{I}C-C\dot{H}P\ddot{e}PrP\ddot{e}PSC-PI$, $PjP\mu C, P^{\circ}P \pm PsP \gg C-P \cdot Pj$
 $P \gg P\mu C\dot{H}PeP^{\circ}PSC-PrP\ddot{e}P\dot{I}C-PSC\acute{r}$ $C\acute{r}P\dot{I}PsPIC-P \gg CH\dot{P}SC\dot{H}C''C, CH\dot{C}\dot{I}C\dot{I} P\dot{I}C-Pr PIP\dot{I}P \gg P\ddot{e}PIPsPj$
 $C\dot{I}PsPeC\acute{r} P\dot{I}C\dot{H}P\mu PN\epsilon P\dot{I}C, C\dot{H}C\acute{r}C, P^{\circ} P \cdot P\dot{I}PsPrP^{\circ}P \gg CH\dot{C}\epsilon P\ddot{e}Pj P\dot{I}C-PrPIP\ddot{e}C\%P\mu PSPSC\dot{I}Pj$
 $C\dot{I}P\ddot{e}C\dot{I}C, P\mu PjPSPsC- PrPsC\dot{I}C, C\acute{r}P\dot{I}PSPsC\dot{I}C, C- P \gg P\mu C\dot{H}PeP^{\circ}PSC-PrP\ddot{e}P\dot{I}C-PSC\acute{r} C, P^{\circ}$
 $P\dot{I}C-PrC\dot{I}P\ddot{e}P \gg P\mu PSPSC\dot{I}Pj P\dot{I}C-P\dot{I}PsC, P\mu PSP \cdot P\ddot{e}PIPsPsC- PrC-C- . P\acute{K}P\mu C\dot{I}P \gg C-Pr$
 $P\dot{I}C\dot{H}P\ddot{e}PN\epsilon PjP^{\circ}C, P\ddot{e} PsPrPSPsC\ddagger P^{\circ}C\dot{I}PSPs P \gg P\mu C\dot{H}PeP^{\circ}PSC-PrP\ddot{e}P\dot{I}C-PS C, P^{\circ}$
 $P\dot{I}C\dot{H}P\mu PN\epsilon P\dot{I}C, C\dot{H}C\acute{r}C, P^{\circ}P \pm Ps P\dot{I}C\dot{H}P\mu PN\epsilon P\dot{I}C, C\dot{H}C\acute{r}C, PsPIP\ddot{e}PN\epsilon C\dot{I}C-Pe.$

$P\acute{K}P\mu C\dot{H}P\mu PePsPjP\mu PSPPrPsPIP^{\circ}PSPs PsPrPSPsC\ddagger P^{\circ}C\dot{I}PSP\mu P \cdot P^{\circ}C\dot{I}C, PsC\dot{I}C\acute{z}PIP^{\circ}PSPSC\dot{I}I$

$P\ddagger PSPrC\acute{z}PeC, PsC\dot{H}P\ddot{e} CYP3A4$

$P\acute{K}P\mu PsP \pm C \dots C-PrPSPs P \cdot PsP \pm P\mu C\dot{H}P\mu P\dot{I}PSC-C\dot{I}C, C\dot{H} P \cdot P^{\circ}C\dot{I}C, PsC\dot{I}PsPIC\acute{r}PIP^{\circ}C, P\ddot{e}$
 $P \gg P\mu C\dot{H}PeP^{\circ}PSC-PrP\ddot{e}P\dot{I}C-PS PsPrPSPsC\ddagger P^{\circ}C\dot{I}PSPs P \cdot C-PSPrC\acute{r}PeC, PsC\dot{H}P^{\circ}PjP\ddot{e} CYP3A4,$
 $C, P^{\circ}PeP\ddot{e}PjP\ddot{e} C\dot{I}Pe P\dot{I}C\dot{H}PsC, P\ddot{e}C\dot{I}C\acute{r}PrPsPjPSC- P \cdot P^{\circ}C\dot{I}PsP \pm P\ddot{e} (C, P\mu PSC-C, PsC-PS,$
 $C, P\mu PSPsP \pm P^{\circ}C\dot{H}P \pm C-C, P^{\circ}P \gg, PeP^{\circ}C\dot{H}P \pm P^{\circ}PjP^{\circ}P \cdot P\mu P\dot{I}C-PS) C, P^{\circ} C\dot{H}P\ddot{e}C, P^{\circ}PjP\dot{I}C-C\ddagger P\ddot{e}PSPsPj$
 $C\ddagger P\mu C\dot{H}P\mu P \cdot PjPsP\dot{I}P \gg P\ddot{e}PIP\mu P \cdot PSP\ddot{e}P\dot{I}P\mu PSPSC\dot{I} P^{\circ}PSC, P\ddot{e}P\dot{I}C-P\dot{I}P\mu C\dot{H}C, P\mu PSP \cdot P\ddot{e}PIPsPsC-$
 $PrC-C- P \gg P\mu C\dot{H}PeP^{\circ}PSC-PrP\ddot{e}P\dot{I}C-PSC\acute{r}. Pj C\ddagger P\ddot{e}C \dots PIP\ddot{e}P\dot{I}P^{\circ}PrPeP^{\circ}C \dots$
 $C\dot{H}P\mu PePsPjP\mu PSPPrPsPIP^{\circ}PSPs C\ddagger P^{\circ}C\dot{I}C, C-C\epsilon P\ddot{e}PN\epsilon PePsPSC, C\dot{H}PsP \gg CH C\dot{H}C-PIPSC\dot{I}I$
 $P^{\circ}C\dot{H}C, P\mu C\dot{H}C-P^{\circ}P \gg CH\dot{P}SPsPiPs C, P\ddot{e}C\dot{I}PeC\acute{r}.$

$P\dot{H}P \gg PePsPiPsP \gg CH$

$P\ddot{Y}P \gg C-Pr C\acute{r}PSP\ddot{e}PeP^{\circ}C, P\ddot{e} PIP\dot{I}P\ddot{e}PIP^{\circ}PSPSC\dot{I}I P^{\circ}P \gg PePsPiPsP \gg CH C\ddagger P\mu C\dot{H}P\mu P \cdot PjPsP\dot{I}P \gg P\ddot{e}PIP\mu$
 $P\dot{I}PsC, P\mu PSC\ddagger C-CHPIP^{\circ}PSPSC\dot{I}I PIP^{\circ}P \cdot PsPrP\ddot{e}P \gg P^{\circ}C, P^{\circ}C\ddagger C-PN\epsilon PSPsPiPs P\mu C, P\mu PeC, C\acute{r}$
 $P^{\circ}PSC, P\ddot{e}P\dot{I}C-P\dot{I}P\mu C\dot{H}C, P\mu PSP \cdot P\ddot{e}PIPsP\ddot{e}C \dots P \gg C-PeP^{\circ}C\dot{H}C\dot{I}CH\dot{H}PeP\ddot{e}C \dots P \cdot P^{\circ}C\dot{I}PsP \pm C-PI.$

$P'P \cdot P^{\circ}C''PjPsPrC-C-, C\dot{I}PeC- P\dot{I}PsC, C\dot{H}P\mu P \pm C\acute{z}CHC, CH PePsC\dot{H}P\mu PeC\ddagger C-C- PrPsP \cdot P\ddot{e}$

