

Р†РќРЎРЎР РЈРЉРІР†РЇ

**PrP»CЦ PjPμPrPëC±PSPsPiPs P•P°CÍC,PsCÍCřPIP°PSPSCЦ P»C-PeP°CЦCÍCHPePsPiPs
P•P°CÍPsP±Cř**

P'P†P'PℏP™P Pℏ

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

$$PrC-C\bar{H}C\ddagger P^{\circ} \ C\bar{H}P\mu C\ddagger PsPIP\ddot{e}PSP^{\circ}: \ C\acute{r}P\ddot{e}P \gg PrP\mu PSP^{\circ}C,,C-P \gg C\acute{r} \ C\ddagger P\ddot{e}C,C\bar{H}P^{\circ}C,,$$

1 C,P°P±P»PμC,PeP° P¶CřPIP°P»CHbPSP° PjC-CřC,PëC,CH CřPëP»PrPμPSP°C,,C-P»Cř 50 PjPi Cř C,,PsCřPjC- CřPëP»PrPμPSP°C,,C-P»Cř C†PëC,CřP°C,Cř 70,24 PjPi P°P±Ps

1 C,P°P±P»P_μC,P_eP° P¶C_rPIP°P»CH₅PSP° P_jC-C_rC,P_eC,CH₅ C_rP_eP»PrP_μPSP°C,,C-P»C_r 100 P_jP_i C_r C,,P_sC₇P_jC- C_rP_eP»PrP_μPSP°C,,C-P»C_r C_rP_eC,C₇P°C,C_r 140,48 P_jP_i;

PrPsPiPsPjC-P¶PSC- C¶PµC¶PsPIPëPSPë: PeP°P»C-Ç¶ PiPsP»P°PeC¶PëP»C-PS; PjP°PiPSC-Ç¶ C¶C, PµP°C¶P°C,; PeC¶PµPjPSC-Ç¶ PrC-PsPeC¶PëPr PePsP»PsC—PrPSPëPñ P±PµP·PiPsPrPSPëPñ; P°C¶PiP°C¶C,P°Pj (P· 951); PSP°C,C¶C-Ç¶ PeC¶PsC¶PeP°C¶PjPµP»PsP·P°; P°C¶PsPjP°C,PëP·P°C,PsC¶ PjB¶™ C¶C,P° PiPµC¶C†PµPIP°; P»P°PeC,PsP·P°, PjPsPSPsPiC-PrC¶P°C,; PiPsPIC-PrPsPS Pñ-30.

$$\text{P}^{\circ}\text{C}-\text{PeP}^{\circ}\text{C}\dot{\text{H}}\text{C}\dot{\text{H}}\text{C}\text{H}\text{PeP}^{\circ}\text{C}_{\text{H}}\text{PsC}\dot{\text{H}}\text{PjP}^{\circ}\text{C}_{\text{H}}\text{P}^{\circ}\text{P}\pm\text{P}\gg\text{P}_{\text{H}}\text{C}_{\text{H}}\text{PeP}^{\circ}\text{P}\|\text{C}\dot{\text{H}}\text{P}\text{IP}^{\circ}\text{P}\gg\text{C}\text{H}\text{P}\text{SC}-.$$

PhCl'PSPsPIPSC- C,,C-P·PëPePs-C...C-PjC-C‡PSC- PIP»P°Cl'C,PëPIPsCl'C,C-: C,CḡPëPeCr'C,PSC- PrPIPsPsPïCr'PeP»C- C,P°P±P»PuC,PePë P±C-P»PsPiPs PePsP»CHbPsCḡCr' P· C,PëCl'PSPµPSPSCḡPj B«50B» P°P±Ps B«100B» P· PsPrPSPsPiPs P±PsPeCr'.

**P_μP°C₇PjP°PePsC,P_μC₇P°PiP_μPIc,PëC†PSP° PiC₇C†PiP°. P—P°C†PsP±Pë, C%Ps
P·P°C†C,Psc†PsPICrC₇C,C₇C†C† PiC₇Pë P_μC₇P_μPeC,PëP»CH₂PSC-PN₂
PrPëC†C,,C†PSPeC†C-C—. PÿPëP»PrP_μPSP°C,,C-P». P₇PsPr P₇PÿP† G04B E03.**

$P \rightleftharpoons P^{\circ} C \bar{H} P j P^{\circ} P e P s P \gg P s P i C - C \nmid P S C - P I P \gg P^{\circ} C \bar{I} C, P \ddot{e} P I P s C \bar{I} C, C - .$

$$P \propto P^{\circ} C_{\text{H}} P_j P^{\circ} P_{\epsilon} P_s P_r P_{\ddot{e}} P S P^{\circ} P_j C - P_{\epsilon} P^{\circ}.$$

$P\ddot{Y}P\ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg$ бҗ“ $PiC_{\eta}P_{\mu}PiP^{\circ}C_{\eta}P^{\circ}C, PrP \gg C_{\eta}PiP_{\mu}C_{\eta}PsC_{\eta}P^{\circ}P \gg C_{\eta}PSPsPiPs$
 $P \cdot P^{\circ}C_{\eta}^{\circ}C, PsC_{\eta}^{\circ}C_{\eta}PI P^{\circ}PSPSC_{\eta}, PiC_{\eta}P\ddot{e}P \cdot PSP^{\circ}C \ddagger P_{\mu}PSP\ddot{e}P_{\eta}$ $PrP \gg C_{\eta}P \gg C-PeC_{\eta}PI P^{\circ}PSPSC_{\eta}$
 $P_{\mu}C_{\eta}P_{\mu}PeC, P\ddot{e}P \gg C_{\eta}PSPsC - PrP\ddot{e}C_{\eta}^{\circ}C, C_{\eta}PSPeC_{\eta}^{\circ}C -$ $P_{\mu}C_{\eta}P\ddot{e} C_{\eta}^{\circ}P_{\mu}PeC_{\eta}^{\circ}C_{\eta}P^{\circ}P \gg C_{\eta}PSPsPjC_{\eta}$
 $P \cdot P \pm C_{\eta}PrP_{\eta}P_{\mu}PSPSC - PiC_{\eta}P_{\mu}PiP^{\circ}C_{\eta}P^{\circ}C, PIC - PrPSPsPI P \gg C_{\eta}C^{\circ}$ $P \cdot PSP\ddot{e}P_{\eta}P_{\mu}PSC_{\eta}$
 $P_{\mu}C_{\eta}P_{\mu}PeC, P\ddot{e}P \gg C_{\eta}PSC_{\eta} C, C_{\eta}PSPeC_{\eta}^{\circ}C - C_{\eta} C_{\eta}P \gg C_{\eta}C \dots PsPj PiPsC_{\eta}^{\circ}P\ddot{e}P \gg P_{\mu}PSPSC_{\eta}$
 $PiC_{\eta}P\ddot{e}C, PsPeC_{\eta}^{\circ}C_{\eta}C_{\eta}PsPIC - PrPs C_{\eta}^{\circ}C, P^{\circ}C, P_{\mu}PIPsPiPs C \ddagger P \gg P_{\mu}PSP^{\circ}$.

PαC-P·C-PsP»PsPiC-C‡PSPëPNë PjPμC...P°PSC-P·Pj, C%Ps P·CfPjPsPIP»CηC” PμCηPμPeC‡C-Cη,
PIPeP»CηC‡P°C” PIPëPIC-P»CHbPSPμPSPSC‡ PsPeC‡PëPrCf P°P·PsC,Cf (NO) Cf
PeP°PIPμCηPSPsP·PSPëC... C,C-P»P°C... Pïc-Pr C‡P°C‡ C‡PμPeC‡CfP°P»CHbPSPsPiPs
P·P±CfPrP¶PμPSPSC‡. P’PëPIC-P»CHbPSPμPSPëPNë NO P°PeC,PëPICfC” C,,PμCηPjPμPSC,
PiCfP°PSC-P»P°C,C‡PëPeP»P°P·Cf, C%Ps C‡C,PëPjCfP»CηC” Pïc-PrPIPëC%PμPSPSC‡
CηC-PIPSC‡ C‡PëPeP»C-C‡PSPsPiPs PiCfP°PSPsP·PëPSPjPsPSPsC,,PsC‡C,,P°C,Cf (C‡P”PηPα),
PIPëPeP»PëPeP°C” CηPsP·C‡P»P°P±P»PμPSPSC‡ PiP»P°PrPePsC— PjCfC‡PeCfP»P°C,CfCηPë
PeP°PIPμCηPSPsP·PSPëC... C,C-P» C,P° C‡PïcCηPëC‡C” PïcCηPëC,PsPeCf PeCηPsPIC-.

PŸPëP»PrPμPSP°C,,C-P» C” PïPsC,CfP¶PSPëPj C,P° C‡PμP»PμPeC,PëPIPSPëPj
C-PSPiC-P±C-C,PsCηPsPj C‡P”PηPα-C‡PïPμC‡PëC,,C-C‡PSPsC—
C,,PsC‡C,,PsPrC-PμC‡C,PμCηP°P·PëB B B B B B B B 5 (PαP”P·5) Cf PeP°PIPμCηPSPsP·PSPëC...
C,C-P»P°C..., PrPμ PαP”P·5 PIC-PrPiPsPIC-PrP°C” P·P° CηPsP·PiP°Pr C‡P”PηPα. P·C,,PμPeC,Pë
C‡PëP»PrPμPSP°C,,C-P»Cf PSP° PμCηPμPeC‡C-Cη PjP°CηC,CHb PïPμCηPëC,,PμCηPëC‡PSPëPNë
C...P°CηP°PeC,PμCη. PŸPëP»PrPμPSP°C,,C-P» PSPμ C‡PëPSPëC,CHb
P±PμP·PiPsC‡PμCηPμPrPSCηPsC— CηPμP»P°PeC‡P°C‡C-PNëPSPsC— PrC-C— PSP°
C-P·PsP»CHbPsPIP°PSC- PeP°PIPμCηPSPsP·PSC- C,C-P»P° P»CηPrPëPSPë, P°P»Pμ
PïPsC,CfP¶PSPs PïPsC‡PëP»CηC” CηPsP·C‡P»P°P±P»CηPIP°P»CHbPSCf PrC-Cη NO PSP° C‡Cη
C,PeP°PSPëPSCf. PμCηPë P°PeC,PëPIP°C‡C-C— PjPμC,P°P±PsP»C-C‡PSPsPiPs CëP»C‡C...Cf
NO/C‡P”PηPα, C%Ps PIC-PrP±C‡PIP°C”C,CHbC‡C‡ PïcCηPë C‡PμPeC‡CfP°P»CHbPSC-PNë
C‡C,PëPjCfP»C‡C‡C-C—, C-PSPiC-P±C‡PIP°PSPSC‡ C‡PëP»PrPμPSP°C,,C-P»PsPj PαP”P·5
C‡PïcCηPëC‡PëPSC‡C” Pïc-PrPIPëC%PμPSPSC‡ CηC-PIPSC‡ C‡P”PηPα Cf
PeP°PIPμCηPSPsP·PSPëC... C,C-P»P°C.... PŸP°PePëPj C‡PëPSPsPj, PrP»C‡C,C,PsPiPs, C%PsP±
Pïc-C‡P»C‡C P·P°C‡C,PsC‡C‡PIP°PSPSC‡ C‡PëP»PrPμPSP°C,,C-P»Cf PIPëPSPëPe
PSPμPsP±C...C-PrPSPëPNë C,,P°CηPjP°PePsP»PsPiC-C‡PSPëPNë PμC,,PμPeC,,
PSPμPsP±C...C-PrPSPμ C‡PμPeC‡CfP°P»CHbPSPμ P·P±CfPrP¶PμPSPSC‡.

P”PsC‡P»C-PrP¶PμPSPSC‡ in vitro PïcCηPsPrPμPjPsPSC‡C,CηCfPIP°P»Pë, C%Ps
C‡PëP»PrPμPSP°C,,C-P» C” C‡PμP»PμPeC,PëPIPSPëPj PrPs PαP”P·5, C%Ps P±PμCηPμ
P°PeC,PëPIPSCf C‡C‡P°C‡C,CHb Cf PïcCηPsC‡PμC‡C- PμCηPμPeC‡C-C—. P·C,,PμPeC,
C‡PëP»PrPμPSP°C,,C-P»Cf PSP° PαP”P·5 PïPsC,CfP¶PSC-CëPëPNë, PSC-P¶ PSP° C-PSCëC-
PIC-PrPsPjC- C,,PsC‡C,,PsPrC-PμC‡C,PμCηP°P·Pë. PïPμPNë PμC,,PμPeC, Cf 10 CηP°P·C-PI
PïPsC,CfP¶PSC-CëPëPNë, PSC-P¶ PμC,,PμPeC, PSP° PαP”P·6, C%Ps P±PμCηPμ C‡C‡P°C‡C,CHb Cf
PïcCηPsC‡PμC‡P°C... C,,PsC,PsPïPμCηPμC,PIPsCηPμPSPSC‡ Cf C‡C-C,PeC-PIC‡C-. PμCηPë
P·P°C‡C,PsC‡C‡PIP°PSPSC- PjP°PeC‡PëPjP°P»CHbPSPëC... CηPμPePsPjPμPSPPrPsPIP°PSPëC...
PrPsP· C‡PμP»PμPeC,PëIPSC-C‡C,CHb C‡PëP»PrPμPSP°C,,C-P»Cf PrPs PαP”P·5 Cf 80
CηP°P·C-PI PïPμCηPμPIPëC%CfC” PNëPsPiPs C‡PμP»PμPeC,PëIPSC-C‡C,CHb PrPs PαP”P·1, Cf
700 CηP°P·C-PI PIPëC%P°, PSC-P¶ PrPs PαP”P·2, PαP”P·3, PαP”P·4, PαP”P·7, PαP”P·8,
PαP”P·9, PαP”P·10 C,P° PαP”P·11. P—PsPeCηPμPjP°, C‡PμP»PμPeC,PëIPSC-C‡C,CHb
C‡PëP»PrPμPSP°C,,C-P»Cf PrPs PαP”P·5 Cf 4000 CηP°P·C-PI PïPμCηPμPIPëC%CfC” PNëPsPiPs
C‡PμP»PμPeC,PëIPSC-C‡C,CHb PrPs PαP”P·3 Bη” C‡PηPηPα-C‡PïPμC‡PëC,,C-C‡PSPsC—
C-P·PsC,,PsCηPjPë C,,PsC‡C,,PsPrC-PμC‡C,PμCηP°P·Pë, C%Ps P±PμCηPμ C‡C‡P°C‡C,CHb Cf
CηPμPiCfP»C‡C‡C-C— C‡PμCηC‡PμIPsC— C‡PePsCηPsC,P»PëIPsC‡C,C-.

PαP°CηPjP°PePsPeC-PSPμC,PëPeP°.

PηP±C‡PsCηP±C‡C-C‡C. PŸPëP»PrPμPSP°C,,C-P» CëPIPëPrPePs P°P±C‡PsCηP±CfC”C,CHbC‡C‡C.
PηP°PeC‡PëPjP°P»CHbPSC- PïP»P°P·PjPsPIC- PePsPSC‡PμPSC,CηP°C‡C-C— PïcCηPμPiP°CηP°C,Cf
PrPsC‡C‡PiP°CηC,CHbC‡C‡C PïcCηPsC,C‡PiPsPj 30Bη”120 C...PIPëP»PëPS (C-P·
PjPμPrC-P°PSPsCη 60 C...PIPëP»PëPS) Pïc-C‡P»C‡C PNëPsPiPs PïPμCηPsCηP°P»CHbPSPsPiPs
P·P°C‡C,PsC‡C‡PIP°PSPSC‡ PSP°C,C%Pμ. PŸPμCηPμPrPSC‡ P°P±C‡PsP»CηC,PSP°
P±C-PsPrPsC‡C,CfPïPSC-C‡C,CHb Pïc-C‡P»C‡C PïPμCηPsCηP°P»CHbPSPsPiPs

P·P°CfC,PsCfCfPIP°PSPSCf CfC,P°PSPsPIPëC,CH 41 % (C-P· PrC-P°PïP°P·PsPSPsPj
P·PSP°CfPµPSCf PIC-Pr 25 PrPs 63 %). Pj CfPµPePsPjPµPSPPrPsPIP°PSPsPjCf
PrC-P°PïP°P·PsPSC- PrPsP· (PIC-Pr 25 PrPs 100 PjPi) PïP»PsC%°P° PïC-Pr
C„P°CfPjP°PePsPeC-PSPµC,PëCfPSPsCf PeCfPëPIPsCf B«PePsPSCfPµPSC,CfP°CfC-Cf-
CfP°CfB» (AUC) C,P° PjP°PeCfPëPjP°P»CHfPSP° PePsPSCfPµPSC,CfP°CfC-Cf (PŸ_{max})
CfPëP»PrPµPSP°C„C-P»Cf PïC-CfP»Cf PNePsPiPs PïPµCfPsCfP°P»CHfPSPsPiPs
P·P°CfC,PsCfCfPIP°PSPSCf PïC-PrPIPëC%°CfCfC,CHCfCf PïCfPsPiPsCfCfC-PNePSPs PrPs
PrPsP·Pë.

PµCfPë P·P°CfC,PsCfCfPIP°PSPSC- CfPëP»PrPµPSP°C„C-P»Cf PïC-Pr CfP°Cf
PIP¶PëPIP°PSPSCf C—P¶C- CfC,CfPïC-PSCH P°P±CfPsCfP±CfC-C— P·PSPëP¶CfC”C,CHCfCf
P·C- CfPµCfPµPrPSC-Pj PïPsPrPsPIP¶PµPSPSCfPj CfP°CfCf PrPsCfCfPiPSPµPSPSCf
PjP°PeCfPëPjP°P»CHfPSPsC— PePsPSCfPµPSC,CfP°CfC-C— (T_{max}) PrPs 60 C...PIPëP»PëPS C-
CfPµCfPµPrPSC-Pj P·PSPëP¶PµPSPSCfPj C_{max} PSP° 29 %.

P PsP·PïPsPrC-P». PŸPµCfPµPrPSC-PNe CfC-PIPSPsPIP°P¶PSPëPNe PsP±Bf™ C”Pj
CfPsP·PïPsPrC-P»Cf (V_d) CfC,P°PSPsPIPëC,CH 105 P»C-C,CfC-PI, C%°Ps CfPIC-PrCfPëC,CH
PïCfPs CfPsP·PïPsPrC-P» PïCfPµPïP°CfP°C,Cf PI C,PeP°PSPëPSP°C... PsCfPiP°PSC-P·PjCf.
PµC-CfP»Cf PsPrPSPsCfP°P·PsPiPsPiPs PïPµCfPsCfP°P»CHfPSPsPiPs
P·P°CfC,PsCfCfPIP°PSPSCf CfPëP»PrPµPSP°C„C-P»Cf PI PrPsP·C- 100 PjPi CfPµCfPµPrPSCf
PjP°PeCfPëPjP°P»CHfPSP° P·P°PïP°P»CHfPSP° PïP»P°P·PjPsPIP° PePsPSCfPµPSC,CfP°CfC-Cf
CfPëP»PrPµPSP°C„C-P»Cf CfC,P°PSPsPIPëC,CH PïCfPëP±P»PëP·PSPs 440 PSPi/PjP»
(PePsPµC„C-CfC-C”PSC, PIP°CfC-P°CfC-C— CfC,P°PSPsPIPëC,CH 40 %). PhCfPeC-P»CHfPePë
P·PiBf™ CfP·CfPIP°PSPSCf CfPëP»PrPµPSP°C„C-P»Cf C,P° PNePsPiPs PiPsP»PsPiPSPsPiPs N-
PrPµCfPjPµC,PëP»PjPµC,P°P±PsP»C-C,Cf P· P±C-P»PeP°PjPë PïP»P°P·PjPë PeCfPsPIC-
PrPsCfCfPiP°C” 96 %, CfPµCfPµPrPSCf PjP°PeCfPëPjP°P»CHfPSP° PïP»P°P·PjPsPIP°
PePsPSCfPµPSC,CfP°CfC-Cf PIC-P»CHfPSPsPiPs CfPëP»PrPµPSP°C„C-P»Cf PrPsCfCfPiP°C” 18
PSPi/PjP» (38 PSPjPsP»CH). PŸC,CfPïC-PSCH P·PiBf™ CfP·CfPIP°PSPSCf P· P±C-P»PeP°PjPë
PïP»P°P·PjPë PeCfPsPIC- PSPµ P·P°P»PµP¶PëC,CH PIC-Pr P·P°PïP°P»CHfPSPëC...
PePsPSCfPµPSC,CfP°CfC-PNe CfPëP»PrPµPSP°C„C-P»Cf.

Pj P·PrPsCfPsPIPëC... PrPsP±CfPsPiPsP»CHCfC-PI, CfPeC- P·P°CfC,PsCfPsPICfPIP°P»Pë
CfPëP»PrPµPSP°C„C-P» PsPrPSPsCfP°P·PsPiPs PI PrPsP·C- 100 PjPi, CfPµCfPµP·
90 C...PIPëP»PëPS PI PµCfPeCfP»CfC,C- PIPëP·PSP°CfP°P»PsCfCf PjPµPSCfPµ 0,0002 % (Cf
CfPµCfPµPrPSCfPsPjCf 188 PSPi) P·P°CfC,PsCfPsPIP°PSPsC— PrPsP·Pë.

P’C-PsC,CfP°PSCfC„PsCfPjP°CfC-Cf. PµPµC,P°P±PsP»C-P·Pj CfPëP»PrPµPSP°C„C-P»Cf
PIC-PrP±CfPIP°C”C,CHCfCf PiPsP»PsPiPSPëPj CfPëPSPsPj P·P° CfCfP°CfC,CH
PjC-PeCfPsCfPsPjP°P»CHfPSPëC... C-P·PsC„PµCfPjPµPSC,C-PI PïPµCfC-PSPëPë CYP3A4
(PiPsP»PsPiPSPëPNe CëP»CfC...) C,P° CYP2C9 (PrCfCfPiPsCfCfPrPSPëPNe CëP»CfC...)
P”PsP»PsPiPSPëPNe CfPëCfPeCfP»CfCfCfPëPNe PjPµC,P°P±PsP»C-C,
CfC,PIPsCfCfC”C,CHCfCf CëP»CfC...PsPj N-PrPµPjPµC,PëP»CHfPIP°PSPSCf
CfPëP»PrPµPSP°C„C-P»Cf. PŸPµP»PµPeC,PëPIPSC-CfC,CH PjPµC,P°P±PsP»C-C,Cf C%°PsPrPs
PµP”P·5 PïPsCfC-PIPSCfPSPSP° P·C- CfPµP»PµPeC,PëPIPSC-CfC,Cf
CfPëP»PrPµPSP°C„C-P»Cf, P° P°PeC,PëPIPSC-CfC,CH PjPµC,P°P±PsP»C-C,Cf C%°PsPrPs
PµP”P·5 CfC,P°PSPsPIPëC,CH PïCfPëP±P»PëP·PSPs 50 % P°PeC,PëPIPSPsCfC,C-
PIPëC...C-PrPSPsC— CfPµCfPsPiPëPSPë. PµP»P°P·PjPsPIC- PePsPSCfPµPSC,CfP°CfC-C—
CfCHfPsPiPs PjPµC,P°P±PsP»C-C,Cf CfC,P°PSPsPIP»CfC,CH PïCfPëP±P»PëP·PSPs 40 %
PePsPSCfPµPSC,CfP°CfC-C— CfPëP»PrPµPSP°C„C-P»Cf Cf PïP»P°P·PjC- PeCfPsPIC-. N-
PrPµPjPµC,PëP»CHfPsPIP°PSPëPNe PjPµC,P°P±PsP»C-C, P·P°P·PSP°C” PïPsPrP°P»CHCëPsPiPs
PjPµC,P°P±PsP»C-P·PjCf, P° PïPµCfC-PsPr PNePsPiPs PSP°PïC-PIPIPëPIPµPrPµPSPSCf

СГГ,Р°PSPsPIPëC,CH ПГГПëP±P»PëP·PSPs 4 PiPsPrPëPSPë.

Р'PëPIPμPrPμPSPSCЦ. Р—Р°PiP°P»CHPSPëPNë PeP»C-ГГPμPSCГГ ГГ'PëP»PrPμPSP°C,,C-P»Cr
СГГ,Р°PSPsPIPëC,CH 41 P»/PiPsPr, P·CrPjPsPIP»CHCHC±Pë PiPμГГC-PsPr PNëPsPiPs
PSP°PiC-PIPIPëPIPμPrPμPSPSCЦ C,ГГPëPIP°P»C-СГГ,CH 3БГ "5 PiPsPrPëPS. ПГPe PiC-ГГ'P»ЦЦ
PiPμГГPsГГP°P»CHPSPsPiPs, C,Р°Pe C- PiC-ГГ'P»ЦЦ PIPSCГГ,ГГC-СëPSCHPsPIPμPSPSPsPiPs
P·Р°СГГ,PsСГГCrPIP°PSPSCЦ PμPeСГГPeГГPμCГC-ЦЦ ГГ'PëP»PrPμPSP°C,,C-P»Cr Cr
PIPëPiP»ЦЦPrC- PjPμC,Р°P±PsP»C-C,C-PI P·PrC-PNëСГГPSCГГ"C,CHCГГЦЦ PiPsP»PsPIPSPëPj
C±PëPSPsPj C-P· PeP°P»PsPj (PiГГPëP±P»PëP·PSPs 80 % PIPIPμPrPμPSPsC—
PiPμГГPsГГP°P»CHPSPsC— PrPsP·Pë) C,Р° PjPμPSCëPsCH PjC-ГГPsCH P·C- ГГ'PμC±PμCH
(PiГГPëP±P»PëP·PSPs 13 % PIPIPμPrPμPSPsC— PiPμГГPsГГP°P»CHPSPsC— PrPsP·Pë).

PαP°ГГPjP°PePsPeC-PSPμC,PëPeP° Cž PsСГГPsP±P»PëPIPëC... PiГГCžPi PiP°CГC-C"PSC,C-PI.

PμP°CГC-C"PSC,Pë P»C-C,PSCHPsPiPs PIC-PeCr. PJ P·PrPsГГPsPIPëC...
PrPsP±ГГPsPIPsP»CHCГC-PI P»C-C,PSCHPsPiPs PIC-PeCr (PIC-PePsPj PIC-Pr 65 ГГPsPeC-PI)
PIC-PrP·PSP°C±P°P»PsСГГЦЦ P·PSPëPμPμPSPSCЦ PeP»C-ГГPμPSCГГCr
ГГ'PëP»PrPμPSP°C,,C-P»Cr, C%oPs P·CrPjPsPIP»CHPIP°P»Ps PiC-PrPIPëC%oPμPSPSCЦ
PiP»P°P·PjPsPIPëC... PePsPSCГPμPSC,ГГP°CГC-PNë ГГ'PëP»PrPμPSP°C,,C-P»Cr C,Р° PNëPsPiPs
Р°PeC,PëPIPSPsPiPs N-PrPμPjPμC,PëP»CHPsPIP°PSPsPiPs PjPμC,Р°P±PsP»C-C,Cr
PiГГPëP±P»PëP·PSPs PSP° 90 % PiPsГГC-PIPSCЦPSPs P· PIC-PrPiPsPIC-PrPSPëPjPë
PePsPSCГPμPSC,ГГP°CГC-ЦЦPjPë Cr P·PrPsГГPsPIPëC... PrPsP±ГГPsPIPsP»CHCГC-PI
PjPsP»PsPrCëPsPiPs PIC-PeCr (18БГ "45 ГГPsPeC-PI). PJ P·PIБГ™ ЦЦP·PeCr P· PIC-PePsPIPëPjPë
PIC-PrPjC-PSPSPсГГC,ЦЦPjPë Cr P·PIБГ™ ЦЦP·CrPIP°PSPSC- P· P±C-P»PeP°PjPë PiP»P°P·PjPë
PeГГPsPIC- PIC-PrPiPsPIC-PrPSPμ PiC-PrPIPëC%oPμPSPSCЦ PiP»P°P·PjPsPIPsC—
PePsPSCГPμPSC,ГГP°CГC-C— PIC-P»CHPSPsPiPs ГГ'PëP»PrPμPSP°C,,C-P»Cr
СГГ,Р°PSPsPIPëP»Ps PiГГPëP±P»PëP·PSPs 40 %.

РКPëГГPePsPIP° PSPμPrPsСГГ,Р°C,PSC-СГГ,CH. PJ PrPsP±ГГPsPIPsP»CHCГC-PI C-P·
PiPsГГCrCëPμPSPSCЦPjPë C,,CrPSPeCГC-C— PSPëГГPsPe P»PμPiPePsPiPs C,Р°
PiPsPjC-ГГPSPsPiPs СГГ,CrPiPμPSCЦ (PeP»C-ГГPμPSCГГ PeГГPμP°C,PëPSC-PSCr 30БГ "80
PjP»/C...PI) C,,Р°ГГPjP°PePsPeC-PSPμC,PëPeP° ГГ'PëP»PrPμPSP°C,,C-P»Cr
P·Р°P»PëCëP°P»Р°СГГЦЦ PSPμP·PjC-PSPμPSPsCH PiC-ГГ'P»ЦЦ PNëPsPiPs
PsPrPSPsГГP°P·PsPIPsPiPs PiPμГГPsГГP°P»CHPSPsPiPs P·Р°СГГ,PsСГГCrPIP°PSPSCЦ Cr
PrPsP·C- 50 PjPi. PŸPμГГPμPrPSC- AUC C,Р° C_{max} N-PrPμPjPμC,PëP»CHPsPIP°PSPsPiPs
PjPμC,Р°P±PsP»C-C,Cr PiC-PrPIPëC%oCrPIP°P»PëСГГЦЦ PSP° 126 % C,Р° 73 %
PIC-PrPiPsPIC-PrPSPs PiPsГГC-PIPSCЦPSPs P· C,Р°PePëPjPë PiPsPeP°P·PSPëPeP°PjPë Cr
PrPsP±ГГPsPIPsP»CHCГC-PI C,Р°PePsPiPs Pμ PIC-PeCr P±PμP· PiPsГГCrCëPμPSCH
C,,CrPSPeCГC-C— PSPëГГPsPe. PhPrPSP°Pe C±PμГГPμP· PIPëСГГPsPeCr
C-PSPPrPëPIC-PrCrP°P»CHPSCr PIP°ГГC-Р°P±PμP»CHPSC-СГГ,CH CГC-
PIC-PrPjC-PSPSPсГГC,C- PSPμ P±CrP»Pë СГГ,Р°C,PëСГГ,C,PëC±PSPs P·PSP°C±CrC%oPëPjPë.

PJ PrPsP±ГГPsPIPsP»CHCГC-PI C-P· C,ЦЦPμPePëPjPë PiPsГГCrCëPμPSPSCЦPjPë
C,,CrPSPeCГC-C— PSPëГГPsPe (PeP»C-ГГPμPSCГГ PeГГPμP°C,PëPSC-PSCr PSPëPμC±Pμ 30
PjP»/C...PI) PeP»C-ГГPμPSCГГ ГГ'PëP»PrPμPSP°C,,C-P»Cr P·PSPëPμCrPIP°PICГГЦЦ, C%oPs
СГГPiГГPëC±PëPSCЦP»Ps ГГ'PμГГPμPrPSC" PiC-PrPIPëC%oPμPSPSCЦ AUC C,Р° C_{max} PSP° 100 %
C,Р° 88 % PIC-PrPiPsPIC-PrPSPs PiPsГГC-PIPSCЦPSPs C-P· PrPsP±ГГPsPIPsP»CHCГCЦЦPjPë
C,Р°PePsPiPs Pμ PIC-PeCr P±PμP· PiPsГГCrCëPμPSCH C,,CrPSPeCГC-C— PSPëГГPsPe.
PлГГC-Pj C,PsPiPs, P·PSP°C±PμPSPSCЦ AUC C,Р° C_{max} N-PrPμPjPμC,PëP»CHPsPIP°PSPsPiPs
PjPμC,Р°P±PsP»C-C,Cr P·PSP°C±CrC%oPμ PiC-PrPIPëC%oCrPIP°P»PëСГГЦЦ БГ " PSP° 200 % C,Р°
79 % PIC-PrPiPsPIC-PrPSPs.

РμРμC±C-PSPePsPIP° PSPμPrPsCfC, P° C, PSC-CfC, CH. PJ PrPsP±CfPsPIPsP»CHC†C-PI C-P· C†PëCfPsP·PsPj PīPμC±C-PSPePë P»PμPiPePsPiPs C, P° PīPsPjC-CfPSPsPiPs CfC, C†PīPμPSCfC (PeP»P°CfPë Ph C, P° P' P·P° PeP»P°CfPëC,, C-PeP°C†C-C" Cф P§P°PNëP»PrP°Bf "PμBf™ Cф) PeP»C-CfPμPSCf C†PëP»PrPμPSP° C,, C-P»C† P·PSPëP¶C†PIP°PICfCfC, C%Ps PīCfPëP·PIPsPrPëP»Ps PrPs PīC-PrPIPëC%PμPSPSCfC AUC (84 %) C, P° C_{max} (47 %) PīPsCfC-PIPSCfCPSPs P· PrPsP±CfPsPIPsP»CHC†CfPjPë C, P°PePsPiPs P¶ PIC-PeC† P±PμP· PīPsCfC†CëPμPSCф C,, C†PSPeC†C-C— PīPμC±C-PSPePë. PαP°CfPjP°PePsPeC-PSPμC, PëPeP° C†PëP»PrPμPSP° C,, C-P»C† C† PīP°C†C-C" PSC, C-PI C-P· PīPsCfC†CëPμPSPSCfCfPjPë C,, C†PSPeC†C-C— PīPμC±C-PSPePë C, CfP¶PePsPiPs CfC, C†PīPμPSCfC PSPμ PIPëPIC†P°P»P°CfCfC.