$P\ddot{Y}C\acute{z}P \pm C\dot{I}C, C\dot{H}P^{\circ}C, P\ddot{e} CYP3A4$

$P\ddot{Y}P \gg C-Pr P \pm C\acute{r}C, P\ddot{e} PsP \pm P\mu C\dot{H}P\mu P\dot{I}PSP\ddot{e}Pj P\dot{I}C\dot{H}P\ddot{e} PsPrPSPsC\ddagger P^{\circ}C\dot{I}PSPsPjC\acute{r}$
 $P \cdot P^{\circ}C\dot{I}C, PsC\dot{I}C\acute{r}PIP^{\circ}PSPSC- P \gg P\mu C\dot{H}PeP^{\circ}PSC-PrP\ddot{e}P\dot{I}C-PSC\acute{r} P \cdot C-PSC\epsilon P\ddot{e}PjP\ddot{e}$
 $C\dot{I}C\acute{r}P \pm C\dot{I}C, C\dot{H}P^{\circ}C, P^{\circ}PjP\ddot{e} CYP3A4, C, P^{\circ}PeP\ddot{e}PjP\ddot{e} C\dot{I}Pe C, P\mu C\dot{H}C, P\mu PSP^{\circ}PrP\ddot{e}PS,$
 $P^{\circ}C\dot{I}C, P\mu PjC-P \cdot PsP \gg, P^{\circ}PSC, P\ddot{e}P^{\circ}C\dot{H}P\ddot{e}C, PjC-C\ddagger PSP\ddot{e}PjP\ddot{e} P\dot{I}C\dot{H}P\mu P\dot{I}P^{\circ}C\dot{H}P^{\circ}C, P^{\circ}PjP\ddot{e} P\ddagger P\ddagger P\ddagger$
 $PeP \gg P^{\circ}C\dot{I}C\acute{r}, C, P^{\circ}PeP\ddot{e}PjP\ddot{e} C\dot{I}Pe P^{\circ}PjC-PsPrP^{\circ}C\dot{H}PsPS, PePIC-PSC-PrP\ddot{e}PS, C\dot{I}PsC, P^{\circ}P \gg PsP \gg.$

$P\dot{H}C-PrP^{\circ}P \cdot PsP \gg P^{\circ}Pj$

$P\mu C\dot{H}P\ddot{e} PsPrPSPsC\ddagger P^{\circ}C\dot{I}PSPsPjC\acute{r} P \cdot P^{\circ}C\dot{I}C, PsC\dot{I}C\acute{r}PIP^{\circ}PSPSC- 20 PjPi$
 $P \gg P\mu C\dot{H}PeP^{\circ}PSC-PrP\ddot{e}P\dot{I}C-PSC\acute{r} C, P^{\circ} PjC-PrP^{\circ}P \cdot PsP \gg P^{\circ}PjC\acute{r} C\acute{r} PrPsP \pm C\dot{H}PsPIPsP \gg CH\dot{C}\ddagger C-PI$
 $P \gg C-C, PSCH\dot{H}PsPiPs PIC-PeC\acute{r} P^{\circ}P \pm C\dot{I}PsC\dot{H}P \pm C\ddagger C-C\dot{I}I P \gg P\mu C\dot{H}PeP^{\circ}PSC-PrP\ddot{e}P\dot{I}C-PSC\acute{r}$
 $P\dot{I}C-PrPIP\ddot{e}C\%C\acute{r}PIP^{\circ}P \gg P^{\circ}C\dot{I}CH\dot{H} (P\dot{I}C\dot{H}P\ddot{e}P \pm P \gg P\ddot{e}P \cdot PSPs PSP^{\circ} 40 \%), P^{\circ} C\epsilon PIP\ddot{e}PrPeC-C\dot{I}C, CH$
 $P^{\circ}P \pm C\dot{I}PsC\dot{H}P \pm C\ddagger C-C- P \cdot PSP\ddot{e}P\dot{I}C\acute{r}PIP^{\circ}P \gg P^{\circ}C\dot{I}CH\dot{H} (t_{\max} P\dot{I}PsPrPsPIP\dot{I}C\acute{r}PIP^{\circ}P \gg P^{\circ}C\dot{I}CH\dot{H} P \cdot 1,75$
 $PrPs 3 P\dot{I}PsPrP\ddot{e}PS). P\dot{L}PsPSC\ddagger P\mu PSC, C\dot{H}P^{\circ}C\ddagger C-C\dot{I}I PjC-PrP^{\circ}P \cdot PsP \gg P^{\circ}PjC\acute{r} PSP\mu$
 $P \cdot PjC-PSCH\dot{H}PIP^{\circ}P \gg P^{\circ}C\dot{I}CH\dot{H}.$

$P\dot{H}P\mu C, PsP\dot{I}C\dot{H}PsP \gg PsP \gg$

$P\dot{H}PrPSPsC\ddagger P^{\circ}C\dot{I}PSP\mu P \cdot P^{\circ}C\dot{I}C, PsC\dot{I}C\acute{r}PIP^{\circ}PSPSC\dot{I}I P \gg P\mu C\dot{H}PeP^{\circ}PSC-PrP\ddot{e}P\dot{I}C-PSC\acute{r} P \cdot$
 $PjP\mu C, PsP\dot{I}C\dot{H}PsP \gg PsP \gg PsPj \text{ в } \dot{H} \beta-P \pm P \gg PsPeP^{\circ}C, PsC\dot{H}PsPj, C\dot{I}PeP\ddot{e}PN\epsilon PIP\ddot{e}PIPsPrP\ddot{e}C, CH\dot{C}\dot{I}C\dot{I}$
 $P\dot{I}P\mu C\dot{H}P\mu PIP^{\circ}P\dot{I}PSPs C\ddagger P\mu C\dot{H}P\mu P \cdot P\dot{I}P\mu C\ddagger C-PSPeC\acute{r}, \text{ в } \dot{H} \text{ " } PSP\mu P \cdot PjC-PSCH\dot{H}C''$
 $P \pm C-PsPrPsC\dot{I}C, C\acute{r}P\dot{I}PSC-C\dot{I}C, CH PjP\mu C, PsP\dot{I}C\dot{H}PsP \gg PsP \gg C\acute{r}, P^{\circ}P \gg P\mu P\dot{I}C\dot{H}P\ddot{e}P \cdot PIPsPrP\ddot{e}C, CH$

С†РµPN_ε P¶Pµ PīPsPeP°P·PSPëPe PrP»CΠ PN_εPsPiPs P°PeC,PëPIPSsPiPs PjPµC,P°P±PsP»C-C,С† β-
PiC-PrCḡPsPeCḡPëPePëCḡP»PsC,Pë Bḡ“ PSP° 28 %. PḡP°P»PsPN_εPjPsPIC-CḡPSPs, C%_oPs
C,P°PeC- P·PjC-PSPë PjP°CḡC,CH PeP»C-PSC-C†PSPµ P·PSP°C†PµPSPSCΠ. PќPµ
PsC†C-PeCḡC“C,CHCḡCΠ PIP·P°C”PjPsPrC-C— PjC-P¶ C†PëPjPë PīCḡPµPīP°CḡP°C,P°PjPë,
CΠPeC%_oPs P»PµCḡPeP°PSC-PrPëPiC-PS PīCḡPëPN_εPjP°C,Pë P·CḡP°PSPeCḡ, P°
CḡPëPjPIP°CḡC,P°C,PëPS Bḡ“ PIPµC†PµCḡC-, CΠPe PIPeP°P·P°PSPs PrP»CΠ C†CHPsPiPs
PīCḡPµPīP°CḡP°C,Cḡ.