РлР»C-PSC-C†PSC- C...P°CfP°PeC, PμCfPëCfC, PëPePë.

PμPsPeP°P·P°PSPSCfC.

РμCfPμPīP°CfP° C, CfPμPePsPjPμPSPrC†C" C, CHCfCfC P·P°CfC, PsC†PsPIC†PIP° C, Pë C†PsP»PsPIC-PeP°Pj C-P· PμCfPμPeC, PëP»CHPSPsCф PrPëCfC,, C†PSPeC†C-C" Cф, CfPeP° PIPëP·PSP° C†P° C, CHCfCfC CfPe PSPμP·PrP° C, PSC-CfC, CH PrPsCfCfCfC, Pë P°P±Ps PīC-PrC, CfPëPjP° C, Pë PμCfPμPeC†C-Cф CfC, P° C, PμPIPsPiPs C†P»PμPSP°, PSPμPsP±C...C-PrPSC† PrP»CfC C†C†PīC-CëPSPsPiPs CfC, P° C, PμPIPsPiPs P°PeC, C†.

P" P»CfC PμC,, PμPeC, PëPIPSPsC— PrC-C— PīCfPμPīP°CfP° C, C† PīPsC, CфC-P±PSPμ C†PμPeC†C†P°P»CHPSPμ P·P±C†PrP¶PμPSPSCfC.

PμCfPsC, PëPīPsPeP°P·P°PSPSCfC.

- PμC-PrPIPëC%PμPSP° C†C†C, P»PëPIC-CfC, CH PrPs P°PeC, PëPIPSPsC— CфPμC†PsPIPëPSPë P°P±Ps PrPs P±C†PrCH-CfPePsC— P· PrPsPīPsPjC-P¶PSPëC... CфPμC†PsPIPëPS PīCfPμPīP°CfP° C, C†.
- PhPrPSPsC†P°CfPSPμ P·P°CfC, PsC†C†PIP°PSPSCfC C-P· PrPsPSPsCfP°PjPë PsPeC†PëPrC† P°P·PsC, C† (C, P°PePëPjPë CfPe P°PjC-P»PSC-C, CфPëC,) P°P±Ps PSC-C, CфP° C, P°PjPë C† P±C†PrCH-CfPeC-PNë C,, PsCfPjC- PīCfPsC, PëPīPsPeP°P·P°PSPμ, PsC†PeC-P»CHPePë PIC-PrPsPjPs, C%Ps C†PëP»PrPμPSP° C,, C-P» PjP° C" PIPīP»PëPI PSP° CëP»CfC...Pë PjPμC, P°P±PsP»C-P·PjC† NO/C†P"PhPα C, P° PīPsC, PμPSC†C-CфC" PīC-PīPsC, PμPSP·PëPIPSPëPNë PμC,, PμPeC, PSC-C, CфP° C, C-PI.
- PhPrPSPsC†P°CfPSPμ P·P°CfC, PsC†C†PIP°PSPSCfC C-PSPiC-P±C-C, PsCfC-PI PαP" P·5 (C† C, PsPjC† C†PëC†P»C- C†PëP»PrPμPSP° C,, C-P»C†) P·C- CfC, PëPjC†P»CfC, PsCfP°PjPë PīC†P°PSC-P»P° C, C†PëPeP»P°P·Pë, C, P°PePëPjPë CfPe CфC-PsC†PëPīC†P° C,, PīCfPsC, PëPīPsPeP°P·P°PSPμ, PsC†PeC-P»CHPePë PjPsP¶Pμ PīCfPëP·PIPμCfC, Pë PrPs C†PëPjPīC, PsPjP° C, PëC†PSPsC— PīC-PīPsC, PμPSP·C-C— (PrPëPI. CфPsP·PrC-P» B«P'P·P° C" PjPsPrC-CfC P· C-PSCëPëPjPë P»C-PeP°CfC†CHPePëPjPë P·P°C†PsP±P°PjPë C, P° C-PSCëC- PIPëPrPë PIP·P° C" PjPsPrC-PNëB»).
- PŸC, P°PSPë, PīCfPë CfPePëC... PSPμ CфPμPePsPjPμPSPrPsPIP°PSP° C†PμPeC†C†P°P»CHPSP° P°PeC, PëPIPSC-CfC, CH (PSP°PīCfPëPeP»P°Pr, C, CfP¶PeC- C†PμCfC†PμPIPs-C†C†PrPëPSPSC- CфPsP·P»P°PrPë, C, P°PeC- CfPe PSPμCfC, P°P±C-P»CHPSP° CfC, PμPSPsPeP°CfPrC-CfC P°P±Ps C†PμCfC†PμPIP° PSPμPrPsCfC, P° C, PSC-CfC, CH

C, CIP¶PePsPiPs CÍC, CíPíPµPSCÍ).

- P°C, CĤP°C, P° P·PsCĤCí PSP° PsPrPSPµ PsPePs PIPSP°CÍP»C-PrPsPe
PSPµP°CĤC, PµCĤC-P°P»CHBPSPsC— PíPµCĤPµPrPSCĤPsC— C-C€PµPjC-C‡PSPsC—
PSPµPICĤPsPíP°C, C-C— P·PsCĤPsPiPsPSPµCĤPIP°, PSPµP·P°P»Pµ¶PSPs PIC-Pr C, PsPiPs,
PíPsPiBĤ™ CIP·P°PSP° C†CÍ PíP°C, PsP»PsPiC-CÍ C-P· PíPsPíPµCĤPµPrPSC-Pj
P·P°CÍC, PsCÍCíPíP°PSPSCÍPj C-PSPiC-P±C-C, PsCĤC-PI PαP”P·5 C‡Pě PSC-.

- PkP°CÍPIPSC-CÍC, CH C, P°PePěC... P·P°C... PIPsCĤCĤPIP°PSCĤ: PíPsCĤCíC€PµPSPSCÍ
C,, CíPSPeC†C-C— PíPµC‡C-PSPePě C, CIP¶PePsPiPs CÍC, CíPíPµPSCÍ,
P°CĤC, PµCĤC-P°P»CHBPSP° PiC-PíPsC, PµPSP·C-CÍ (P°CĤC, PµCĤC-P°P»CHBPSPěPNē C, PěCÍPe
PSPěP¶C‡Pµ 90/50 PjPj CĤC,, CÍC,,), PSPµC%oPsPrP°PIPSPs PíPµCĤPµPSPµCÍPµPSPěPNē
C-PSCÍCíP»CHC, P°P±Ps C-PSC,, P°CĤPeC, PjC-PsPeP°CĤPrP° C, P° CÍPíP°PrPePsPIC-
PrPµPíPµPSPµCĤP°C, PěPIPSC- P·P°C... PIPsCĤCĤPIP°PSPSCÍ CÍC-C, PeC-PIPePě, C, P°PeC- CÍPe
PíC-PíPjPµPSC, PSPěPNē CĤPµC, PěPSC-C, (PSPµPIPµP»PěPeP° PeC-P»CHPeC-CÍC, CH
C, P°PePěC... PíP°C†C-C”PSC, C-PI PjP°C” PíPµPSPµC, PěC‡PSC- CĤPsP·P»P°PrPě
C,, PsCÍC,, PsPrC-PµCÍC, PµCĤP°P· CÍC-C, PeC-PIPePě), BĤ” PsCÍPeC-P»CHPePě P±PµP·PíPµPeP°
CÍPěP»PrPµPSP°C,, C-P»Cí PSPµ PrPsCÍP»C-PrP¶CíPíP°P»P°CÍCÍ Cí CĤP°P·C- C, P°PePěC...
P·P°C... PIPsCĤCĤPIP°PSCĤ.

**P’P·P°C”PjPsPrC-CÍ P· C-PSC€PěPjPě P»C-PeP°CĤCÍCHPePěPjPě P·P°CÍPsP±P°PjPě C, P°
C-PSC€C- PIPěPrPě PIP·P°C”PjPsPrC-PNē.**

P·C,, PµPeC, Pě C-PSC€PěC... P»C-PeP°CĤCÍCHPePěC... P·P°CÍPsP±C-PI PSP°
CÍPěP»PrPµPSP°C,, C-P»

P”PsCÍP»C-PrP¶PµPSPSCÍ in vitro

PĤPµC, P°P±PsP»C-P·Pj CÍPěP»PrPµPSP°C,, C-P»Cí PIC-PrP±CíPIP°C”C, CHCÍCÍ
PíPµCĤPµPIP°P¶PSPs P·P° CíC‡P°CÍC, CĤ C-P·PsC,, PsCĤPjPě 3PĤ4 (PíPsP»PsPIPSPěPNē
C€P»CÍC...) C, P° C-P·PsC,, PsCĤPjPě 2PŸ9 (PrCĤCíPíPsCĤCÍPrPSPěPNē C€P»CÍC...) C†PěC, PsC... CĤPsPjCí P 450 (CYP). PŸPsPjCí C-PSPiC-P±C-C, PsCĤPě C†PěC...
C-P·PsC,, PµCĤPjPµPSC, C-PI PjPsP¶CíC, CH P·PSPěP¶CíPIP°C, Pě PeP»C-CĤPµPSCÍ
CÍPěP»PrPµPSP°C,, C-P»Cí, P° C-PSPrCíPeC, PsCĤPě C†PěC... C-P·PsC,, PµCĤPjPµPSC, C-PI
PjPsP¶CíC, CH PíC-PrPIPěC%oCíPIP°C, Pě PeP»C-CĤPµPSCÍ CÍPěP»PrPµPSP°C,, C-P»Cí.

P”PsCÍP»C-PrP¶PµPSPSCÍ in vivo

PµPsPíCíP»CÍC†C-PNēPSPěPNē C,, P°CĤPjP°PePsPeC-PSPµC, PěC‡PSPěPNē P°PSP°P»C-P·
PrP°PSPěC... PeP»C-PSC-C‡PSPěC... PrPsCÍP»C-PrP¶PµPSCĤ
PíCĤPsPrPµPjPsPSCÍC, CĤCíPíP°PI P·PSPěP¶PµPSPSCÍ PeP»C-CĤPµPSCÍCí
CÍPěP»PrPµPSP°C,, C-P»Cí PíCĤPě PNēPsPiPs PsPrPSPsC‡P°CÍPSPsPjCí
P·P°CÍC, PsCÍCíPíP°PSPSC- P· C-PSPiC-P±C-C, PsCĤP°PjPě CYP3A4 (C, P°PePěPjPě CÍPe
PePµC, PsPePsPSP°P·PsP», PµCĤPěC, CĤPsPjC-C†PěPS, C†PěPjPµC, PěPrPěPS). PÍPsC‡P° PíCĤPě
PsPrPSPsC‡P°CÍPSPsPjCí P·P°CÍC, PsCÍCíPíP°PSPSC- CÍPěP»PrPµPSP°C,, C-P»Cí C, P°
C-PSPiC-P±C-C, PsCĤC-PI CYP3A4 P·CĤPsCÍC, P°PSPSCÍ C‡P°CÍC, PsC, Pě PíPsP±C-C‡PSPěC...
CÍPIPěC%o PSPµ CÍPíPsCÍC, PµCĤC-PíP°P»PsCÍCÍ, CÍP»C-Pr CĤPsP·PíP»CÍPSCíC, Pě
PjPsP¶P»PěPIC-CÍC, CH P·P°CÍC, PsCÍCíPíP°PSPSCÍ PíPsC‡P°C, PePsPiPsC— PrPsP·Pě
CÍPěP»PrPµPSP°C,, C-P»Cí 25 PjPi.

PĤPrPSPsC‡P°CÍPSPµ P·P°CÍC, PsCÍCíPíP°PSPSCÍ C-PSPiC-P±C-C, PsCĤP° P’P†P·-

$PiC_{\eta}PsC, P_{\mu}P^{\circ}P \cdot P \ddot{e} C_{\eta}P \ddot{e}C, PsPSP^{\circ}PIC-C_{\eta}C_{\eta}$, $PrC_{\eta}P \parallel P_{\mu} P \ddot{e}PsC, C_{\eta}P \parallel PSPsPiPs$
 $C-PSPiC-P \pm C-C, PsC_{\eta}P^{\circ} P 450$, $C_{\eta} C_{\eta}C, P^{\circ}PSC- C_{\eta}C-PIPSPsPIP^{\circ}P \parallel PSPsC-$
 $PePsPSC \dagger P_{\mu}PSC, C_{\eta}P^{\circ}C \dagger C-C-$ (500 $PjPi 2 C_{\eta}P^{\circ}P \cdot PSP^{\circ} PrPsP \pm C_{\eta}$) $C, P^{\circ} C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta}$
 $(PsPrPSPsC_{\eta}P^{\circ}P \cdot PsPIP^{\circ} PrPsP \cdot P^{\circ} 100 PjPi) PiC_{\eta}P \ddot{e}P \cdot PIPsPrP \ddot{e}P \gg Ps PrPs PiC-PrPIP \ddot{e}C \% P_{\mu}PSPSC \parallel$
 $C_{\max} C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} PSP^{\circ} 300 \% (C_{\eta} 4 C_{\eta}P^{\circ}P \cdot P \ddot{e}) C, P^{\circ} PiC-PrPIP \ddot{e}C \% P_{\mu}PSPSC \parallel$
 $PiP \gg P^{\circ}P \cdot PjPsPIP sC-$ AUC $C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} PSP^{\circ} 1000 \% (C_{\eta} 11 C_{\eta}P^{\circ}P \cdot C-PI)$.
 $PSP_{\mu}C_{\eta}P_{\mu}P \cdot 24 PiPsPrP \ddot{e}PSP \ddot{e} PiP \gg P^{\circ}P \cdot PjPsPIC- C_{\eta}C-PIPSC- C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} PIC_{\eta}P_{\mu}$
 $C \% P_{\mu} C_{\eta}C, P^{\circ}PSPsPIP \ddot{e}P \gg P \ddot{e} PiC_{\eta}P \ddot{e}P \pm P \gg P \ddot{e}P \cdot PSPs 200 PSPi/PjP \gg$, $PiPsC_{\eta}C-PIPSC \parallel PSPs P \cdot$
 $C_{\eta}C-PIPSP_{\mu}Pj PiC_{\eta}P \ddot{e}P \pm P \gg P \ddot{e}P \cdot PSPs 5 PSPi/PjP \gg$, $C...P^{\circ}C_{\eta}P^{\circ}PeC, P_{\mu}C_{\eta}PSP \ddot{e}Pj PrP \gg C \parallel$
 $P \cdot P^{\circ}C_{\eta}C, PsC_{\eta}C_{\eta}PIP^{\circ}PSPSC \parallel C, C-P \gg C_{\eta}PeP \ddot{e} C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta}$, $C \% Ps$
 $PIC-PrPiPsPIC-PrP^{\circ}C" P \cdot PSP^{\circ}C \dagger PSPsPjC_{\eta} PIPiP \gg P \ddot{e}PIC_{\eta} C_{\eta}P \ddot{e}C, PsPSP^{\circ}PIC-C_{\eta}C_{\eta} PSP^{\circ}$
 $C \ddot{e}P \ddot{e}C_{\eta}PsPeP \ddot{e}PN_{\eta} C_{\eta}PiP_{\mu}PeC, C_{\eta} C_{\eta}C_{\eta}P \pm C_{\eta}C, C_{\eta}P^{\circ}C, C-PI P 450$. $P \ddot{e}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg PSP_{\mu}$
 $PIPiP \gg P \ddot{e}PIP^{\circ}C" PSP^{\circ} C, P^{\circ}C_{\eta}PjP^{\circ}PePsPeC-PSP_{\mu}C, P \ddot{e}PeC_{\eta} C_{\eta}P \ddot{e}C, PsPSP^{\circ}PIC-C_{\eta}C_{\eta}$. $P-$
 $PsPiP \gg C \parallel PrC_{\eta} PSP^{\circ} C \dagger C- C, P^{\circ}C_{\eta}PjP^{\circ}PePsPeC-PSP_{\mu}C, P \ddot{e}C \dagger PSC- PrP^{\circ}PSC- PsPrPSPsC \dagger P^{\circ}C_{\eta}PSP_{\mu}$
 $P \cdot P^{\circ}C_{\eta}C, PsC_{\eta}C_{\eta}PIP^{\circ}PSPSC \parallel C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} C, P^{\circ} C_{\eta}P \ddot{e}C, PsPSP^{\circ}PIC-C_{\eta}C_{\eta} PSP_{\mu}$
 $C_{\eta}P_{\mu}PePsPjP_{\mu}PSPPrC_{\eta}C" C, C_{\eta}C_{\eta}C \parallel (PrP \ddot{e}PI. C_{\eta}PsP \cdot PrC-P \gg B \langle PhC_{\eta}PsP \pm P \gg P \ddot{e}IPsC_{\eta}C, C-$
 $P \cdot P^{\circ}C_{\eta}C, PsC_{\eta}C_{\eta}PIP^{\circ}PSPSC \parallel B \rangle$; $C_{\eta} PePsP \parallel PSPsPjC_{\eta} C_{\eta}P^{\circ}P \cdot C- PjP^{\circ}PeC_{\eta}P \ddot{e}PjP^{\circ}P \gg C_{\eta}PSP^{\circ}$
 $PrPsP \cdot P^{\circ} C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} PSP_{\mu} PiPsPIP \ddot{e}PSPSP^{\circ} PiP_{\mu}C_{\eta}P_{\mu}PIP \ddot{e}C \% C_{\eta}PIP^{\circ}C, P \ddot{e} 25 PjPi$
 $PiC_{\eta}PsC, C \parallel PiPsPj 48 PiPsPrP \ddot{e}PS$.