P”C-CḡCḡPµC,PëPePë, C-PSPiC-P±C-C,PsCḡPë
P°PSPiC-PsC,PµPSP·PëPSPiPµCḡPµC,PIPsCḡCHPIP°P»CHPSPsPiPs C,,PµCḡPjPµPSC,Cḡ (PḡPµPḡ)

P>PµCḡPeP°PSC-PrPëPiC-PS PjPsP¶PSP° P·P°CḡC,PsCḡPsPICḡPIP°C,Pë PsPrPSPsC†P°CḡPSPs P·
PrC-CḡCḡPµC,PëPeP°PjPë C,P° C-PSPiC-P±C-C,PsCḡP°PjPë PḡPµPḡ.

P†PSCεC- P»C-PeP°CḡCḡCHPeC- P·P°CḡPsP±Pë, CΠPeC- PIPiP»PëPIP°CḡC,CH PSP°
P°CḡC,PµCḡC-P°P»CHPSPëPN_ε C,PëCḡPe

PīPe C- PrP»CΠ PICḡC-C... P°PSC,PëPiC-PiPµCḡC,PµPSP·PëPIPSsPiPs PīCḡPµPīP°CḡP°C,C-PI,
PjPsP¶P»PëPIPµ PīPsCḡPëP»PµPSPSCΠ PiC-PiPsC,PµPSP·PëPIPSsPiPs PµC,,PµPeC,Cḡ PīCḡPë
PsPrPSPsC†P°CḡPSPsPjCḡ P·P°CḡC,PsCḡCḡPIP°PSPSC- P»PµCḡPeP°PSC-PrPëPiC-PSCḡ P·
C-PSCεPëPjPë P»C-PeP°CḡCḡCHPePëPjPë P·P°CḡPsP±P°PjPë, CΠPeC- PIPiP»PëPIP°CḡC,CH
PSP° P°CḡC,PµCḡC-P°P»CHPSPëPN_ε C,PëCḡPe, C,P°PePëC... CΠPe α-P±P»PsPeP°C,PsCḡPë
PrP»CΠ CḡPëPjPiC,PsPjP°C,PëC†PSPsPiPs P»C-PeCḡPIP°PSPSCΠ P·P°C...PIPsCḡCHPIP°PSCḡ
CḡPµC†PsPIPsPiPs PjC-C...CḡCḡP°, C,CḡPëC†PëPeP»C-C†PSC- P°PSC,PëPrPµPīCḡPµCḡP°PSC,Pë,
PSPµPN_εCḡPsP»PµPiC,PëPePë.

PќP°PIPiP°PePë, PjPsP¶Pµ CḡPiPsCḡC,PµCḡC-PiP°C,PëCḡCΠ P·PSPëP¶PµPSPSCΠ
PiC-PiPsC,PµPSP·PëPIPSsPiPs PµC,,PµPeC,Cḡ PīCḡPë PsPrPSPsC†P°CḡPSPsPjCḡ
P·P°CḡC,PsCḡCḡPIP°PSPSC- P· PePsCḡC,PëPePsCḡC,PµCḡPsC—PrP°PjPë.

PḡCḡPsP±P»PëPIPsCḡC,C- P·P°CḡC,PsCḡCḡPIP°PSPSCΠ

PḡPëPSPPrCḡPsPj CḡP»P°P±PePsCḡC,C- CḡPëPSCḡCḡPsPIPsPiPs PICḡP·P»P°

P>PµCḡPeP°PSC-PrPëPiC-PS CḡP»C-Pr P· PsP±PµCḡPµP¶PSC-CḡC,CH
P·P°CḡC,PsCḡPsPICḡPIP°C,Pë PīP°C†C-C”PSC,P°Pj C-P· CḡPëPSPPrCḡPsPjPsPj
CḡP»P°P±PePsCḡC,C- CḡPëPSCḡCḡPsPIPsPiPs PICḡP·P»P° (P±PµP· C-PjPīP»P°PSC,PsPIP°PSPsPiPs
PeP°CḡPrC-PsCḡC,PëPjCḡP»CΠC,PsCḡP°).

P”PëCḡC,,CḡPSPeC†C-CΠ P»C-PIPsPiPs CεP»CḡPSPsC†PeP° CḡPµCḡC†CΠ

PḡPsC†P° PiPµPjPsPrPëPSP°PjC-C†PSPs PePsPSC,CḡPsP»CHPsPIP°PSC-
PrPsCḡP»C-PrP¶PµPSPSCΠ PSPµ PIPëCΠPIPëP»Pë PīPsPiC-CḡCεPµPSPSCΠ
CεP»CḡPSPsC†PePsPIPsC— C,,CḡPSPeC†C-C—, P· PsP±PµCḡPµP¶PSC-CḡC,CH CḡP»C-Pr
PīCḡPëP·PSP°C†P°C,Pë PīCḡPµPīP°CḡP°C, PīCḡPë PrPëCḡC,,CḡPSPeC†C-C— P»C-PIPsPiPs
CεP»CḡPSPsC†PeP° CḡPµCḡC†CΠ.

P†CεPµPjC-C†PSP° C...PIPsCḡPsP±P° CḡPµCḡC†CΠ

P’CḡP»Ps PIPëCḡP»PsPIP»PµPSPs PīCḡPëPiCḡC%_oPµPSPSCΠ, C%_oPs PrPµCΠPeC-
PrPëPiC-PrCḡPsPiC-CḡPëPrPëPSPë PePsCḡPsC,PePsC— PrC-C— PjPsP¶CḡC,CH P±CḡC,Pë

PĭPsPI'CIIP·P°PSC- P· PĭC-PrPIPĕC%oPμPSPĕPj CĭPμCĭC†PμPIPs-CĭC†PrPĕPSPSPĕPj
 CĭPĕP·PĕPePsPj Cĭ PĭP°C†C-C"°PSC,C-PI P· C-CĕPμPjC-C†PSPsCĭ C...PIPsCĭPsP±PsCĭ
 CĭPμCĭC†Cĭ. PĭPsC†P° P·P—PĭPĕPĭPĕP C" PĭCĭPμPĭP°CĭP°C,PSPj
 PrPsPIPIPsC,CĭPĕPIP°P»PSC— PrC-C—, PĭCĭPsC,Pμ C,P°PePĕPj C...PIPsCĭPĕPj
 PĭCĭPμPĭP°CĭP°C, CĭP»C-Pr P·P°CĭC,PSCĭPsPICĭPIP°C,Pĕ P· PsP±PμCĭPμP¶PSC-CĭC,Cĭ.
 P"°PμCĭPeC- PrPĕPiC-PrCĭPsPĭC-CĭPĕPrPĕPSPĕ CĭC-PrPePs PjPsP¶CĭC,Cĭ
 PĭCĭPĕP·PIPsPrPĕC,Pĕ PrPs PĭCĭPμPePsCĭPrC-P°P»CHPSPsPiPs P±PsP»Cĭ P°P±Ps
 CĭC,PμPSPsPeP°CĭPrC-C—. P"°CĭP¶Pμ CĭC-PrPePs Cĭ PĭP°C†C-C"°PSC,C-PI P· CĭP°PSC-CĕPμ
 C-CĭPSCĭCĭC†PsCĭ CĭC,PμPSPsPeP°CĭPrC-C"°Cĭ PjPsP¶P»PĕPIPμ P·P±C-P»CHCĕPμPSPSCĭ
 C†P°CĭC,PSC,Pĕ, C,CĭPĕPIP°P»PSCĭC,C- P°P±Ps C,CĭP¶PePsCĭC,C- C†PĕC... PSP°PĭP°PrC-PI.
 PĕPsP¶CĭC,Cĭ PIC-PrP·PSP°C†P°C,PĕCĭCĭ PĭPsPsPrPĕPSPsPeC- PIPĕPĭP°PrPePĕ
 C-PSC,,P°CĭPeC,Cĭ PjC-PsPeP°CĭPrP°.