$PhPrPSPsC \dagger P^{\circ}C_{\eta}PSP_{\mu} P \cdot P^{\circ}C_{\eta}C, PsC_{\eta}C_{\eta}PIP^{\circ}PSPSC \parallel C-PSPiC-P \pm C-C, PsC_{\eta}P^{\circ} P' \dagger P \gg$
 $PiC_{\eta}PsC, P_{\mu}P^{\circ}P \cdot P \ddot{e} C_{\eta}P^{\circ}PePIC-PSP^{\circ}PIC-C_{\eta}C_{\eta}$, $C-PSPiC-P \pm C-C, PsC_{\eta}P^{\circ} CYP3A4$, $C_{\eta} PrPsP \cdot C-$,
 $C \% Ps P \cdot P^{\circ}P \pm P_{\mu}P \cdot PiP_{\mu}C \dagger C_{\eta}C" C_{\eta}C-PIPSPsPIP^{\circ}P \parallel PSC_{\eta} PePsPSC \dagger P_{\mu}PSC, C_{\eta}P^{\circ}C \dagger C-C_{\eta} (1200 PjPi 3$
 $C_{\eta}P^{\circ}P \cdot P \ddot{e} PSP^{\circ} PrPsP \pm C_{\eta})$, $C, P^{\circ} C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} (100 PjPi PsPrPSPsC_{\eta}P^{\circ}P \cdot PsPIP s)$
 $PiC_{\eta}P \ddot{e}P \cdot PIPsPrP \ddot{e}P \gg Ps PrPs PiC-PrPIP \ddot{e}C \% P_{\mu}PSPSC \parallel C_{\max} C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} PSP^{\circ}$
 $140 \% C, P^{\circ} P \cdot P \pm C-P \gg C_{\eta}C \ddot{e}P_{\mu}PSPSC \parallel AUC C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} PSP^{\circ} 210 \%$. PkP_{μ}
 $PIP \ddot{e}C \parallel PIP \gg P_{\mu}PSPs PIPiP \gg P \ddot{e}PIC_{\eta} C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} PSP^{\circ}$
 $C, P^{\circ}C_{\eta}PjP^{\circ}PePsPeC-PSP_{\mu}C, P \ddot{e}PeC_{\eta} C_{\eta}P^{\circ}PePIC-PSP^{\circ}PIC-C_{\eta}C_{\eta} (PrP \ddot{e}PI. C_{\eta}PsP \cdot PrC-P \gg$
 $B \langle P \ddot{e}PiPsC_{\eta}C-P \pm P \cdot P^{\circ}C_{\eta}C, PsC_{\eta}C_{\eta}PIP^{\circ}PSPSC \parallel C, P^{\circ} PrPsP \cdot P \ddot{e}B \rangle$).
 $P_{\mu}P_{\mu}C_{\eta}P_{\mu}PrP \pm P^{\circ}C \dagger P^{\circ}C" C, C_{\eta}C_{\eta}C \parallel, C \% Ps P \pm C-P \gg C_{\eta}C \ddot{e} PiPsC, C_{\eta}P \parallel PSC-$
 $C-PSPiC-P \pm C-C, PsC_{\eta}P \ddot{e} CYP3A4$, $C, P^{\circ}PeC- C \parallel Pe PeP_{\mu}C, PsPePsPSP^{\circ}P \cdot PsP \gg C, P^{\circ}$
 $C-C, C_{\eta}P^{\circ}PePsPSP^{\circ}P \cdot PsP \gg$, $P \pm C_{\eta}PrC_{\eta}C, C_{\eta} PjP^{\circ}C, P \ddot{e} P \pm C-P \gg C_{\eta}C \ddot{e} PIP \ddot{e}C_{\eta}P^{\circ}P \parallel P_{\mu}PSP \ddot{e}PN_{\eta}$
 $PIPiP \gg P \ddot{e}PI$.

$P_{\mu}C_{\eta}P \ddot{e} P \cdot P^{\circ}C_{\eta}C, PsC_{\eta}C_{\eta}PIP^{\circ}PSPSC- C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} (100 PjPi PsPrPSPsC_{\eta}P^{\circ}P \cdot PsPIP s)$
 $C, P^{\circ} P_{\mu}C_{\eta}P \ddot{e}C, C_{\eta}PsPjC-C \dagger P \ddot{e}PSC_{\eta}$, $PiPsPjC-C_{\eta}PSPsPiPs C-PSPiC-P \pm C-C, PsC_{\eta}P^{\circ} CYP3A4$, C_{η}
 $C_{\eta}C-PIPSPsPIP^{\circ}P \parallel PSPsPjC_{\eta} C_{\eta}C, P^{\circ}PSC- (500 PjPi 2 C_{\eta}P^{\circ}P \cdot P \ddot{e} PSP^{\circ} PrPsP \pm C_{\eta} PiC_{\eta}PsC, C \parallel PiPsPj$
 $5 PrPSC-PI) C_{\eta}PiPsC_{\eta}C, P_{\mu}C_{\eta}C-PiP^{\circ}P \gg PsC_{\eta}C \parallel PiC-PrPIP \ddot{e}C \% P_{\mu}PSPSC \parallel P_{\eta}UC$
 $C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} PSP^{\circ} 182 \%$. $Pj P \cdot PrPsC_{\eta}PsPIP \ddot{e}C... PrPsP \pm C_{\eta}PsPIP sP \gg C_{\eta}C \dagger C-PI$
 $C \dagger PsP \gg PsPIC-C \dagger PsC- C_{\eta}C, P^{\circ}C, C- PSP_{\mu} C_{\eta}PiPsC_{\eta}C, P_{\mu}C_{\eta}C-PiP^{\circ}P \gg PsC_{\eta}C \parallel PIPiP \gg P \ddot{e}PIC_{\eta}$
 $P^{\circ}P \cdot P \ddot{e}C, C_{\eta}PsPjC-C \dagger P \ddot{e}PSC_{\eta} (500 PjPi PSP^{\circ} PrPsP \pm C_{\eta} PiC_{\eta}PsC, C \parallel PiPsPj 3 PrC-P \pm) PSP^{\circ} AUC$,
 $P \ddot{e}_{\max}, T_{\max}, PePsPSC_{\eta}C, P^{\circ}PSC, C_{\eta} C \ddot{e}PIP \ddot{e}PrPePsC_{\eta}C, C- P_{\mu}P \gg C-PjC-PSPP^{\circ}C \dagger C-C- C, P^{\circ}$
 $PiPsPrP^{\circ}P \gg C_{\eta}C \ddot{e}P \ddot{e}PN_{\eta} PiP_{\mu}C_{\eta}C-PsPr PSP^{\circ}PiC-PIPiP \ddot{e}PIPiPrP_{\mu}PSPSC \parallel$
 $C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} P^{\circ}P \pm Ps PN_{\eta}PsPiPs PiPsP \gg PsPIPSPsPiPs C \dagger P \ddot{e}C_{\eta}PeC_{\eta}P \gg C_{\eta}C_{\eta}C \dagger PsPiPs$
 $PjP_{\mu}C, P^{\circ}P \pm PsP \gg C-C, C_{\eta}$. $PiP \ddot{e}PjP_{\mu}C, P \ddot{e}PrP \ddot{e}PS (C-PSPiC-P \pm C-C, PsC_{\eta} C \dagger P \ddot{e}C, PsC...C_{\eta}PsPjC_{\eta} P 450$
 $C, P^{\circ} PSP_{\mu}C_{\eta}PiP_{\mu}C \dagger P \ddot{e}C, C-C \dagger PSP \ddot{e}PN_{\eta} C-PSPiC-P \pm C-C, PsC_{\eta} CYP3A4) C_{\eta} PrPsP \cdot C- 800 PjPi PiC_{\eta}P \ddot{e}$
 $PsPrPSPsC \dagger P^{\circ}C_{\eta}PSPsPjC_{\eta} P \cdot P^{\circ}C_{\eta}C, PsC_{\eta}C_{\eta}PIP^{\circ}PSPSC- P \cdot C- C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg PsPj C_{\eta}$
 $PrPsP \cdot C- 50 PjPi C_{\eta} P \cdot PrPsC_{\eta}PsPIP \ddot{e}C... PrPsP \pm C_{\eta}PsPIP sP \gg C_{\eta}C \dagger C-PI PiC_{\eta}P \ddot{e}P \cdot PIPsPrP \ddot{e}PI PrPs$
 $PiC-PrPIP \ddot{e}C \% P_{\mu}PSPSC \parallel PiP \gg P^{\circ}P \cdot PjPsPIP \ddot{e}C... PePsPSC \dagger P_{\mu}PSC, C_{\eta}P^{\circ}C \dagger C-PN_{\eta}$
 $C_{\eta}P \ddot{e}P \gg PrP_{\mu}PSP^{\circ}C, C-P \gg C_{\eta} PSP^{\circ} 56 \%$.

PjPePjPsP»CH, PIPiP»PëPI PíCḡPμPīP°CḡP°C,Cr PSP° PeP»C-CḡPμPSCÍ CÍCrP±CÍC,CḡP°C,C-PI C†PëC... C-P·PsC,,PμCḡPjPμPSC,C-PI PjP°P»PsPNëPjPsPIC-CḡPSPëPNë.

P'C-PrCÍCrC,PSC- PrP°PSC- C%PsPrPs PIP·P°C"PjPsPrC-C— CÍPëP»PrPμPSP°C,,C-P»Cr C- C,P°PePëC... PSPμCÍPīPμC†PëC,,C-C†PSPëC... C-PSPiC-P±C-C,PSCḡC-PI C,,PsCÍC,,PsPrC-PμCÍC,PμCḡP°P·Pë, C‡Pe C,PμPsC,,C-P»C-PS C,P° PrPëPiC-CḡPëPrP°PjPsP».

P"PsCÍP»C-PrP¶PμPSPSC‡ in vivo

PhCÍPeC-P»CHPePëB B PIC-PrPsPjPs,B B B C%PsB B CÍPëP»PrPμPSP°C,,C-P»B B PjP°C"B B PIPiP»PëPI B B PSP° B B PjPμC,P°P±PsP»C-P·Pj B NO/C†P"PhPα, P±CÍP»Ps PICÍC,P°PSPsPIP»PμPSPs, C%Ps CÍPëP»PrPμPSP°C,,C-P» PiPsC,PμPSC†C-CḡC" PiC-PiPsC,PμPSP·PëPIPSCr PrC-Cḡ PSC-C,CḡP°C,C-PI, C,PSPjCr PNëPsPiPs PsPrPSPsC†P°CÍPSPμ P·P°CÍC,PSCÍCrPIP°PSPSC‡ P· PrPsPSPsCḡP°PjPë PsPeCÍPëPrCr P°P·PsC,Cr P°P±Ps P· PSC-C,CḡP°C,P°PjPë Cr P±CÍPrCH-C‡PeC-PNë C,,PsCḡPjC- PíCḡPsC,PëPiPsPeP°P·P°PSPμ (PrPëPI. CḡPsP·PrC-P» B«PμCḡPsC,PëPiPsPeP°P·P°PSPSC‡B»).

P C-PsC†PëPiC¿P°C,. P"PsPeP»C-PSC-C†PSC- PrPsCÍP»C-PrP¶PμPSPSC‡ PíCḡPsPrPμPjPsPSCÍC,CḡCrPIP°P»Pë P°PrPëC,PëPIPSPëPNë CÍPëCÍC,PμPjPSPëPNë PμC,,PμPeC, P·PSPëP¶PμPSPSC‡ P°CḡC,PμCḡC-P°P»CHPSPsPiPs C,PëCÍPeCr PíCḡPë PsPrPSPsC†P°CÍPSPsPjCr P·P°CÍC,PSCÍCrPIP°PSPSC- C-PSPiC-P±C-C,PSCḡC-PI PαP"P·5 P· CḡC-PsC†PëPiCrP°C,PSPj. PḡP»C-PSC-C†PSC- PrPsCÍP»C-PrP¶PμPSPSC‡ PíCḡPsPrPμPjPsPSCÍC,CḡCrPIP°P»Pë, C%Ps CḡC-PsC†PëPiCrP°C, PiPsCÍPëP»CḡC" PiC-PiPsC,PμPSP·PëPIPSCr PrC-Cḡ C-PSPiC-P±C-C,PSCḡC-PI PαP"P·5.

PJ PīP°C†C-C"PSC,C-PI, C‡PeC- P±CḡP°P»Pë CrC†P°CÍC,CH Cr PrPsCÍP»C-PrP¶PμPSPSC-, PSPμ CÍPiPsCÍC,PμCḡC-PiP°P»PsCÍCH PiPsP·PëC,PëPIPSPsPiPs PeP»C-PSC-C†PSPsPiPs PμC,,PμPeC,Cr PIC-Pr PsPrPSPsC†P°CÍPSPsPiPs P·P°CÍC,PSCÍCrPIP°PSPSC‡ C-PSPiC-P±C-C,PSCḡC-PI PαP"P·5 P· CḡC-PsC†PëPiCrP°C,PSPj. PμCḡPsC,PëPiPsPeP°P·P°PSPμ PsPrPSPsC†P°CÍPSPμ P·P°CÍC,PSCÍCrPIP°PSPSC‡ CḡC-PsC†PëPiCrP°C,Cr P· C-PSPiC-P±C-C,PSCḡP°PjPë PαP"P·5 (Cr C,PSPjCr C†PëCÍP»C- P·C- CÍPëP»PrPμPSP°C,,C-P»PsPj) (PrPëPI. CḡPsP·PrC-P»B B«PμCḡPsC,PëPiPsPeP°P·P°PSPSC‡B»).

PhPrPSPsC†P°CÍPSPμ P·P°CÍC,PSCÍCrPIP°PSPSC‡ CÍPëP»PrPμPSP°C,,C-P»Cr C,P° P±P»PsPeP°C,PSCḡC-PI α-P°PrCḡPμPSPsCḡPμC†PμPīC,PSCḡC-PI PjPsP¶Pμ PíCḡPëP·PIPμCÍC,Pë PrPs CḡPsP·PIPëC,PeCr CÍPëPjPiC,PSPjP°C,PëC†PSPsC— PiC-PiPsC,PμPSP·C-C— Cr PrPμC‡PePëC... CÍC...PëP»CHPSPëC... PrPs C†CHPsPiPs PīP°C†C-C"PSC,C-PI. PŷP°PeP° CḡPμP°PeC†C-C‡ PSP°PNëC†P°CÍC,C-C€Pμ PIPëPSPëPeP°P»P° PíCḡPsC,C‡PiPsPj 4 PiPsPrPëPS PíC-CÍP»C‡ P·P°CÍC,PSCÍCrPIP°PSPSC‡ CÍPëP»PrPμPSP°C,,C-P»Cr (PrPëPI. CḡPsP·PrC-P»Pë B«PhCÍPsP±P»PëPIPsCÍC,C- P·P°CÍC,PSCÍCrPIP°PSPSC‡B» C,P° B«PŷPiPsCÍC-P± P·P°CÍC,PSCÍCrPIP°PSPSC‡ C,P° PrPsP·PëB»). PJ PíCḡPsC†PμCÍC- 3 PrPsCÍP»C-PrP¶PμPSCḡ CÍPiPμC†PëC,,C-C†PSPsC— PIP·P°C"PjPsPrC-C— P»C-PeP°CḡCÍCHPePëC... P·P°CÍPsP±C-PI P±P»PsPeP°C,PSCḡ α-P°PrCḡPμPSPsCḡPμC†PμPīC,PSCḡC-PI PrPsPeCÍP°P·PsP·PëPS (4 PjPi C,P° 8 PjPi) C,P° CÍPëP»PrPμPSP°C,,C-P» (25 PjPi, 50 PjPi C,P° 100 PjPi) P·P°CÍC,PSCÍPsPICrPIP°P»Pë PsPrPSPsC†P°CÍPSPs PīP°C†C-C"PSC,P°Pj C-P· PrPsP±CḡPsC‡PeC-CÍPSPsCḡ PiC-PiPμCḡPīP»P°P·C-C"ḡ PīPμCḡPμPrPjC-C...CrCḡPsPIPSC— P·P°P»PsP·Pë, CÍC,P°P±C-P»C-P·P°C†C-C‡ CÍC,P°PSCr C‡PePëC... P±CÍP»P° PrPsCÍC‡PiPSCrC,P° PíCḡPë P·P°CÍC,PSCÍCrPIP°PSPSC- PrPsPeCÍP°P·PsP·PëPSCr. PJ C†PëC... PiPsPiCrP»C‡C†C-C‡C... CÍPiPsCÍC,PμCḡC-PiP°P»PsCÍC‡ CÍPμCḡPμPrPSC" PrPsPrP°C,PePsPIPμ P·PSPëP¶PμPSPSC‡ P°CḡC,PμCḡC-P°P»CHPSPsPiPs C,PëCÍPeCr Cr PiPsP»PsP¶PμPSPSC- P»PμP¶P°C†Pë PSP° 7/7 PjPj CḡC,. CÍC,.. 9/5 PjPj CḡC,. CÍC,. C,P° 8/4 PjPj CḡC,. CÍC,. C,P° CÍPμCḡPμPrPSC" P·PSPëP¶PμPSPSC‡ P°CḡC,PμCḡC-P°P»CHPSPsPiPs C,PëCÍPeCr Cr PiPsP»PsP¶PμPSPSC-

СГГ,PsCЦC±Pë PSP° 6/6 PjPj CГC,, СГГ,, 11/4 PjPj CГC,, СГГ,, 4/5 PjPj CГC,, СГГ,,
PIC-PrPіPsPIC-PrPSPs. PμCГPë PsPrPSPsC±P°CГPSPsPjCр P·P°CГC,PsCГCрPIP°PSPSC-
CГPëP»PrPμPSP°C,,C-P»Cр C,P° PrPsPeCГP°P·PsP·PëPSCр PіP°CГC-C”PSC,P°Pj,
CГC,P°P±C-P»C-P·P°CГC-CЦ CГC,P°PSCр CЦPePëC... P±CрP»P° PrPsCГCЦPіPSCрC,P° PіCГPë
P·P°CГC,PsCГCрPIP°PSPSC- PrPsPeCГP°P·PsP·PëPSCр, C-PSPsPrC-
PіPsPIC-PrPsPjP»CЦP»PsCГCЦ PіCГPs CГPsP·PIPëC,PsPe CГPëPjPіC,PsPjP°C,PëC±PSPsC—
PsCГC,PsCГC,P°C,PëC±PSPsC— PіC-PіPsC,PμPSP·C-C—. PJ CГPëC...
PіPsPIC-PrPsPjP»PμPSPSCЦC... P№CєP»PsCГCЦ PіCГPs PIPëPіP°PrPePë
P·P°PіP°PjPsCГPsC±PμPSPSCЦ C,P° CГC,P°PS PіPμCГPμPr PSPμPіCГPëC,PsPjPSC-CГC,CГ,
P°P»Pμ P±PμP· CГPëPSPePsPіPμ.

PкPμ CГPіPsCГC,PμCГC-PіP°P»PsCГCЦ P¶PsPrPSPëC... P·PSP°C±CрC%PëC...
PIP·P°C”PjPsPrC-P№ PіCГPë PsPrPSPsC±P°CГPSPsPjCр P·P°CГC,PsCГCрPIP°PSPSC-
CГPëP»PrPμPSP°C,,C-P»Cр (50 PjPi) C- C,PsP»P±CрC,P°PjC-PrCр (250 PjPi) P°P±Ps
PIP°CГC,,P°CГPëPSCр (40 PjPi), C%Ps PjPμC,P°P±PsP»C-P·CрCГC,CГCГCЦ CYP2C9.

PŸPëP»PrPμPSP°C,,C-P» (50 PjPi) PSPμ PіCГPëP·PIPsPrPëPI PrPs PіPsPrPsPIP¶PμPSPSCЦ
C±P°CГCр PeCГPsPIPsC,PμC±C-, CГPіCГPëC±PëPSPμPSPsC— P·P°CГC,PsCГCрPIP°PSPSCЦPj
P°CГPμC,PëP»CГP°P»C-CГPëP»PsPIPsC— PePëCГP»PsC,Pë (150 PjPi).

PŸPëP»PrPμPSP°C,,C-P» (50 PjPi) PSPμ PіPsC,PμPSCГC-CГPIP°PI PіC-PіPsC,PμPSP·PëPIPsCр
PrC-CГ P°P»PePsPіPsP»CГ Cр P·PrPsCГPsPIPëC... PrPsP±CГPsPIPsP»CГCГC-PI PіCГPë
CГPμCГPμPrPSCГPsPjCр PjP°PeCГPëPjP°P»CГPSPsPjCр CГC-PIPsC- PμC,P°PSPsP»Cр
PeCГPsPIC- 80 PjPi/PrP».

PJ PіP°CГC-C”PSC,C-PI, CЦPeC- P·P°CГC,PsCГPsPICрPIP°P»Pë CГPëP»PrPμPSP°C,,C-P», PSPμ
CГPіPsCГC,PμCГC-PіP°P»PsCГCЦ P¶PsPrPSPëC... PIC-PrPjC-PSPSPsCГC,PμP№ PіCГPsC,,C-P»CГ
PіPsP±C-C±PSPëC... PμC,,PμPeC,C-PI PіPsCГC-PIPsCЦPSPs P· PіP»P°CГPμP±Ps PіCГPë
PsPrPSPsC±P°CГPSPsPjCр P·P°CГC,PsCГCрPIP°PSPSC- C,P°PePëC... PeP»P°CГC-PI
PіC-PіPsC,PμPSP·PëPIPsPëC... P»C-PeP°CГCГCГPePëC... P·P°CГPsP±C-PI, CЦPe
PrC-CрCГPμC,PëPePë, P±P»PsPeP°C,PsCГPë β-P°PrCГPμPSPsCГPμCГPμPіC,PsCГC-PI,
C-PSPіC-P±C-C,PsCГPë PђPұPα, P°PSC,P°PіPsPSC-CГC,Pë P°PSPіC-PsC,PμPSP·PëPSCр PГPГ,
P°PSC,PëPіC-PіPμCГC,PμPSP·PëPIPsC- P»C-PeP°CГCГCГPeC- P·P°CГPsP±Pë
(CГCрPrPëPSPsCГPsP·CєPëCГCГPIP°P»CГPSC- C,P° CГPμPSC,CГP°P»CГPSPsC— PrC-C—),
P±P»PsPeP°C,PsCГPë P°PrCГPμPSPμCГPіC-C±PSPëC... PSPμP№CГPsPSC-PI,
P±P»PsPeP°C,PsCГPë PeP°P»CГCГC-C”PIPëC... PeP°PSP°P»C-PI C,P° P±P»PsPeP°C,PsCГPë α-
P°PrCГPμPSPsCГPμCГPμPіC,PsCГC-PI.

PJ CГPіPμCГC-P°P»CГPSPsPjCр PrPsCГP»C-PrP¶PμPSPSC- PIP·P°C”PjPsPrC-C— PіCГPë
PsPrPSPsC±P°CГPSPsPjCр P·P°CГC,PsCГCрPIP°PSPSC- CГPëP»PrPμPSP°C,,C-P»Cр (100 PjPi) C,P°
P°PjP»PsPrPëPіC-PSCр PіP°CГC-C”PSC,P°Pj C-P· P°CГC,PμCГC-P°P»CГPSPsCГ
PіC-PіPμCГC,PμPSP·C-C”CГ CГPіPsCГC,PμCГC-PіP°P»PsCГCЦ PrPsPrP°C,PePsPIPμ
P·PSPëP¶PμPSPSCЦ CГPëCГC,PsP»C-C±PSPsPіPs P°CГC,PμCГC-P°P»CГPSPsPіPs C,PëCГPeCр PI
PіPsP»PsP¶PμPSPSC- P»PμP¶P°C±Pë PSP° 8 PjPj CГC,, CГC,. P’C-PrPіPsPIC-PrPSPμ
P·PSPëP¶PμPSPSCЦ PrC-P°CГC,PsP»C-C±PSPsPіPs P°CГC,PμCГC-P°P»CГPSPsPіPs C,PëCГPeCр
CГC,P°PSPsPIPëP»Ps 7 PjPj CГC,, CГC,. PіC- PrPsPrP°C,PePsPIC- P·PSPëP¶PμPSPSCЦ
P°CГC,PμCГC-P°P»CГPSPsPіPs C,PëCГPeCр P±CрP»Pë PіPsCГC-PIPsCЦPSPSPëPjPë P·
C,P°PePëPjPë, C%Ps CГPіPsCГC,PμCГC-PіP°P»PëCГCЦ PіCГPë P·P°CГC,PsCГCрPIP°PSPSC-
P»PëCєPμ CГPëP»PrPμPSP°C,,C-P»Cр Cр P·PrPsCГPsPIPëC... PrPsP±CГPsPIPsP»CГCГC-PI.

PŸPëP»PrPμPSP°C,,C-P» Cр PrPsP·C- 100 PjPi PSPμ PIPіP»PëPIP°PI PSP°
C,,P°CГPjP°PePsPeC-PSPμC,PëC±PSC- PіPsPeP°P·PSPëPePë C-PSPіC-P±C-C,PsCГC-PI P’PГP>-

РісЧіРсС,РіР°Р·Рө, СГР°РөРіС-РSP°РіС-СГіСр С,Р° СГіРөС,РсРSP°РіС-СГіСр, СіРөС- С"
СГіСрР±СГіС,СГіР°С,Р°РјРө СYP3A4.

PJ P·PrPsCГіРсРіРөС... PrPsP±СГіРсРіРсР»СНС†С-Рі С†РсР»РсРіС-С†РсС— СГіС,Р°С,С-
Р·Р°СГіС,РсСГіСрРіР°РSPSCі СГіРөР»PrРіРSP°С,,С-Р»Ср Ср СГіС-РіРSPсРіР°Р¶РSPсРјСр
СГіС,Р°РSC- (80 РјРі 3 СГіР°Р·Рө РSP° PrPsP±Ср) РіСГіРөР·РіРсРрРөР»Рс РrPs
РіС-РrРіРөС%РіРSPSCі AUC С,Р° C_{max} Р±РсСГіРіРSC,Р°РSCр (125 РјРі 2 СГіР°Р·Рө РSP°
PrPsP±Ср) РSP° 49,8 % С,Р° 42 % РіС-РrРіРсРіС-РrРSPс.

Р"РсРrР°РіР°РSPSCі РсРrРSPсСГіР°Р·РсРіРсС— РrPsP·Рө СГіРөР»PrРіРSP°С,,С-Р»Ср РrPs
СГіР°РөСрР±С-С,СГіРөР»Ср / РіР°Р»СГіР°СГіС,Р°РSCр Ср СГіС-РіРSPсРіР°Р¶РSPсРјСр СГіС,Р°РSC-
Ср РіР°С†С-С"РSC,С-Рі Р· Р°СГіС,РіСГіС-Р°Р»СНРSPсСРі РіС-РіРіСГіС,РіРSP·С-С"СР Р±СрР»Рс
РіРсРіСіР·Р°Р·Р°Р·Рі Р·С- Р·РSP°С†РSPс Р±С-Р»СНС€ РіРөСГіР°Р¶РіРSPөРј Р·РSPөРіРіРSPSCіРј
Р°СГіС,РіСГіС-Р°Р»СНРSPсРіРс С,РөСГіРөСр РіРсСГіС-РіРSCіРSPс Р· РіСГіРөРіРсРјРсРјРсРј
С,С-Р»СНРөРө СГіР°РөСрР±С-С,СГіРөР»Ср/РіР°Р»СГіР°СГіС,Р°РSCр. РйР°РөРөРј С†РөРSPсРј,
СГіР»С-Рr РrPsС,СГіРөРјСрРіР°Р°С,РөСГіСі РсР±РіСГіРіРіРSPсСГіС,С- РіСГіРө РіРсС†Р°С,РөСр
Р·Р°СГіС,РсСГіСрРіР°РSPSCі СГіРөР»PrРіРSP°С,,С-Р»Ср Ср РіР°С†С-С"РSC,С-Рі, СіРөС-
РсС,СГіРөРјСрСРіС,СН Р»С-РөСрРіР°РSPSCі
СГіР°РөСрР±С-С,СГіРөР»РсРј/РіР°Р»СГіР°СГіС,Р°РSPсРј.

РіСГіРсР±Р»РөРіРсСГіС,С- Р·Р°СГіС,РсСГіСіРіР°РSPSCі.

Р"Рс РіРсС†Р°С,РөСр С,РіСГіР°РіС-С— СГіР»С-Рr Р·С-Р±СГіР°С,Рө РјРіРrРөС†РSPөРіР
Р°РSP°РјРSPіР· РіР°С†С-С"РSC,Р° С,Р° РіСГіРсРіРіСГіС,Рө С,,С-Р·РөРөР°Р»СНРSPі
РсР±СГіС,РіРіРіРSPSCі РrР»Сі РrС-Р°РіРSPсСГіС,РөРөРө РіСГіРіРөС,РөР»СНРSPсС—
РrРөСГіС,,СрРSPөС†С-С— С,Р° РіРөР·РSP°С†РіРSPSCі С—С— РјРсРіР»РөРіРөС...
РіСГіРөС†РөРс.

*Р»Р°РөС,РсСГіРө СГіРөР·РөРөСі СГіРіСГіРіРіРіРс-СГіСіРrРөРSPSPөС...
Р·Р°С...РіРсСГіСРіР°РSCі*

РіСГіРөС-Р»СНРөРө СГіРіРөСГіСрР°Р»СНРSP° Р°РөС,РөРіРSC-СГіС,СН
СГіСрРіСГіРсРіРсРrРіСГіС,СНСГіСі РіРіРіРSPөРј СГіРөР·РөРөРсРј Р· Р±РсРөСр СГіРіСГіС†Сі,
РrPs РіРсС†Р°С,РөСр Р±СрРrСН-СіРөРсРіРс Р»С-РөСрРіР°РSPSCі РіСГіРіРөС,РөР»СНРSPсС—
РrРөСГіС,,СрРSPөС†С-С— Р»С-РөР°СГі РјР°С" РсС†С-РSPөС,Рө СГіС,Р°РSP СГіРіСГіС†РіРіРс-
СГіСрРrРөРSPSPсС— СГіРөСГіС,РіРјРө РіР°С†С-С"РSC,Р°. РйРөР»PrРіРSP°С,,С-Р»
С†РөРSPөС,СН СГіСрРrРөРSPсСГіРсР·СРөРөСГіРіР°Р»Р»СНРSCр РrС-СР, С%Рс
РіСГіРсСіРіР»СіС"С,СНСГіСі Р»РіРіРөРөРј С,Р° РөРсСГіРсС,РөРсС†Р°СГіРSPөРј
Р·РSPөРіРіРSPSCіРј Р°СГіС,РіСГіС-Р°Р»СНРSPсРіРс С,РөСГіРөСр. Р"Рс
РіСГіРөР·РSP°С†РіРSPSCі СГіРөР»PrРіРSP°С,,С-Р»Ср Р»С-РөР°СГі РјР°С" СГіРіС,РіР»СНРSPс
Р·РіР°РіРөС,Рө, С†Рө РјРсРіРі С,Р°РөРөРіРі РіС,,РіРөС, РSPіСГіРіРөСіС,Р»РөРіРс
РіРіР»РөРіР°С,Рө РSP° РіР°С†С-С"РSC,С-Рі С-Р· РіРіРіРSPөРј РсСГіРSPсРіРSPөРјРө
Р·Р°С...РіРсСГіСРіР°РSPSCіРө, РсСГіРсР±Р»РөРіРс Ср РөРсРјР±С-РSP°С†С-С— С-Р·
СГіРіРөСГіСрР°Р»СНРSPсСРі Р°РөС,РөРіРSC-СГіС,СР. Р"Рс РіР°С†С-С"РSC,С-Рі С-Р·
РіС-РrРіРөС%РіРSPсСРі С†СрС,Р»РөРіС-СГіС,СР РrPs РіР°Р·РсРrРөР»СіС,Р°С,РсСГіС-Рі
РSP°Р»РіРіР°С,СН РіР°С†С-С"РSC,Рө С-Р· РсР±СГіС,СГіСрРөС†С-С"СР РіРөРіС-РrРSPсРіРс
С,СГіР°РөС,Ср Р»С-РіРсРіРс СРөР»СрРSPсС†РөР° (РSP°РіСГіРөРөР»Р°Рr, СГіС,РіРSPсР·
Р°РсСГіС,Рө, РіС-РіРіСГіС,СГіРс,,С-С†РSP° РсР±СГіС,СГіСрРөС,РөРіРSP°
РөР°СГіРrС-РсРјС-РсРіР°С,С-Сі) С,Р° РіР°С†С-С"РSC,Рө С-Р· СГіС-РrРөС-СГіРSPөРј
СГіРөРSPrСГіРсРјРсРј РјСрР»СНС,РөСГіРөСГіС,РіРјРSPсС— Р°С,СГіРс,,С-С—, РсРrРSPөРј С-Р·

PhPrPSPsC†P°C†PSPμ P·P°C†C,PsC†C¿PIP°PSPSC‡ P· C-PSC€PëPjPë C-PSPiC-P±C-C,PsC‡P°PjPë
PαP°P·5 P°P±Ps C-PSC€PëPjPë PiC‡PμPiP°C‡P°C,P°PjPë PrP»C‡ P»C-PeC¿PIP°PSPSC‡
PμC‡PμPeC,PëP»CHbPSPsC— PrPëC†C„C¿PSPeC†C-C—

P'PμP·PīPμPeP° C,P° PμC,,PμPeC,PēPIPSC-Cl'C,CH PsPrPSPsC‡P°Cl'PSPsPiPs
P·P°Cl'C,PsCl'CfPIP°PSPSClI Cl'PēP»PrPμPSP°C,,C-P»Cr P· C-PSC€PēPjPē
C-PSPIc-P±C-C,PsCḡP°PjPē PαP''P•5 P°P±Ps C-PSC€PēPjPē PīCḡPμPīP°CḡP°C,P°PjPē PrP»ClI
P»C-PeCfPIP°PSPSClI PiC-PīPμCḡC,PμPSP·C-C— P»PμPīPμPSPμPIPsC— P°CḡC,PμCḡC-C—,
ClIPeC- PjC-Cl'C,ClIc,CH Cl'PēP»PrPμPSP°C,,C-P», C‡Pē P· C-PSC€PēPjPē
PīCḡPμPīP°CḡP°C,P°PjPē PrP»ClI P»C-PeCfPIP°PSPSClI PμCḡPμPeC,PēP»CHPSPsC—
PrPēCl'C,,CfPSPeC‡C-C— PSPμ PIPēPIC‡P°P»PēCl'ClI. PŷPsPjCf P·P°Cl'C,PsCl'CfPIP°PSPSClI
C,P°PePēC... PePsPjP±C-PSP°C‡C-PNē PSPμ CḡPμPePsPjPμPSPPrPsPIP°PSPμ.