PμPμCĭPĕC,PSPSPμP°P»CHPSPĕPN Prc-P°P»C-P.

P—P°CĭC,PSCĭCĭPIP°PSPSCĭ P»PμCĭPeP°PSC-PrPĕPiC-PSCĭ PĭPsPIĕĭ™CĭP·CĭC"°C,CĭCĭCĭ P·
 CĭPsP·PIPĕC,PePsPj PĭPsPjCĭC,PSC-PSPSCĭ PĭPμCĭPĕC,PSPSPμP°P»CHPSPsPiPs
 PμPeCĭCĭPrP°C,Cĭ Cĭ PĭP°C†C-C"°PSC,C-PI PSP° PĭPμCĭPĕC,PSPSPμP°P»CHPSPsPjCĭ
 PrC-P°P»C-P·C-. PμPsPjCĭC,PSC-PSPSCĭ PsP±CĭPjPsPIP»PμPSPs PĭC-PrPIPĕC%oPμPSPsCĭ
 PePsPSC†PμPSC,CĭP°C†C-C"°Cĭ C,CĭPĕPiP»C-C†PμCĭPĕPrC-PI PI PμPeCĭCĭPrP°C,C-
 PsC†PμCĭPμPIPĕPSPĕ. PĭPsC†P° PjPμC...P°PSC-P·Pj PSPμPIC-PrPsPjPĕPN, C†PμPN PμC,,PμPeC,
 PjP°C" C,PμPSPPrPμPSC†C-Cĭ P·PSPĕPeP°C,Pĕ PSPμPIPrPsPIP·C- PĭC-CĭP»Cĭ PIC-PrPjC-PSPĕ
 P»PμCĭPeP°PSC-PrPĕPiC-PSCĭ. P|PμPN PIP·P°C"PjPsP·PIĕĭ™CĭP·PsPe CĭP»C-Pr PIP·CĭC,Pĕ
 PrPs CĭPIP°PiPĕ P·P°PrP»Cĭ CĭPSPĕPePSPμPSPSCĭ PIPĕPĭP°PrPeC-PI, PePsP»Pĕ
 PĭPsPjCĭC,PSC-PSPSCĭ PĭPμCĭPĕC,PSPSPμP°P»CHPSPsPiPs PμPeCĭCĭPrP°C,Cĭ PjPsP¶Pμ
 P±CĭC,Pĕ PĭPsPjPĕP»PePsPIPs PĭCĭPĕPN PSCĭC,Ps P·P° C-PSC,,PμPeC†C-PN PSPĕPN
 PĭPμCĭPĕC,PSPSC-C, P· PĭPsPrP°P»CHCĕPsCĭ PSPμPĭPsC,CĭC-P±PSPsCĭ
 PĭPsCĭPĭC-C,P°P»C-P·P°C†C-C"°Cĭ C,P° PμPjPĭC-CĭPĕC†PSPĕPj PIPμPrPμPSPSCĭPj
 P°PSC,PĕP±C-PsC,PĕPeC-PI.

P»P°PeC,PSP·P°

P|PμPN P»C-PeP°CĭCĭCHPePĕPN P·P°CĭC-P± PjC-CĭC,PĕC,Cĭ P»P°PeC,PSP·Cĭ.

PμP°C†C-C"°PSC,P°Pj C-P· CĭC-PrPeC-CĭPSPĕPjPĕ CĭPĭP°PrPePsPIPĕPjPĕ PĭCĭPsP±P»PμPjP°PjPĕ
 PSPμPĭPμCĭPμPSPsCĭPĕPjPsCĭC,C- PĭP°P»P°PeC,PSP·Pĕ, P·P°PĭP°P»CHPSPsC—
 P»P°PeC,P°P·PSPsC— PSPμPrPsCĭC,P°C,PSPsCĭC,C- P°P±Ps PĭP»CĭPePsP·Ps-
 PĭP°P»P°PeC,PSP·PSPsC— PjP°P»CHP°P±CĭPsCĭP±C†C-C— PSPμ CĭP»C-Pr
 P·P°CĭC,PSCĭPsPICĭPIP°C,Pĕ C†PμPN P»C-PeP°CĭCĭCHPePĕPN P·P°CĭC-P±.

***P—P°CĭC,PSCĭCĭPIP°PSPSCĭ Cĭ PĭPμCĭC-PsPr PIP°PiC-C,PSPsCĭC,C- P°P±Ps
 PĭPsPrCĭPIP°PSPSCĭ PĭCĭCĭPrPrCĭ***

P'°PiC-C,PSC-CĭC,Cĭ

PĕP»C-PSC-C†PSPĕPN PrPsCĭPIC-Pr P·P°CĭC,PSCĭCĭPIP°PSPSCĭ
 P»PμCĭPeP°PSC-PrPĕPiC-PSCĭ Cĭ PĭPμCĭC-PsPr PIP°PiC-C,PSPsCĭC,C- PIC-PrCĭCĭC,PSC-PN.
 P"°PSCĭP»C-PrPĕ PSP° C,PIP°CĭPĕPSP°C... PSPμ PIPĕCĭPIPĕP»Pĕ C,PμCĭP°C,PSPμPSPsPiPs
 PμC,,PμPeC,Cĭ, P°P»Pμ PIC-PS CĭPĭPsCĭC,PμCĭC-PĭP°PICĭCĭ PĭC-Pr C†P°Cĭ
 P·P°CĭC,PSCĭCĭPIP°PSPSCĭ C-PSCĕPĕC... PrPĕPiC-PrCĭPsPĭC-CĭPĕPrPĕPSPsPIPĕC...

СГРiPcP»CpPе. PкPм СГРiPмPеPcPjPмPSPpPcPIP°PSPc P·P°CГC,PsCГPcPICpPIP°C,Pë
P»PмCГPеP°PSC-PrPëPc-PS CГ PиPмCГC-PcPr PIP°Pc-C,PSPcCГC,C- P°P±Ps P¶C-PSPeP°Pj
CГPмPиCГPcPrCpPеC,PëPIPSPcPiPs PIC-PeCГ, C¶PеC%Ps PIPcPSPë PSPм
P·P°CГC,PsCГPcPICpCГC,CH PмC,,PмPеC,PëIPSCГ PеPcPSC,CГP°C†PмPиC†C-CГ.

PмPмCГC-PcPr PiPsPrCзPIP°PSPSC¶ PiCГCзPrPrCГ

PÿPcC±PSPc PSPмPIC-PrPcPjPs, C±Pë PиCГPcPSPëPеP°C” P»PмCГPеP°PSC-PrPëPc-PS P°P±Ps
PмPcPiPs PjPмC,P°P±PsP»C-C,Pë CГ PiCГCГPrPSPм PjPsP»PsPеPs, C,PsPjCГ CГPëP·PëPе PrP»C¶
PrPëC,PëPSPë PSPм PjPsP¶Pм P±CГC,Pë PIPëPеP»CГC±PмPSPëPм. PкPм CГP»C-Pr
P·P°CГC,PsCГPcPICpPIP°C,Pë P»PмCГPеP°PSC-PrPëPc-PS PиC-Pr C±P°CГ PiPsPrCГPIP°PSPSC¶
PiCГCГPrPrCГ.

PмPмCГC,PëP»CHPSC-CГC,CH

PмP»C-PSC-C±PSC- PrP°PSC- C%PsPrPs PIPиP»PëPICp P»PмCГPеP°PSC-PrPëPc-PSCp PSP°
C,,PмCГC,PëP»CHPSC-CГC,CH PIC-PrCГCГC,PSC-. PкP°C¶IPSC- PrP°PSC- PиCГPc
PsP±PsCГPcPSC,PSC- P±C-PcC...C-PjC-C±PSC- P·PjC-PSPë PI PiPsP»C-PIC†C-
CГPиPмCГPjP°C,PsP·Pc—PrC-PI, C%Ps PjPsP¶CГC,CH PIPиP»PëPSCpC,Pë PSP°
PjPsP¶P»PëPIC-CГC,CH P·P°PиP»C-PrPSPмPSPSC¶, CГ PиP°C†C-C”PSC,C-PI, C¶PеC-
P»C-PeCГPIP°P»PëCГC¶ P±P»PsPеP°C,PsCГP°PjPë PеP°P»CHC†C-C”PIPëC... PеP°PSP°P»C-PI. Pj
CГP°P·C- PиPsPIC,PsCГPSPcPiPs PSPмCГCГPиC-C€PSPcPiPs P·P°PиP»C-PrPSPмPSPSC¶ *in vitro*
C,P° P·P° PIC-PrCГCГC,PSPcCГC,C- C-PSC€PëC... PиPsC¶CГPSPмPSCH C%PsPrPs C†CHPcPiPs
CГP»C-Pr CГPcP·PiP»C¶PSCpC,Pë P·P°CГC,PsCГCГPIP°PSPSC¶ P±P»PsPеP°C,PsCГC-PI
PеP°P»CHC†C-C”PIPëC... PеP°PSP°P»C-PI C¶Pе PjPsP¶P»PëPICp PиCГPëC±PëPSCp.

**P—PrP°C,PSC-CГC,CH PIPиP»PëPIP°C,Pë PSP° C€PIPëPrPеC-CГC,CH CГPмP°PеC†C-C—
PиCГPë PеPмCГCзPIP°PSPSC- P°PIC,PsC,CГP°PSCГPиPsCГC,PsPj P°P±Ps C-PSC€PëPjPë
PjPмC...P°PSC-P·PjP°PjPë**

P’PиP»PëPI P»PмCГPеP°PSC-PrPëPc-PSCp PSP° P·PrP°C,PSC-CГC,CH PеPмCГCГPIP°C,Pë
C,CГP°PSCГPиPsCГC,PSPëPj P·P°CГPcP±PsPj P°P±Ps C-PSC€PëPjPë PjPмC...P°PSC-P·PjP°PjPë
PSPмP·PSP°C±PSPëPм. PhPrPSP°Pе PSPмPsP±C...C-PrPSPc PICГP°C...PsPICpPIP°C,Pë
PjPsP¶P»PëPIC-CГC,CH P·P°PиP°PjPsCГPcC±PмPSPSC¶, CГP»P°P±PеPsCГC,C-,
PиC-PrPIPëC%PмPSPc— PIC,PsPjP»CГPIP°PSPcCГC,C-, CГC-PrPеPs бГ”
CГPcPSP»PëIPcCГC,C-.

PÿPиPsCГC-P± P·P°CГC,PsCГCзPIP°PSPSC¶ C,P° PrPsP·Pë

P PмPеPcPjPмPSPpPcPIP°PSP° PrPsP·P° CГC,P°PSPcPIPëC,CH 10 PjPi PиPмCГPcCГP°P»CHPSPc 1
CГP°P· PSP° PrPsP±CГ, P·P°CГC,PsCГPcPICpCГC,CH PSPм CГP°PSC-C€Pм PSC-P¶ P·P° 15
C...PIPëP»PëPS PrPs PIP¶PëPIP°PSPSC¶ C—P¶C-. P—P°P»PмP¶PSPc PIC-Pr
C-PSPrPëPIC-PrCГP°P»CHPSPc— CГPмP°PеC†C-C— PиP°C†C-C”PSC,P° PSP°
P»C-PeCГPIP°PSPSC¶ PrPsP·CГ PjPsP¶PSP° PиC-PrPIPëC%PëC,Pë PrPs 20 PjPi.

PмC-PrP±C-CГ PrPsP·Pë PjP°C” P±CГC,Pë PиPsCГC,CГPиPsPIPëPj, PsCГPеC-P»CHPеPë
PjP°PеCГPëPjP°P»CHPSP° P°PSC,PëPiC-PиPмCГC,PмPSP·PëIPSP° PrC-C¶
CГPcP·PIPëPIP°C”C,CHCГC¶ PиCГPcC,C¶PiPsPj PrPIPc... C,PëP¶PSC-PI P»C-PeCГPIP°PSPSC¶.

- $P \cdot P^{\circ}Cl^{\circ}C$, $PsCl^{\circ}PsPICr^{\circ}PIP^{\circ}C$, $P\ddot{e} \ P\ddot{i}CH_P\mu P\ddot{i}P^{\circ}CH_P^{\circ}C$, $P\pm P^{\circ}P^{\circ}P^{\circ}PSPs \ PICr^{\circ}P^{\circ}PSC^{\circ}C-$, $PSP\mu$

PjPμPSC€Pμ PSC-P¶ P·P° 15 C...PIPëP»PëPS PrPs CÍPSC-PrP°PSPeCf;

- C†PμPNë P»C-PeP°CĤCÍCHPePëPNë P·P°CÍC-P± PSPμ PjPsP¶PSP° PíCĤPëPNëPjP°C,Pë P·
PiCĤPμPNëPíC,,CĤCfC,PsPj C,P° PiCĤPμPNëPíC,,CĤCfC,PsPIPëPj CÍPsPePsPj.

P''C-C,Pë

P'PμP·PíPμPeP° C,P° PμC,,PμPeC,PëPIPSC-CÍC,CH P·P°CÍC,PsCÍCfPIP°PSPSCİ
P»C-PeP°CĤCÍCHPePsPiPs P·P°CÍPsP±Cf PrC-C,CİPj PIC-PePsPj PrPs 18 CĤPsPeC-PI PSPμ
PrPsCÍP»C-PrP¶CfPIP°P»PëCÍCİ, PrP°PSC- C%oPsPrPs P·P°CÍC,PsCÍCfPIP°PSPSCİ
PrC-C,CİPj PIC-PrCÍCfC,PSC-.

PμPμCĤPμPrPsP·C¿PIP°PSPSCİ

PμCĤPsC,CİPiPsPj PíPμCĤC-PsPrCf PíC-CÍP»CİCĤPμC''CÍC,CĤP°C†C-PNëPSPsPiPs
P·P°CÍC,PsCÍCfPIP°PSPSCİ P»PμCĤPeP°PSC-PrPëPíC-PSCf PíPsPIC-PrPsPjP»CİP»PsCÍCİ
PíCĤPs PrPμCİPeC- PIPëPíP°PrPePë PíPμCĤPμPrPsP·CfPIP°PSPSCİ (PIC-Pr 30BĤ"40 PjPi PrPs
800 PjPi, Cf C,PsPjCf C†PëCÍP»C- CÍPíCĤPsP±P° CÍCfC—C†PëPrCf).