P'PīP»PēPI PSP° P·C-Cḡ

PμPsPIC-PrPsPjP»ClI»PsCl'CH PīCḡPs PIPēPSPēPePSPμPSPSClI PrPμC,,PμPeC,C-PI P·PsCḡCf
C,P° PIPēPīP°PrPePē PSPμP°CḡC,PμCḡC-P°P»CHPSPsC— PīPμCḡPμPrPSCḡPsC—
C-C€PμPjC-C‡PSPsC— PSPμPICḡPsPīP°C,C-C— P·PsCḡPsPIPsPiPs PSPμCḡPIP° Cf
P·PIBḡ™ ClI·PeCf C-P· P·P°Cl'C,PsCl'CfPIP°PSPSClI Pj Cl'PēP»PrPμPSP°C,,C-P»Cr C,P°
C-PSC€PēC... C-PSPIc-P±C-C,PsCḡC-PI PαP''P•5 (PrPēPI. CḡPsP·PrC-P» B«PμPsP±C-C‡PSC-
CḡPμP°PeC‡C-C—B»). PμP°C‡C-C''PSC,C-PI Cl'P»C-Pr PīPsPīPμCḡPμPrPēC,Pē, C‰Ps Cf
CḡP°P·C- CḡP°PīC,PsPIPsPiPs PīPsCḡCfC€PμPSPSClI P·PsCḡCf P·P°Cl'C,PsCl'CfPIP°PSPSClI
PīCḡPμPīP°CḡP°C,Cf Cl'P»C-Pr PīCḡPēPīPēPSPēC,Pē C,P° PSPμPīP°PNēPSPs
P·PIPμCḡPSCfC,PēCl'ClI PrPs P»C-PeP°CḡClI (PrPēPI. CḡPsP·PrC-P»
B«PμCḡPsC,PēPīPsPeP°P·P°PSPSClI»).

PḡPrPSPsC‡P°Cl'PSPμ P·P°Cl'C,PsCl'CfPIP°PSPSClI P· CḡPēC,PsPSP°PIC-CḡPsPj

PḡPrPSPsC‡P°Cl'PSPμ P·P°Cl'C,PsCl'CfPIP°PSPSClI Cl'PēP»PrPμPSP°C,,C-P»Cr C,P°
CḡPēC,PsPSP°PIC-CḡCf PSPμ CḡPμPePsPjPμPSPPrCfC''C,CHCl'ClI (PrPēPI. CḡPsP·PrC-P»
B«P'P·P°C''PjPsPrC-ClI P· C-PSC€PēPjPē P»C-PeP°CḡCl'CHPePēPjPē P·P°Cl'PsP±P°PjPē C,P°
C-PSC€C- PIPēPrPē PIP·P°C''PjPsPrC-PNēB»).

*PḡPrPSPsC‡P°Cl'PSPμ P·P°Cl'C,PsCl'CfPIP°PSPSClI P· P±P»PsPeP°C,PsCḡP°PjPē α-
P°PrCḡPμPSPsCḡPμC‡PμPīC,PsCḡC-PI*

PμP°C‡C-C''PSC,P°Pj, ClIPeC- P·P°Cl'C,PsCl'PsPICfCḡC,CH P±P»PsPeP°C,PsCḡPē α-
P°PrCḡPμPSPsCḡPμC‡PμPīC,PsCḡC-PI, P·P°Cl'C,PsCl'PsPICfPIP°C,Pē Cl'PēP»PrPμPSP°C,,C-P»
Cl'P»C-Pr P· PsP±PμCḡPμP¶PSC-Cl'C,CH, PsCl'PeC-P»CHPePē C,P°PeP° PePsPjP±C-PSP°C‡C-ClI
PjPsP¶Pμ PīCḡPēP·PIPμCl'C,Pē PrPs Cl'PēPjPīC,PsPjP°C,PēC‡PSPsC— PiC-PīPsC,PμPSP·C-C— Cf
PrPμClIPePēC... Cl'C...PēP»CHPSPēC... PrPs C‡CHPsPiPs PīP°C‡C-C''PSC,C-PI.
PŷPēPjPīC,PsPjP°C,PēC‡PSP° PiC-PīPsC,PμPSP·C-ClI P·P°P·PIPēC‡P°PNē PIPēPSPēPeP°C''
PīCḡPsC,ClIPiPsPj 4 PiPsPrPēPS PīC-Cl'P»ClI P·P°Cl'C,PsCl'CfPIP°PSPSClI
Cl'PēP»PrPμPSP°C,,C-P»Cr. PŷPsPjCf PrPs PīPsC‡P°C,PeCf P·P°Cl'C,PsCl'CfPIP°PSPSClI
Cl'PēP»PrPμPSP°C,,C-P»Cr PiPμPjPsPrPēPSP°PjC-C‡PSC- PīPsPeP°P·PSPēPePē PīP°C‡C-C''PSC,P°
PīCḡPē C,PμCḡP°PīC-C— P±P»PsPeP°C,PsCḡP°PjPē α-P°PrCḡPμPSPsCḡPμC‡PμPīC,PsCḡC-PI
PjP°CḡC,CH P±CfC,Pē Cl'C,P°P±C-P»CHPSPēPjPē. PŷP»C-Pr CḡPsP·PīP»ClIPSCfC,Pē
PjPsP¶P»PēPIC-Cl'C,CH P·P°Cl'C,PsCl'CfPIP°PSPSClI PīPsC‡P°C,PePsPIPsC— PrPsP·Pē
Cl'PēP»PrPμPSP°C,,C-P»Cr 25 PjPi (PrPēPI. CḡPsP·PrC-P» B«PŷPīPsCl'C-P±
P·P°Cl'C,PsCl'CfPIP°PSPSClI C,P° PrPsP·PēB»). PḡCḡC-Pj C,PsPiPs, PīPsC,CḡC-P±PSPs
PīCḡPsC-PSC,,PsCḡPjCfPIP°C,Pē PīP°C‡C-C''PSC,C-PI, ClIPe PrC-ClIc,Pē Cf CḡP°P·C-
PīPsClIPIPē Cl'PēPjPīC,PsPjC-PI PsCḡC,PsCl'C,P°C,PēC‡PSPsC— PiC-PīPsC,PμPSP·C-C—.

P'PīP»PēPI PSP° PeCḡPsPIPsC,PμC‡Cž

P''PsCl'P»C-PrP¶PμPSPSClIB B C,CḡPsPjP±PsC‡PēC,C-PI B P»CHPrPēPSPē

В РіСґPsPrPμPjPsPSCГC,СґCрPIP°P»Pө, В C%Ps В in B vitro В CГPөP»PrPμPSP°C,„C-P»B
 В PіPsC,РμPSCГC-СґC” P°PSC,PөP°PiCґPμPiP°CГC-PMөPSC- PμC,„PμPөC,Pө PSP°C,СґC-Сґ
 PSC-C,СґPsPiCґCрCГPөPrCр. PкPμPjP°C” P¶PsPrPSPsC— C-PSC,„PsCґPjP°CГC-C— C%PsPrPs
 P±PμP·PiPμPөPө P·P°CГC,PsCГCрPIP°PSPSCГ CГPөP»PrPμPSP°C,„C-P»Cр PiP°CГC-C”PSC,P°Pj
 C-P· PiPsCґCрCөPμPSPSCГPjPө P·PiPsCґC,P°PSPSCГ PeCґPsPIC- P°P±Ps PiPsCГC,СґPsCґ
 PiPμPiC,PөC±PSPsCґ PIPөCґP°P·PePsCґ. PҗP°PePөPj C±PөPSPsPj, P·P°CГC,PsCГCрPIP°PSPSCГ
 CГPөP»PrPμPSP°C,„C-P»Cр PiP°CГC-C”PSC,P°Pj CГC-C”C— PiCґCрPiPө PjPsP¶P»PөPIPμ
 P»PөCөPμ PiC-CГP»CГ CґPμC,РμP»СґPSPsC— PsCГC-PSPөPө CГPiC-PIPIC-PrPSPsCөPμPSPSCГ
 PePsCґPөCГC,C- C,P° CґPөP·PөPeC-PI.

P’C,CґP°C,P° CГP»CзC...Cз

P>C-PөP°CґCГPj CГP»C-Pr PiPsCґP°PrPөC,Pө PiP°CГC-C”PSC,P°Pj PiCґPөPiPөPSPөC,Pө
 P·P°CГC,PsCГCрPIP°PSPSCГ C-PSPiC-P±C-C,PsCґC-PI PαP”P•5, PIPөP»CґC±P°CґC±Pө
 PiCґPμPiP°CґP°C, P’C-PIP°PMөCґP°, C,P° PSPμPiP°PMөPSPs P·PIPμCґPSCрC,PөCГCГ P·P°
 PjPμPrPөC±PSPsCґ PrPsPiPsPjPsPiPsCґ Cр CґP°P·C- CґP°PiC,PsPIPsPiPs P·PSPөP¶PμPSPSCГ
 C±Pө PIC,CґP°C,Pө CГP»CрC...Cр. PμCґPs CГC- CГPIPөC%P°, CГPeC- C,P°PePsP¶
 PjPsP¶CрC,Сґ CГCрPiCґPsPIPsPrP¶CрPIP°C,PөCГCГ PrP·PIPsPSPsPj Cр PICрC...P°C... C,P°
 P·P°PiP°PjPsCґPsC±PμPSPSCГPj, PiPsPIC-PrPsPjP»CГP»PsCГCГ P· P°CГPsCГC-P°CГC-C”Cґ Cр
 C±P°CГC- C-P· P·P°CГC,PsCГCрPIP°PSPSCГPj C-PSPiC-P±C-C,PsCґC-PI PαP”P•5,
 PIPөP»CґC±P°CґC±Pө PiCґPμPiP°CґP°C, P’C-PIP°PMөCґP°. P’PөP·PSP°C±PөC,Pө, C±Pө CГC-
 CГPIPөC%P° PiCґCГPjPs PiPsPiCґ™ CГP·P°PSC- C-P· P·P°CГC,PsCГCрPIP°PSPSCГPj
 C-PSPiC-P±C-C,PsCґC-PI PαP”P•5, C±Pө P· C-PSCөPөPjPө C,„P°PeC,PsCґP°PjPө,
 PSPμPjPsP¶P»PөPIPs.

*PкPrPSPsC±P°CГPSPμ P·P°CГC,PsCГCзPIP°PSPSCГ C-P· PiC-PiPsC,РμPSP·PөPIPSPөPjPө
 PiCґPμPiP°CґP°C,P°PjPө*

P’C-PIP°PMөCґP° C±PөPSPөC,Сґ CГPөCГC,РμPjPSCр
 CГCрPrPөPSPsCґPsP·CөPөCґCґPIP°P»СґPSCр PrC-Сґ C,P° PjPsP¶Pμ PrP°P»C-
 P·PSPөP¶CрPIP°C,Pө P°CґC,РμCґC-P°P»СґPSPөPMө C,PөCГPe Cр PiP°CГC-C”PSC,C-PI, CГPeC-
 P·P°CГC,PsCГCрPICрCґC,Сґ PiC-PiPsC,РμPSP·PөPIPSC- P>C-PөP°CґCґCґPeC- P·P°CГPsP±Pө.
 Pj PsPeCґPμPjPsPjCр PrPsCГP»C-PrP¶PμPSPSC- P»C-PөP°CґCґCґPePsC— PIP·P°C”PjPsPrC-C—
 PiCґPө PsPrPSPsC±P°CГPSPsPjCр P·P°CГC,PsCГCрPIP°PSPSC- P°PjP»PsPrPөPiC-PSCр (5 PjPi
 P°P±Ps 10 PjPi) C,P° PiCґPμPiP°CґP°C,Cр P’C-PIP°PMөCґP° (100 PjPi) PiPμCґPsCґP°P»СґPSPs
 CГPiPsCГC,РμCґC-PiP°P»PsCГCГ CГPμCґPμPrPSC” PrPsPrP°C,PePsPIPμ P·PSPөP¶PμPSPSCГ
 CГPөCГC,PsP»C-C±PSPsPiPs C,PөCГPeCр PSP° 8 PjPj CґC,.. CГC,.. PrC-P°CГC,PsP»C-C±PSPsPiPs
 Bґ” PSP° 7 PjPj CґC,.. CГC..

*P—P°C...PIPсCґCґPIP°PSPSCГ, C%Ps PiPμCґPμPrP°CґC,СґCГCГ CГC,P°C,РμPIPөPj
 CөP»CГC...PsPj*

P—P°CГC,PsCГCрPIP°PSPSCГ PiCґPμPiP°CґP°C,Cр P’C-PIP°PMөCґP° PSPμ P·P°C...PөC%P°C”
 PIC-Pr P·P°C...PIPsCґCґPIP°PSCґ, C%Ps PiPμCґPμPrP°CґC,СґCГCГ CГC,P°C,РμPIPөPj
 CөP»CГC...PsPj. PҗP»C-Pr PiCґPsC-PSCГC,CґCрPeC,CрPIP°C,Pө PiP°CГC-C”PSC,C-PI C%PsPrPs
 PSPμPsP±C...C-PrPSPөC... P·P°PiPsP±C-P¶PSPөC... P·P°CГPsP±C-PI PrP»CГ P·P°C...PөCГC,Cр
 PIC-Pr P·P°C...PIPsCґCґPIP°PSCґ, C%Ps PiPμCґPμPrP°CґC,СґCГCГ CГC,P°C,РμPIPөPj
 CөP»CГC...PsPj, PIPөP»CґC±P°CґC±Pө PIC-CґCрCГ C-PjCрPSPsPrPμC,„C-C±PөC,Cр
 P»CґPrPөPSPө.

PαPμCґC,PөP»СґPSC-CГC,Сґ

РѹС-СІР»СІ Р·Р°СІС,РсСІСІРІР°РSPSCІ РrPsP·Рё 100 РjРi Р·РrPsСђРsРІРёРjРё
РrPsP±СђРsРІРsР»СђСІСІРjРё РSPµ СІРiРsСІС,РµСђС-РiР°Р»РsСІСІ РІРiР»РёРІСІ РSP°
РjРsСђС,,РsР»РsРiС-Сђ СІРё СђСІС...РsРjС-СІС,Сђ СІРiРµСђРjР°С,РsР·РsС—РrС-РІ.

Р"Рs СІРёР»Р°РrСІ Р'С-РІР°РNСђСђР° РІС...РsРrРёС,Сђ Р»Р°РёС,РsР·Р°, СІРёС%Рs СІ Р'Р°СІ
РІСІС,Р°РSPsРІР»РµРSPs РSPµРiРµСђРµРSPsСІРёРjС-СІС,Сђ РrРµСІРёРёС... СІСІРёСђС-РІ,
РiСђРsРёРsРSPСІСІР»СђС,СІРNС,РµСІСІ Р· Р»С-РёР°СђРµРj, РiРµСђСё РSC-Р¶
РiСђРёРNСРjР°С,Рё СІРµРNС Р»С-РёР°СђСІСђРёРёРNС Р·Р°СІС-Р±. РђСІРiР°СђС,Р°Рj (Р· 951)
РІ СІРёР»Р°РrС- Р»С-РёР°СђСІСђРёРsРiРs Р·Р°СІСРs±СІ С" РiРsС...С-РrРSPёРj
С,,РµРSC-Р»Р°Р»Р°РSC-РSCІ, С%Рs СІРІР»»СІС" РSPµР±РµР·РiРµСІ РrР»СІ
С...РІРsСђРёС... РSP° С,,РµРSC-Р»РёРµС,РsРSCІСђС-Сђ.