PŸPëPjPíC,PsPjPë

P—P° P°PSP°P»PsPiC-C''CĤ P· C-PSC€PëPjPë PrPëPiC-PrCĤPsPíC-CĤPëPrPëPSP°PjPë, PíCĤPë
PíPμCĤPμPrPsP·CfPIP°PSPSC- P»PμCĤPeP°PSC-PrPëPíC-PSPsPj CÍP»C-Pr PsC†C-PeCfPIP°C,Pë
PIPëPSPëPePSPμPSPSCİ PSP°PrPjC-CĤPSPsC— PíPμCĤPëC,,PμCĤPëC†PSPsC—
PIP°P·PsPrPëP»P°C,P°C†C-C— C- CİPIPSPsC— P°CĤC,PμCĤC-P°P»CHPSPsC—
PiC-PíPsC,PμPSP·C-C— C,P° CĤPμC,,P»PμPeC,PsCĤPSPsC— C,P°C...C-PeP°CĤPrC-C—.
PhPrPSP°Pe PíCĤPë PrCfP¶Pμ PIPëCÍPsPePëC... PrPsP·P°C... PíPμCĤPëC,,PμCĤPëC†PSP°
CÍPμP»PμPeC,PëPIPSC-CÍC,CH PjPsP¶Pμ P±CfC,Pë PIC,CĤP°C†PμPSP°, C%oPs PjPsP¶Pμ
CÍPíCĤPëC†PëPSPëC,Pë P±CĤP°PrPëPeP°CĤPrC-CĤ C- PSPμPiP°C,PëPIPSPëPNë
C-PSPsC,CĤPsPíPSPëPNë PμC,,PμPeC,. PkP°PNëPíPsC€PëCĤPμPSC-C€C- PíPsP±C-C†PSC-
CĤPμP°PeC†C-C—, PíPsPíBĤ™ CİP·P°PSC- P· PíPμCĤPμPrPsP·CfPIP°PSPSCİPj, BĤ"
P°CĤC,PμCĤC-P°P»CHPSP° PiC-PíPsC,PμPSP·C-Cİ, P·P°PíP°PjPsCĤPsC†PμPSPSCİ,
PíPsP»PsPIPSPëPNë P±C-P»CH C,P° PíPsCÍPëP»PμPSPμ CÍPμCĤC†PμP±PëC,C,Cİ.

P>C-PeC¿PIP°PSPSCİ

PμCĤPë C,CİP¶PeC-PNë P°CĤC,PμCĤC-P°P»CHPSC-PNë PiC-PíPsC,PμPSP·C-C—
PSPμPsP±C...C-PrPSPs PIP¶PëC,Pë P·P°C...PsPrC-PI P°PeC,PëPIPSPsC—
PeP°CĤPrC-PsPIP°CÍPeCfP»CİCĤPSPsC— PiC-PrC,CĤPëPjPePë, PIPeP»CĤC†P°CĤC†Pë
C†P°CÍC,PëPNë PjPsPSC-C,PsCĤPëPSPi CÍPμCĤC†PμPIPsC— C,P° PrPëC...P°P»CHPSPsC—
C,,CfPSPeC†C-PNë, PSP°PrP°PSPSCİ PíP°C†C-C''PSC,Cf PíPsCĤPëP·PsPSC,P°P»CHPSPsPiPs
PíPsP»PsP¶PμPSPSCİ P· PíCĤPëPíC-PrPSCİC,PëPjPë PSPëP¶PSC-PjPë PeC-PSC†C-PIPeP°PjPë,
PePsPSC,CĤPsP»CH CĤC-PrPëPSPë, C%oPs C†PëCĤPeCfP»CĤC'', C,P°
CÍPμC†PsPIPëPíCfCÍPeP°PSPSCİ. P— PsPiP»CİPPrCf PSP° PíPsPrPsPIP¶PμPSCf
C,,P°CĤPjP°PePsP»PsPiC-C†PSCf PrC-CĤ P»PμCĤPeP°PSC-PrPëPíC-PSCf Cf CĤP°P·C-
PíPμCĤPμPrPsP·CfPIP°PSPSCİ C'' PSPμPsP±C...C-PrPSPëPj PSP°PiP»CİPPr P·P° CÍC,P°PSPsPj
CÍPμCĤC†PμPIPs-CÍCfPrPëPSPSPsC— CÍPëCÍC,PμPjPë C,P°PePëC... PíP°C†C-C''PSC,C-PI
C%oPsPSP°PNëPjPμPSC€Pμ PíCĤPsC,CİPiPsPj 24 PíPsPrPëPS. P'PμCĤCfC†Pë PrPs CÍPIP°PiPë
PIPëCÍPsPePμ P·PíBĤ™ CİP·CfPIP°PSPSCİ P»PμCĤPeP°PSC-PrPëPíC-PSCf P· P±C-P»PeP°PjPë,
PrC-P°P»C-P· PjPsP¶Pμ P±CfC,Pë PSPμPμC,,PμPeC,PëPIPSPëPj. PμP°C†C-C''PSC,Pë P·

PsC†C–PeC†PIP°PSPsC† P†PsPjC–C†PSPsC† P°P±Ps C,C†P¶PePsC†
C–PSC,PpPeC†PëPeP°C†C–C”C† PjP°C†C,C† P±C†C,Pë P†C–Pr PSP°PiP»C†P†PsPj P†
C†PjPsPIP°C... C–PSC,PµPSC†PëPIPSpSc— C,PµC†P°P†C–C—.

PµPsP±C–C†PSC– C†PµP°PeC†C–C—

P—PiC–PrPSPs P· PrP°PSPëPjPë PeP»C–PSC–C†PSPëC... PrPsC†P»C–PrP¶PµPSC† C,P°
P†C–C†P»C†C†PµC”C†C,C†P°C†C–PµPSPsPiPs P·P°C†C,Psc†C†PIP°PSPSC† P†PsP±C–C†PSC–
C†PµP°PeC†C–C—, C†PeC– P·C†C†C,C†C–C†P°P»PëC†C† PSP°PµC†P°C†C,C–C†Pµ, B†” C†Pµ
P†PµC†PëC„PµC†PëC†PSC– PSP°P±C†C†PePë, PiPsP»PsPIPSPëPµ P±C–P»C†C,
P†C†PëP†P»PëPIPë, C,P°C...C–PeP°C†PrC–C† C– P†PsC†PëP»PµPSPµ C†PµC†C†PµP±PëC,C,C†.

Pj PSPëP¶C†PµPSP°PIPµPrPµPSC–Pµ C,P°P±P»PëC†C– P†PeP°P·P°PSPs P†PsP±C–C†PSC–
C†PµP°PeC†C–C—, P†C†Ps C†PeC– P†PsPIC–PrPsPjP»C†P»Psc†C† P†C–Pr C†P°C†
PeP»C–PSC–C†PSPëC... PrPsC†P»C–PrP¶PµPSC† C,P°
P†C–C†P»C†C†PµC”C†C,C†P°C†C–PµPSPsPiPs P·P°C†C,Psc†C†PIP°PSPSC†
P†C†PµP†P°C†P°C,C† C† PeC†P°C—PSP°C... C†PIC–C,C† C,P° PrP»C† C†PePëC... P±C†PI
PsP±T’C†C†PSC,PsPIP°PSPëPµ P†C†PëC†PëPSPSPs–PSP°C†P»C–PrPePsPIPëPµ
P·P†B†” C†P·PsPe C–P· P·P°C†C,Psc†C†PIP°PSPSC†Pj P†C†PµP†P°C†P°C,C†.
PµPsP±C–C†PSC– C†PµP°PeC†C–C— PSP°PIPµPrPµPSPs P·PiC–PrPSPs P·
PeP»P°C†PëC„C–PeP°C†C–C”C† MedDRA C,P° C†P°C†C,Psc,Psc†
PIPëPSPëPePSPµPSPSC†: PrC†P¶Pµ C†P°C†C,Psc (≥ 1/10), C†P°C†C,Psc (≥ 1/100, < 1/10),
PSPµC†P°C†C,Psc (≥ 1/1000, < 1/100), C†C–PrPePs (≥ 1/10000, < 1/1000), PrC†P¶Pµ C†C–PrPePs
(< 1/10000), C†P°C†C,Psc,P° PSPµPIC–PrPsPjP° (PSPµ PjPsP¶PSP° PIPëP·PSP°C†PëC,Pë
P·PiC–PrPSPs P· PSP°C†PIPSPsC† C–PSC„Psc†PjP°C†C–C”C†). Pj PePsP¶PSC–Pµ
P†C†C†PiC– PIC–PrP†PsPIC–PrPSPs PrPs C†P°C†C,Psc,Pë PIPëPSPëPePSPµPSPSC†
C†PµP°PeC†C–C— P·P°P·PSP°C†PµPSPs P† P†PsC†C†P†PeC† P·PjPµPSC†PµPSPSC† C–C...
C†PµC†PµP·PSPsc†C,C–.

MedDRA PeP»P°C†PëC„C–PeP°C†C–C† C†PëC†C,PµPj C,P° PsC†P†PSC–PI	P§P°C†C,Psc	PkPµC†P°C†C,Psc	P C–PrPePs	P§P°C†C,Psc,P° PSPµPIC–PrPsPjP°
P— P±PsPeC± C–PjC±PSPSPsC— C†PëC†C,PµPjPë	B†”	B†”	P†C–PrPIPëC%PµPSP° C†C†C,P»PëPIC–C†C,C†	B†”
P— P±PsPeC± PSPµC†PIPSPsC— C†PëC†C,PµPjPë	P†PsP»PsPIPSPëPµ P±C–P»C†	P·P°P†P°PjPsC†PµPSPSC†	C†PsPSP»PëPIC–C†C,C†, PSPµP†C†PëC,PspjPSC–C†C,C†	B†”
P— P±PsPeC± C†PµC†C†C†	C,P°C...C–PeP°C†PrC–C†, P†PsC†PëP»PµPSPµ C†PµC†C†PµP±PëC,C,C†	B†”	C†C,PµPSPsPeP°C†PrC–C†	B†”
P— P±PsPeC± C†C±PrPëPS	P†C†PëP†P»PëPIPë	P°C†C,PµC†C–P°P»C†PSP° PiC–PiPsC,PµPSP·C–C†	B†”	B†”
P— P±PsPeC± C†P»C±PSPPePsPIPSPs– PePëC†PePsPIPSPiPs C,C†P°PeC,C±	B†”	PrPëC†P†PµP†C†C–C†, PSC†PrPsC,P°, P±C–P»C† C† P†PµC†C...PSC–Pµ C†P°C†C,PëPSC– P¶PëPiPsC,P°	P±P»C†PIP°PSPSC†, PrC–P°C†PµC†	P†C–P†PµC†C,C†PsC„C–C† C†C†PµPSP*, P†PsPjC†C,PSC–PSPSC† P†PµC†PëC,PspPµP°P»C†PSPs–PiPs PµPëC†C†PrP°C,C†*
P— P±PsPeC± PiPµPiP°C,PspP±C–P»C–P°C†PSPsC— C†PëC†C,PµPjPë	B†”	B†”	B†”	P†C–PrPIPëC%PµPSPSC† C†C–PIPSC† C,C†P°PSC†P°PjC–PSP°P· C†PëC†PsPIP°C,PëPë PeC†PsPIC–*
P— P±PsPeC± C†PeC–C†Pë C,P° PiC–PrC†PeC–C†PSPsC— C,P°PSPëPSPë	B†”	PIPëC†PëP†, C†PIPµC†P±C–P¶	PIPëC†PëP†P°PSPSC†	PSP°P±C†C†Pe*
P— P±PsPeC± C†PëPµP»PµC,PSPs– PjPë” C†P·PsPIPSPsC— C†PëC†C,PµPjPë C,P° C†PiPsP»C±C†PSPsC— C,P°PSPëPSPë	B†”	PjC–P°P»PiC–C†	B†”	B†”
P— P±PsPeC± C†PµC†PsPIPëPrC–P»C†PSPsC— C†PëC†C,PµPjPë	B†”	P†PsP»C–C†C†C–C†	P†PsP»P°PeC–C†C†C–C†	B†”
P— P°PiP°P·C†PSC– P†PsC†C±C†PµPSPSC† C,P° C†PµP°PeC†C–C— C± PjC–C†C†C– PIPIPµPrPµPSPSC†	P†PµC†PëC„PµC†PëC†PSC– PSP°P±C†C†PePë	P°C†C,PµPSC–C†, PiC–PrPIPëC%PµPSP° PIC,PspjP»C†PIP°PSC–C†C,C†	P±C–P»C† C† P†C†C†PrC†C... B†”	B†”

*P_μPsP±C-C†PSC- C₇P_μP°PeC†C-C— P·C- C†P_iPsPSC,P°PSPSPëC... P_iPsPIC-PrPsPjP»P_μPSCH P_iC-Pr C†P°C† P_iC-C†P»C₁C₇P_μC”C†C,C₇P°C†C-PN₆PSPsPiPs P·P°C†C,PsC†C†PIP°PSPSC₁ C† PeC₇P°C—PSP°C... C†PIC-C,C†.

P>P_μC₇PeP°PSC-PrPëP_iC-PS PSP_μ PIP_iP»PëPIP°C” PSP_μPiP°C,PëPIPSPs PSP° C₇C-PIP_μPSCH C†C†PeC₇C† PI PeC₇PsPIC- C,P° C₇C-PIP_μPSCH P»C-P_iC-PrC-PI C† C†PëC₇PsPIP°C,C†C- PeC₇PsPIC-.

PJ P_iP»P°C†P_μP±Ps-PePsPSC,C₇PsP»CHPsPIP°PSPëC... PeP»C-PSC-C†PSPëC... PrPsC†P»C-PrP₁P_μPSPSC₁C... P_iP_μC₇PëC,,P_μC₇PëC†PSC- PSP°P±C₇C₁PePë C†C,P°PSPsPiPëP»PëB B B B B B B B B B B B 0,9 % PSP° C,P»C- P_iC₇PëPN₆PsPjC† P»P_μC₇PeP°PSC-PrPëP_iC-PSC† C† PrPsP·C- 10B₇”20 PjPi C,P° 0,83 % B₇” P_iC₇Pë P_iC₇PëPN₆PsPjC- P_iP»P°C†P_μP±Ps. P_iC₁ C†P°C†C,PsC,P° PrPsC†C₁PiP°P»P° 2 % C† P·P°PiP°P»CHPsPSC-PN₆ P_iPsPiC†P»C₁C†C-C— PrPsC†P»C-PrP₁P_μPSPSC₁, PIPeP»CHC†P°CHC†Pë PrPsPiPiPsC,C₇PëPIP°P»C- PeP»C-PSC-C†PSC- PIPëP_iC₇PsP±C†PIP°PSPSC₁.

P—P°C†C,PsC†C†PIP°PSPSC₁ PrP_μC₁PePëC... PrPëPiC-PrC₇PsPiC-C₇PëPrPëPSC-PI PjPsP₁P_μ C-PSPsPrC- P_iC₇PëP·PiPsPrPëC,Pë PrPs P_iC₇P_μPePsC₇PrC-P°P»CHPsPsPiPs P±PsP»CH P°P±Ps C†C,P_μPSPsPeP°C₇PrC-C—, C† PIPëPSC₁C,PePsPiPëC... PIPëPiP°PrPeP°C... C† P_iP°C†C-C”PSC,C-PI P·C- C†C,P_μPSPsPeP°C₇PrC-C”CH PjPsP₁P_μ P·P±C-P»CHC€PëC,PëC†C₁ C†P°C†C,PsC,P°, C,C₇PëPIP°P»C-C†C,CH P°P±Ps C,C₁P₁PeC-C†C,CH PSP°P_iP°PrC-PI, PjPsP₁C†C,CH PIC-PrP·PSP°C†P°C,PëC†C₁ C-P·PsP»CHPsPiP°PSC- PIPëPiP°PrPePë C-PSC,,P°C₇PeC,C† PjC-PsPeP°C₇PrP°.