*Р—Р°СІС,РsСІСІРІР°РSPSCІ СІ РiРµСђС-РsРr РІР°РiС-С,РSPsСІС,С- Р°Р±Рs
РiРsРrСІРІР°РSPSCІ РiСђСІРrСђ.*

РѹСђРµРiР°СђР°С, РSPµ РiСђРёР·РSP°СІРµРSPёРNС РrР»СІ Р·Р°СІС,РсСІСІРІР°РSPSCІ
Р¶С-РSPёР°РjРё.

*Р—РrР°С,РSC-СІС,Сђ РІРiР»РёРІР°С,Рё РSP° СёРІРёРrРёС-СІС,Сђ СђРµР°РёСІС— РiСђРё
РёРµСђСІРІР°РSPSC- Р°РІС,РsС,СђР°РSPСІРiРsСђС,РsРj Р°Р±Рs С-РSCёРёРjРё
РjРµС...Р°РSC-Р·РjР°РjРё.*

Р"РsСІР»С-РrР¶РµРSCђ РІРiР»РёРёРІСІ РiСђРµРiР°СђР°С,СІ РSP° Р·РrР°С,РSC-СІС,Сђ
РёРµСђСІРІР°С,Рё С,СђР°РSPСІРiРsСђС,РSPёРjРё Р·Р°СІСРs±Р°РjРё С,Р° РiСђР°СІСђРІР°С,Рё
Р· С-РSCёРёРjРё РjРµС...Р°РSC-Р·РjР°РjРё РSPµ РiСђРsРІРsРrРёР»РsСІСІ.

РђСІРёС-Р»СђРёРё СІ РiСђРsСІРµСІС- РёР»С-РSC-СІРSPёС... РrPsСІР»С-РrР¶РµРSCђ
Р·Р°СІС,РсСІСІРІР°РSPSCІ СІРёР»Р»РrРµРSP°С,,С-Р»СІ РiРsРІС-РrPsРjР»СІР»РsСІСІ РiСђРs
РІРёРiР°РrРёРё Р·Р°РiР°РjРsСђСІРµРSPSCІ С,Р° РiРsСђСІСёРµРµРSPSCІ Р· Р±РsРёСІ
РsСђРiР°РSC-РІ Р·РsСђСІ, РiРµСђРµРr С,РёРj, СІРё СІС-РrР°С,Рё Р·Р° РёРµСђРjРs
С,СђР°РSPСІРiРsСђС,РSPsРiРs Р·Р°СІСРs±СІ Р°Р±Рs РiСђР°СІСђРІР°С,Рё Р·
РjРµС...Р°РSC-Р·РjР°РjРё, РiР°СІС-С"РSC,Р°Рj РSPµРsР±С...С-РrРSPs Р·Бђ™СІСІСІРІР°С,Рё,
СІРёРsСђ С" С—С...РSCІ С-РSPrРёРІС-РrСІР°Р»СђРSP° СђРµР°РёСІС-СІ РSP°
Р·Р°СІС,РсСІСІРІР°РSPSCІ РiСђРµРiР°СђР°С,СІ.

РŸРiРsСІС-Р± Р·Р°СІС,РsСІСІРІР°РSPSCІ С,Р° РrPsP·Рё.

РѹСђРµРiР°СђР°С, Р·Р°СІС,РсСІРsРІСІСђС,Сђ РiРµСђРsСђР°Р»СђРSPs.

РŸР°Р±Р»РµСІ СІР»С-Рr РiРsРІРSC-СІС,Сђ СђРsР·Р¶СІРІР°С,Рё С,Р°
РiСђРsРёРsРІС,РSCІС,Рё.

Р"РsСђРsСІР»С-

РѹСђРё РSPµРsР±С...С-РrРSPsСІС,С- РiСђРµРiР°СђР°С, Р'С-РІР°РNСђСђР° СІР»С-Рr

P·P°CÍ'C,PSCÍ'PsPICÍ'PIP°C,Pë P·P° 1 PiPsPrPëPSCÍ' PrPs CÍ'PµPeCÍ'CrP°P»CHbPSPsC—
P°PeC,PëPIPsPSCÍ'C,C-. P PµPePsPjPµPSPPrPsPIP°PSP° PrPsP·P° CÍ'C,P°PSPsPIPëC,CH 50 PjPi.
PhCÍ'PeC-P»CHbPePë PiCtjPë PsPrPSPsC†P°CÍ'PSPsPjCr P·P°CÍ'C,PSCÍ'CrPIP°PSPSC-
PiCtjPµPiP°CtjP°C,Cr P·C—P¶PµCt PIC-PrP±CÍ'PIP°C''C,CHbCÍ'Cl P·P°C,CtjPëPjPeP°
P°P±CÍ'PsCtjP±C†C-C— C,P° CrPiPsPIC-P»CHbPSCtC''C,CHbCÍ'Cl PµC,,PµPeC,
PiCtjPµPiP°CtjP°C,Cr (PrPëPI. CtjPsP·PrC-P» B«PαP°CtjPjP°PePsPeC-PSPµC,PëPeP°B»),
CtjPµPePsPjPµPSPPrPsPIP°PSPs P·P°CÍ'C,PSCÍ'PsPICÍ'PIP°C,Pë PiCtjPµPiP°CtjP°C, PSP°C,C%Pµ.

P—P°P»PµP¶PSPs PIC-Pr PµC,,PµPeC,PëPIPsPSCÍ'C,C- C,P° PiPµCtjPµPSPsCÍ'PëPjPsCÍ'C,C-
PrPsP·Cr PiCtjPµPiP°CtjP°C,Cr PjPsP¶PSP° P·P±C-P»CHbCëPëC,Pë PrPs 100 PjPi P°P±Ps
P·PjPµPSCëPëC,Pë PrPs 25 PjPi*. PëP°PeCÍ'PëPjP°P»CHbPSP° CtjPµPePsPjPµPSPPrPsPIP°PSP°
PrPsP·P° CÍ'C,P°PSPsPIPëC,CH 100 PjPi.

PëP°PeCÍ'PëPjP°P»CHbPSP° CtjPµPePsPjPµPSPPrPsPIP°PSP° C†P°CÍ'C,PsC,P°
P·P°CÍ'C,PSCÍ'CrPIP°PSPSCl PiCtjPµPiP°CtjP°C,Cr CÍ'C,P°PSPsPIPëC,CH 1 CtjP°P· PSP°
PrPsP±CÍ'.

PµP°C†C-C''PSC,Pë P»C-C,PSCHbPsPiPs PIC-PeCé

PkPµPsP±C...C-PrPSC-CÍ'C,CH Cr PePsCtjPµPeC†C-C— PrPsP·Pë PiP°C†C-C''PSC,P°Pj
P»C-C,PSCHbPsPiPs PIC-PeCÍ' (PIC-PePsPj PIC-Pr 65 CtjPsPeC-PI) PIC-PrCÍ'CrC,PSCl.

PµP°C†C-C''PSC,Pë P·PSPëCtjPePsPIPsCt PSPµPrPsCÍ'C,P°C,PSC-CÍ'C,Ct

P''P»Cl PiP°C†C-C''PSC,C-PI C-P·PSPëCtjPePsPIPsCt PSPµPrPsCÍ'C,P°C,PSC-CÍ'C,Ct
P»PµPiPePsPiPs C,P° PiPsPjC-CtjPSPsPiPs CÍ'C,CrPiPµPSCl C,C¶P¶PePsCÍ'C,C-
(PeP»C-CtjPµPSCÍ' PeCtjPµP°C,PëPSC-PSCÍ' 30Btj''80 PjP»/C...PI) CtjPµPePsPjPµPSPPrPsPIP°PSP°
PrPsP·P° PiCtjPµPiP°CtjP°C,Cr C'' C,P°PePsCt P¶, ClPe PSP°PIPµPrPµPSPs PIPëC%Pµ Cr
CtjPsP·PrC-P»C- B«P''PsCtjPsCÍ'P»C-B».

PhCÍ'PeC-P»CHbPePë Cr PiP°C†C-C''PSC,C-PI P·PSPëCtjPePsPIPsCt
PSPµPrPsCÍ'C,P°C,PSC-CÍ'C,Ct C,C¶P¶PePsPiPs CÍ'C,CrPiPµPSCl (PeP»C-CtjPµPSCÍ'
PeCtjPµP°C,PëPSC-PSCÍ' < 30 PjP»/C...PI) PeP»C-CtjPµPSCÍ' CÍ'PëP»PrPµPSP°C,,C-P»Cr
P·PSPëP¶PµPSPëPñ, CÍ'P»C-Pr CtjPsP·PiP»ClPSCÍ'C,Pë PjPsP¶P»PëPIC-CÍ'C,CH
P·P°CÍ'C,PSCÍ'CrPIP°PSPSCl PrPsP·Pë B B B B B 25 PjPi*. P—P°P»PµP¶PSPs PIC-Pr
PµC,,PµPeC,PëPIPsPSCÍ'C,C- C,P° PiPµCtjPµPSPsCÍ'PëPjPsCÍ'C,C- PiCtjPµPiP°CtjP°C,Cr PiCtjPë
PSPµPsP±C...C-PrPSPsCÍ'C,C- PrPsP·Cr PjPsP¶PSP° PiPsCÍ'C,CrPiPsPIPs P·P±C-P»CHbCëPëC,Pë
PrPs 50 PjPi C,P° PrPs 100 PjPi.

PµP°C†C-C''PSC,Pë P·PiPµC†C-PSPePsPIPsCt PSPµPrPsCÍ'C,P°C,PSC-CÍ'C,Ct

PhCÍ'PeC-P»CHbPePë Cr PiP°C†C-C''PSC,C-PI C-P·PiPµC†C-PSPePsPIPsCt
PSPµPrPsCÍ'C,P°C,PSC-CÍ'C,Ct (PSP°PiCtjPëPeP»P°Pr, C-P·C†PëCtjPsP·PsPj) PeP»C-CtjPµPSCÍ'
CÍ'PëP»PrPµPSP°C,,C-P»Cr P·PSPëP¶PµPSPëPñ, CÍ'P»C-Pr CtjPsP·PiP»ClPSCÍ'C,Pë
PjPsP¶P»PëPIC-CÍ'C,CH P·P°CÍ'C,PSCÍ'CrPIP°PSPSCl PrPsP·Pë 25 PjPi*. P—P°P»PµP¶PSPs PIC-Pr
PµC,,PµPeC,PëPIPsPSCÍ'C,C- C,P° PiPµCtjPµPSPsCÍ'PëPjPsCÍ'C,C- PiCtjPµPiP°CtjP°C,Cr PiCtjPë
PSPµPsP±C...C-PrPSPsCÍ'C,C- PrPsP·Cr PjPsP¶PSP° PiPsCÍ'C,CrPiPsPIPs P·P±C-P»CHbCëPëC,Pë
PrPs 50 PjPi C,P° PrPs 100 PjPi.

*PµP°C†C-C''PSC,Pë, ClPeC- P·P°CÍ'C,PSCÍ'PsPICéCtC,CH C-PSCëC- P»C-PeP°CtjCÍ'CHbPeC-
P·P°CÍ'PsP±Pë*

PiPeC%Ps PiP°C†C-C''PSC,Pë PsPrPSPsC†P°CÍ'PSPs P·P°CÍ'C,PSCÍ'PsPICÍ'CtC,CH

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

PḡPµCḡPµP· CḡPëP·PëPe CḡPsP·PIPëC,PeCḡ PİPsCḡC,CḡCḡP°P»CHPSPsC— PiC-PİPsC,PµPSP·C-C— Cḡ PİP°C†C-C''PSC,C-PI, CḡPeC- P·P°CḡC,PsCḡPsPICḡCḡC,CH P±P»PsPeP°C,PsCḡPë α-P°PrCḡPµPSPsCḡPµC†PµPiC,PsCḡC-PI, PrPs PİPsC†P°C,PeCḡ P·P°CḡC,PsCḡCḡPIP°PSPSCḡ CḡPëP»PrPµPSP°C,,C-P»Cḡ CḡP»C-Pr CḡPİPµPIPSPëC,PëCḡCḡ, C%Ps C—C...PSC-PNë CḡC,P°PSB CḡC,P°P±C-P»CHPSPëPNë. PḡP°PePsPḡ PİPsC,CḡC-P±PSPs CḡPsP·PiP»CḡPSCrC,Pë PjPsPḡP»PëPIC-CḡC,CH P·P°CḡC,PsCḡCḡPIP°PSPSCḡ PİPsC†P°C,PePsPIPsC— PrPsP·Pë 25 PjPi* (PrPëPI. CḡPsP·PrC-P»Pë B«P'P·P°C''PjPsPrC-Cḡ P· C-PSC€PëPjPë P»C-PeP°CḡCḡCHPePëPjPë P·P°CḡPsP±P°PjPë C,P° C-PSC€C- PIPëPrPë PIP·P°C''PjPsPrC-PNëB» C,P° B«PhCḡPsP±P»PëPIPsCḡC,C- P·P°CḡC,PsCḡCḡPIP°PSPSCḡB»).

*B P—P°CḡC,PsCḡPsPICḡPIP°C,Pë Cḡ PIC-PrPİPsPIC-PrPSPsPjCḡ PrPsP·CḡPIP°PSPSC-.

P''C-C,Pë.

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

PµPµCḡPµPrPsP·CḡPIP°PSPSCḡ.

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

Pj CḡP°P·C- PİPµCḡPµPrPsP·CḡPIP°PSPSCḡ PİCḡPë PSPµPsP±C...C-PrPSPsCḡC,C- PIPrP°CḡC,CHCḡCḡ PrPs P·PIPëC†P°PNëPSPëC... PİC-PrC,CḡPëPjCḡPIP°P»CHPSPëC... P·P°C...PsPrC-PI. PµCḡPëCḡPePsCḡPµPSPSCḡ PeP»C-CḡPµPSCḡCḡ CḡPëP»PrPµPSP°C,,C-P»Cḡ PİCḡPë PİPµPjPsPrC-P°P»C-P·C- PjP°P»PsPNëPjPsPIC-CḡPSPµ C†PµCḡPµP· PIPëCḡPsPePëPNë CḡC,CḡPİC-PSCḡ P·PIBḡ™ CḡP·CḡPIP°PSPSCḡ PİCḡPµPİP°CḡP°C,Cḡ P· P±C-P»PeP°PjPë PİP»P°P·PjPë PeCḡPsPIC- C,P° PIC-PrCḡCḡC,PSC-CḡC,CH PµP»C-PjC-PSP°C†C-C— CḡPëP»PrPµPSP°C,,C-P»Cḡ C-P· CḡPµC†PµCH.

PμPsP±C-C‡PSC- CḡPμP°PeC†C-C—.

PμCḡPsC,,C-P»CH P±PμP·PīPμPePē CÍPēP»PrPμPSP°C,,C-P»CÍ P±P°P·CÍC”C,CHCÍCÍ PSP°
PrP°PSPēC..., PsC,CḡPēPjP°PSPēC... C-P· 74 PīPsPrPIC-PNēPSPēC... CÍP»C-PīPēC...
PīP»P°C†PμP±Ps-PePsPSC,CḡPsP»CHPsPIP°PSPēC... PeP»C-PSC-C‡PSPēC...
PrPsCÍP»C-PrP¶PμPSCH (9570 PīP°C†C-C”PSC,C-PI). PḡP°PNēC‡P°CÍC,C-CēPμ
PīPsPIC-PrPsPjP»CÍP»PsCÍCÍ PīCḡPs C,P°PeC- PīPsP±C-C‡PSC- CḡPμP°PeC†C-C—, CÍPe
PīPsP»PsPIPSēPNē P±C-P»CH, PīCḡPēPīP»PēPIPē PeCḡPsPIC-, PrPēCÍPīPμPīCÍC-CÍ,
P·P°PeP»P°PrPμPSC-CÍC,CH PSPsCÍP°, P±C-P»CH CÍ CÍPīPēPSC-,
P·P°PīP°PjPsCḡPsC‡PμPSPSCÍ, PSCÍPrPsC,P°, PīCḡPēPīP»PēPIPē P¶P°CḡCÍ,
PīPsCḡCÍCēPμPSPSCÍ P·PsCḡCÍ, C†C-P°PSPsPīCÍC-CÍ C,P° P·P°C,CHPjP°CḡPμPSPSCÍ
P·PsCḡCÍ. P†PSC,,PsCḡPjP°C†C-CÍ C%PsPrPs PīPsP±C-C‡PSPēC... CḡPμP°PeC†C-PNē CÍ
CḡP°PjPeP°C... PīPsCÍC,PjP°CḡPePμC,PēSPiPsPIPsPīPs CÍPīPsCÍC,PμCḡPμP¶PμPSPSCÍ
P±CÍP»P° P·C-P±CḡP°PSP° PīCḡPsC,CÍPiPsPj PīPμCḡC-PsPrCÍ C,CḡPēPIP°P»C-CÍC,Cḡ
P±C-P»CHCēPμ PSC-P¶ 10 CḡPsPeC-PI. PhCÍPeC-P»CHPePē P·P°CÍPIPSēPePsPIC-
PīPsPIC-PrPsPjP»CÍP»PsCÍCÍ PSPμ PīCḡPs PICÍC- PīPsP±C-C‡PSC- CḡPμP°PeC†C-C— C,P°
PSPμ PICÍC- PīPsP±C-C‡PSC- CḡPμP°PeC†C-C— P±CÍP»Pē PīPeP»CḡC‡PμPSC- PI P±P°P·CÍ
PrP°PSPēC... P· P±PμP·PīPμPePē, C‡P°CÍC,PsC,P° C,P°PePēC... CḡPμP°PeC†C-PNē PSPμ PjPsP¶Pμ
P±CÍC,Pē PrPsCÍC,PsPIC-CḡPSPs PIPēP·PSP°C‡PμPSP°.