P_μPsPIC-PrPsPjP»P_μPSPSC₁ P_iC₇Ps P_iC-PrPsP·C₇CHPIP°PSC- P_iPsP±C-C†PSC- C₇P_μP°PeC†C-C—. P_μPsPIC-PrPsPjP»P_μPSPSC₁ P_iC₇Ps P_iPsP±C-C†PSC- C₇P_μP°PeC†C-C— P_iC-C†P»C₁ C₇P_μC”C†C,C₇P°C†C-C— P»C-PeP°C₇C†CHPePsPiPs P·P°C†PsP±C† PjP°C” PIP°P₁P»PëPIP_μ P·PSP°C†P_μPSPSC₁. P_iP_μ PrP°C” P·PjPsPiC† P_iC₇PsPiPsPrPëC,Pë PjPsPSC-C,PsC₇PëPSPi C†P_iC-PIPIC-PrPSPsC€P_μPSPSC₁ PePsC₇PëC†C,CH / C₇PëP·PëPe P_iC₇Pë P·P°C†C,PsC†C†PIP°PSPSC- C†CHPsPiPs P»C-PeP°C₇C†CHPePsPiPs P·P°C†PsP±C†. P_μP_μPrPëC†PSPëPj C,P° C,,P°C₇PjP°C†P_μPIC,PëC†PSPëPj P_iC₇P°C†C-PIPSPëPeP°Pj, P° C,P°PePsP₁ P_iP°C†C-C”PSC,P°Pj P°P±Ps C—C... P·P°PePsPSPSPëPj P_iC₇P_μPrC†C,P°PIPSPëPeP°Pj C†P»C-Pr P_iPsPIC-PrPsPjP»C₁C,Pë P_iC₇Ps C†C†C- PIPëPiP°PrPePë P_iC-PrPsP·C₇CHPIP°PSPëC... P_iPsP±C-C†PSPëC... C₇P_μP°PeC†C-PN₆ C,P° PIC-PrC†C†C,PSPsC†C,C- P_μC,,P_μPeC,PëPIPSPsC†C,C- P»C-PeP°C₇C†CHPePsPiPs P·P°C†PsP±C† C†P_μC₇P_μP· P_hPIC,PsPjP°C,PëP·PsPiP°PSC† C-PSC,,PsC₇PjP°C†C-PN₆PSC† C†PëC†C,P_μPjC† P· C,,P°C₇PjP°PePsPSP°PiP»C₁PrC† P·P° P_iPsC†PëP»P°PSPSC₁Pj: <https://aisf.dec.gov.ua>.

P₇P_μC₇PjC-PS P_iC₇PëPrP°C,PSPsC†C,C-

2 C₇PsPePë.

PJPjPsPIPë P·P±P_μC₇C-PiP°PSPSC₁

P—P±P_μC₇C-PiP°C,Pë P_iC₇Pë C,P_μPjPiP_μC₇P°C,C†C₇C- PSP_μ PIPëC‰P_μ 25 B°P₇ PI PsC₇PëPiC-PSP°P»CHPsPSC-PN₆ C†PiP°PePsPIC†C-.

P—P±P_μC₇C-PiP°C,Pë C† PSP_μPrPsC†C,C†PiPSPsPjC† PrP»C₁ PrC-C,P_μPN₆ PjC-C†C†C-.

PJPiP°PePsPIPeP°

PμPs 10 C,P°P±P»PμC,PsPe Cr P±P»C-CÍC,PμCȚC-, PīPs 3 P°P±Ps 6 P±P»C-CÍC,PμCȚC-PI CȚP°P·PsPj P· C-PSCÍC,CȚCrPeC†C-C"CH PrP»CȚ PjPμPrPëC†PSPsPiPs P·P°CÍC,PsCÍCrPIP°PSPSCȚ Cr PīP°C†C- P· PeP°CȚC,PsPSCÍ.

PμPs 30 P°P±Ps 60 C,P°P±P»PμC,PsPe Cr P±P°PSC†C-, PīPs 1 P±P°PSC†C- CȚP°P·PsPj P· C-PSCÍC,CȚCrPeC†C-C"CH PrP»CȚ PjPμPrPëC†PSPsPiPs P·P°CÍC,PsCÍCrPIP°PSPSCȚ Cr PīP°C†C- P· PeP°CȚC,PsPSCÍ.

PљP°C,PμPiPsCȚC-CȚ PIC-PrPīCrCÍPeCr

P—P° CȚPμC†PμPīC,PsPj.

P—P°CȚPIPSPëPe

PŷPhP' PќP'Pα B«PњP†PљP PhPГ†PњB».

PњC-CÍC†PμP·PSP°C...PsPrP¶PμPSPSCȚ P·P°CȚPIPSPëPeP°

PJPeCȚP°C—PSP°, 01013, Pj. PљPëC—PI, PIC†P». P'CrPrC-PSP†CrCÍC,CȚC-C—, P±CrPr. 5.

PμPsPIC-PrPsPjPëC,Pë PīCȚPs PSPμP±P°P¶P°PSPμ CȚPIPëC‰Pμ PīCȚPë P·P°CÍC,PsCÍCrPIP°PSPSC- P»C-PeP°CȚCÍCHPePsPiPs P·P°CÍPsP±Cr P'Pë PjPsP¶PμC,Pμ C†PμCȚPμP· CÍPëCÍC,PμPjCr C„P°CȚPjP°PePsPSP°PiP»CȚPrCr PŷPhP' PќP'Pα B«PњP†PљP PhPГ†PњB» PSP° PSPsPjPμCȚ C,PμP»PμC„PsPSCÍ: 38 (050) 309-83-54 (C†C-P»PsPrPsP±PsPIPs) P°P±Ps PSP° e-mail: pharmacovigilance@microkhim.com.

P'PëCȚPsP±PSPëPe

PŷPhP' PќP'Pα B«PњP†PљP PhPГ†PњB» (PIC-PrPīPsPIC-PrP°P»CHPSPëPN P·P° PIPëPīCrCÍPe CÍPμCȚC-C—, PSPμ PIPeP»CHC†P°CHC†Pë PePsPSC,CȚPsP»CH/PIPëPīCȚPsP±CrPIP°PSPSCȚ CÍPμCȚC-C—).

AT B«P>CrP±PSPëC„P°CȚPjB» (PIC-PrPīPsPIC-PrP°P»CHPSPëPN P·P° PIPëCȚPsP±PSPëC†C,PIPs C,P° PePsPSC,CȚPsP»CH/PIPëPīCȚPsP±CrPIP°PSPSCȚ CÍPμCȚC-C—, PSPμ PIPeP»CHC†P°CHC†Pë PIPëPīCrCÍPe CÍPμCȚC-C—).

PњC-CÍC†PμP·PSP°C...PsPrP¶PμPSPSCȚ PIPëCȚPsP±PSPëPeC-PI C,P° P°PrCȚPμCÍP°

PjC-CfC†CЦ PïCђPsPIP°PrP¶PμPSPSCЦ C—C... PrC-CЦP»CHъPSPsCfC,C-

PJPeCђP°C—PSP°, 01013, Pj. PљPөC—PI, PICfP». P'CfPrC-PSPrCfCfC,CђC—C—, P±CfPr. 5.

PJPeCђP°C—PSP°, 37500, PцPsP»C,P°PICfCHъPeP° PsP±P»., Pj. P>CfP±PSPө, PICfP».
P'P°CђPIC-PSPePsPIP°, 16.