PjCÍC- PeP»C-PSC-C‡PSPs P·PSP°C‡CÍC%C- PīPsP±C-C‡PSC- CḡPμP°PeC†C-C—, C%Ps
CÍPīPsCÍC,PμCḡC-PiP°P»PēCÍCÍ PīC-Pr C‡P°CÍ PeP»C-PSC-C‡PSPēC...
PrPsCÍP»C-PrP¶PμPSCH C‡P°CÍC,C-CēPμ, PSC-P¶ PīCḡPē P·P°CÍC,PsCÍCÍPIP°PSPSC-
PīP»P°C†PμP±Ps, PSP°PIPμPrPμPSPs PSPēP¶C‡Pμ P·P° CÍPēCÍC,PμPjP°PjPē PsCḡPiP°PSC-PI C,P°
C‡P°CÍC,PsC,PsCḡ: PrCÍP¶Pμ C‡P°CÍC,Ps (≥1/10), C‡P°CÍC,Ps (≥1/100 вḡ” <1/10),
PSPμC‡P°CÍC,Ps (≥1/1000 вḡ” <1/100), CḡC-PrPePs (≥1/10000 вḡ” <1/1000), PrCÍP¶Pμ
CḡC-PrPePs (<1/10000), C‡P°CÍC,PsC,P° PSPμPIC-PrPsPjP° (PSPμ PjPsP¶Pμ P±CÍC,Pē
PICÍC,P°PSPsPIP»PμPSP° PIPēC...PsPrCÍC‡Pē P· PSP°CÍPIPSēC... PrP°PSPēC...
PḡP°CÍC,PsC,P° PeP»C-PSC-C‡PSPs P·PSP°C‡CÍC%PēC... PīPsP±C-C‡PSPēC... CḡPμP°PeC†C-PNē,
PīCḡPs CÍPeC- PīPsPIC-PrPsPjP»CÍP»PsCÍCÍ CÍ CḡP°PjPeP°C...
PīPsCÍC,PjP°CḡPePμC,PēSPiPsPIPēC... CÍPīPsCÍC,PμCḡPμP¶PμPSCH, PIPēP·PSP°C‡PμPSP°
CÍPe PSPμPIC-PrPsPjP°. Pj PjPμP¶P°C... PePsP¶PSPsC— PīCḡCÍPīPē P·P° C‡P°CÍC,PsC,PsCḡ
PīPsP±C-C‡PSC- CḡPμP°PeC†C-C— PSP°PIPμPrPμPSC- CÍ PīPsCḡCÍPrPeCÍ
P·PjPμPSCēPμPSPSCÍ C—C... CÍPμCḡPNēPsP·PSPsCÍC,C-.

P†PSC,,PμPeC†C-PNēPSC- C,P° C-PSPIP°P·PēPIPS- P·P°C...PIPsCḡCḡPIP°PSPSCÍ

PḡPμC‡P°CÍC,Ps: CḡPēPSC-C,.

P— P±PsPeCž C-PjCžPSPSPsC— CÍPēCÍC,PμPjPē

PḡPμC‡P°CÍC,Ps: CḡPμP°PeC†C-C— PīC-PrPIPēC%PμPSPsC— C‡CÍC,P»PēPIPsCÍC,C-.

P— P±PsPeCž PSPμCḡPIPsPIPsC— CÍPēCÍC,PμPjPē

P”CÍP¶Pμ C‡P°CÍC,Ps: PīPsP»PsPIPSēPNē P±C-P»CH.

PḡP°CÍC,Ps: P·P°PīP°PjPsCḡPsC‡PμPSPSCÍ.

PḡPμC‡P°CÍC,Ps: CÍPsPSP»PēPIC-CÍC,CH, PīC-PīPμCÍC,PμP·C-CÍ.

P C-PrPePs: C-PSCÍCíP»CHC,, CÍPëPSPePsPíPµ, C,CĤP°PSP·PëC,PsCĤPSP° C-CëPµPjC-C‡PSP°
P°C,P°PeP°, CÍCíPrPsPjPë, CĤPµC‡PëPrPëPIPë CÍCíPrPsPj.

P— P±PsPeCž PsCĤPiP°PSC-PI P·PsCĤCž

PŞP°CÍC,Ps: PíPsCĤCíCëPµPSPSCİ CÍPíCĤPëPñPSCİC,C,Cİ PePsP»CHPsCĤCí*,
CĤPsP·P»P°PrPë P·PsCĤCí, P·P°C,CHPjP°CĤPµPSPSCİ P·PsCĤCí.

PќPµC‡P°CÍC,Ps: CĤPsP·P»P°PrPë CÍP»CHPsP·PsPIPëPrC-P»PµPSPSCİ**, P±C-P»CH PI
PsC‡P°C..., C,,PsC,PsC,,PsP±C-Cİ, C,,PsC,PsPíCÍC-Cİ, PiC-PíPµCĤPµPjC-Cİ PsC‡PµPñ,
CİCÍPëCĤP°PIC-CÍC,CH P·PsCĤCí, PePsPSbĤ™ CĤPSPeC,PëPIC-C,.

P C-PrPePs: PSPµP°CĤC,PµCĤC-P°P»CHPSP° PíPµCĤPµPrPSCİ C-CëPµPjC-C‡PSP°
PSPµPñCĤPsPíP°C,C-Cİ P·PsCĤPsPIPsPíPs PSPµCĤPIP°, PsPeP»CHP·C-Cİ CÍCíPrPëPS
CÍC-C,PeC-PIPePë, CĤPµC,PëPSP°P»CHPSPëPñ PeCĤPsPIPsPIPëP»PëPI,
P°CĤC,PµCĤC-PsCÍPeP»PµCĤPsC,PëC‡PSP° CĤPµC,PëPSPsPíP°C,C-Cİ, PíPsCĤCíCëPµPSPSCİ
P· P±PsPeCí CÍC-C,PeC-PIPePë, PiP»P°CíPePsPjP°, PrPµC,,PµPeC,Pë PíPsP»Cİ P·PsCĤCí,
PrPëPíP»PsPíC-Cİ, P·PSPëP¶PµPSPSCİ PiPsCÍC,CĤPsC,Pë P·PsCĤCí, PjC-PsPíC-Cİ,
P°CÍC,PµPSPsPíC-Cİ, PíP»P°PIP°CĤC‡C- PíPsPjCíC,PSC-PSPSCİ CÍPeP»PëCÍC,PsPiPs
C,C-P»P°, PíPsCĤCíCëPµPSPSCİ P· P±PsPeCí CĤP°PñPrCíP¶PSPsC— PsP±PsP»PsPSPePë,
PjC-PrCĤC-P°P, PíPsCİPIP° CÍCİCĤC‡PëC... PeCĤCíPiC-PI PSP°PIPePsP»Ps PrP¶PµCĤPµP»P°
CÍPIC-C,P»P° (PiP°P»Ps) Cí PíPsP»C- P·PsCĤCí, PSP°P±CĤCİPe PsC‡PµPñ,
PíCĤPëPíCíC...P»C-CÍC,CH PsC‡PµPñ, PíPsCĤCíCëPµPSPSCİ P· P±PsPeCí PsC‡PµPñ,
PiC-PíPµCĤPµPjC-Cİ PePsPSbĤ™ CĤPSPeC,PëPIPë, PíPsPrCĤP°P·PSPµPSPSCİ PsC‡PµPñ,
P°PSPsPjP°P»CHPSC- PIC-PrC‡CíC,C,Cİ PI PsC‡P°C..., PSP°P±CĤCİPe PíPsPIC-Pe,
P·PSPµP±P°CĤPIP»PµPSPSCİ CÍPeP»PµCĤPë.

*P— P±PsPeCž PsCĤPiP°PSC-PI CÍP»CžC...Cž C,P° PIPµCÍC,PëP±CžP»CİCĤPSPsPiPs
P°PíP°CĤP°C,Cž*

PќPµC‡P°CÍC,Ps: P·P°PíP°PjPsCĤPsC‡PµPSPSCİ, PIPµCĤC,PëPiPs, PrP·PIC-PS Cí PICíC...P°C....

P C-PrPePs: PíP»CíC...PsC,P°.

P— P±PsPeCž CÍPµCĤC‡PµPIPs-CÍCžPrPëPSPSPsC— CÍPëCÍC,PµPjPë

PŞP°CÍC,Ps: PíCĤPëPíP»PëPIPë PeCĤPsPIC- PrPs PsP±P»PëC‡C‡Cİ, PíCĤPëPíP»PëPIPë
P¶P°CĤCí.

PќPµC‡P°CÍC,Ps: C,P°C...C-PeP°CĤPrC-Cİ, PíPsCÍPëP»PµPSPµ CÍPµCĤC‡PµP±PëC,C,Cİ,
P°CĤC,PµCĤC-P°P»CHPSP° PiC-PíPµCĤC,PµPSP·C-Cİ, P°CĤC,PµCĤC-P°P»CHPSP°
PiC-PíPsC,PµPSP·C-Cİ.

P C-PrPePs: CĤP°PíC,PsPIP° CÍPµCĤC‡PµPIP° CÍPjPµCĤC,CH, C-PSC,,P°CĤPeC,
PjC-PsPeP°CĤPrP°, CëP»CíPSPsC‡PePsPIP° P°CĤPëC,PjC-Cİ, C,,C-P±CĤPëP»CİC‡C-Cİ
PíPµCĤPµPrCÍPµCĤPrCH, PSPµCÍC,P°P±C-P»CHPSP° CÍC,PµPSPsPeP°CĤPrC-Cİ.

*P— P±PsPeCž PrPëC...P°P»CHPSPsC— CÍPëCÍC,PµPjPë, PiCĤCžPrPSPsC— PeP»C-C,PePë C,P°
CÍPµCĤPµPrPsCÍC,C-PSPSCİ*

PŞP°CÍC,Ps: P·P°PeP»P°PrPµPSC-CÍC,CH PSPsCÍP°.

PќPµC‡P°CÍC,Ps: PSPsCÍPsPIP° PeCĤPsPIPsC,PµC‡P°, P·P°PeP»P°PrPµPSC-CÍC,CH

ΠΪCΉPēPrP°C,PePsPIPēC... ΠiP°P·CfC... PSPScfP°.

P C-PrPePs: PIC-PrC‡CfC,C,Cf CfC,PēCfPSPμPSPSCf Cf ΠiPsCΉP»C-, PSP°P±CΉCfPe CfP»PēP·PsPSC— PsP±PsP»PsPSPePē PSPScfP°, CfCfC...C-CfC,CΉ Cf PSPScfC-.

P— P±PsPeCζ CēP»CζPSPePsPIPs-PePēCēPePsPIPsPiPs C,CΉP°PeC,Cζ

P§P°CfC,Ps: PSCfPrPsC,P°, PrPēCfΠiPμΠiCfC-Cf.

PκPμC‡P°CfC,Ps: ΠiP°CfC,CΉPsPμP·PsC,,P°ΠiPμP°P»CΉPSP° CΉPμC,,P»CΉPeCfPSP° C...PIPsCΉPsP±P°, P±P»CΉPIP°PSPSCf, P±C-P»CΉ Cf ΠiPμCΉC...PSC-PNē C‡P°CfC,PēPSC-P¶PēPIPsC,P°, CfCfC...C-CfC,CΉ Cf CΉPsC,C-.

P C-PrPePs: ΠiC-ΠiPμCfC,PμP·C-Cf CΉPsC,PsPSC— ΠiPsCΉPsP¶PSPēPSPē.

P— P±PsPeCζ CēPeC-CΉPē C,P° ΠiC-PrCēPeC-CΉPSPSc— PeP»C-C,PePsPIPēPSPē

PκPμC‡P°CfC,Ps: PIPēCfPēΠiP°PSPSCf.

P C-PrPePs: CfPēPSPPrCΉPsPj PŸC,C-ΠiPμPSCfP°bΉ“P”P¶PsPSCfPsPSP°, C,PsPeCfPēC‡PSPēPNē PμΠiC-PrPμCΉPjP°P»CΉPSPēPNē PSPμPeCΉPsP»C-P·.

P— P±PsPeCζ PsΠiPsCΉPSPSc-CΉCζC...PsPIPsPiPs P°ΠiP°CΉP°C,Cζ C,P° CfΠiPsP»CζC‡PSPSc— C,PeP°PSPēPSPē

PκPμC‡P°CfC,Ps: PjC-P°P»ΠiC-Cf, P±C-P»CΉ Cf PeC-PSC†C-ΠiPeP°C....

P— P±PsPeCζ CfPμC‡PsPIPēPrC-P»CΉPSPSc— CfPēCfC,PμPjPē

PκPμC‡P°CfC,Ps: ΠiPμPjP°C,CfCΉC-Cf.

P— P±PsPeCζ CΉPμΠiCΉPsPrCζPeC,PePIPSPsC— CfPēCfC,PμPjPē C,P° PjPsP»PsC‡PSPēC... P·P°P»PsP·

P C-PrPePs: PeCΉPsPIPsC,PμC‡P° P·C- CfC,P°C,PμPIPsPiPs C‡P»PμPSP°, ΠiCΉC-P°ΠiC-P·Pj, ΠiPμPjP°C,PsCfΠiPμCΉPjC-Cf, ΠiPsPrPsPIP¶PμPSP° PμCΉPμPeC†C-Cf.

P— P°ΠiP°P»CΉPSC- CΉPsP·P»P°PrPē C,P° CΉPμP°PeC†C-C— Cζ PjC-CfC†C- ΠiΠiPμPrPμPSPSCf

PκPμC‡P°CfC,Ps: P±C-P»CΉ Cf ΠiCΉCfPrCfC..., ΠiC-PrPIPēC%PμPSP° PIC,PsPjP»CΉPIP°PSC-CfC,CΉ, PIC-PrC‡CfC,C,Cf P¶P°CΉCf.

P C-PrPePs: ΠiPsPrCΉP°P·PSPμPSPSCf.

P>P°P±PsCΉP°C,PsCΉPSC- C,PμCfC,Pē

PκPμC‡P°CfC,Ps: ΠiC-PrPIPēC%PμPSP° C‡P°CfC,PsC,P° CfPμCΉC‡PμPIPēC... CfPePsCΉPsC‡PμPSCΉ.

* B PμPsCΉCfCēPμPSPSCf CfΠiCΉPēPNēPSCfC,C,Cf PePsP»CΉPsCΉCf: C...P»PsCΉPsΠiCfC-Cf, C...CΉPsPjP°C,PsΠiCfC-Cf, C†C-P°PSPsΠiCfC-Cf, PμCΉPēC,CΉPsΠiCfC-Cf, PeCfP°PSC,PsΠiCfC-Cf.

** PμPsCΉCfCēPμPSPSCf CfP»CΉPsP·PsPIPēPrC-P»PμPSPSCf: CfCfC...C-CfC,CΉ Πi

PsC†P°C..., PīPsCḡCfC€PμPSPSCḡ C†P»CHbPsP·PsPIPëPrC-P»PμPSPSCḡ C,P°
PīC-PrPIPëC%PμPSPSCḡ C†P»CHbPsP·PsPIPëPrC-P»PμPSPSCḡ.

PkPëP¶C†PμP·P°P·PSP°C†PμPSC- CḡIPIPëC%P° C†PīPsC†C,PμCḡC-PiP°P»PëC†Cḡ C† < 2 %
PīP°C†C-C"°PSC,C-PI C† PīCḡPsC†PμC†C- PePsPSC,CḡPsP»CHbPsPIP°PSPëC...
PeP»C-PSC-C†PSPëC... PrPsC†P»C-PrP¶PμPSCḡ; PīCḡPëC†PëPSPSPëPNë
PIP·P°C"°PjPsP·PIbḡ™ CḡP·PsPe PSPμ PIPëP·PSP°C†PμPSPëPNë. PμPsPIC-PrPsPjP»PμPSPSCḡ
PIPeP»CHC†P°P»Pë CḡIPIPëC%P°, C%P°P° PjP°P»Pë PIC-CḡPsPiC-PrPSPëPNë P·PIbḡ™ CḡP·PsPe
C-P· P·P°C†C,P°C†C†PIP°PSPSCḡPj PīCḡPμPīP°CḡP°C,C†. PīIPIPëC%P°, CḡPeC- PSPμ P±C†P»Pë
P·P°P·PSP°C†PμPSC-, P±C†P»Pë P»PμPiPePëPjPë C,P° PīPsPIC-PrPsPjP»PμPSPSCḡ P±C†P»Pë
PrC†P¶Pμ PSPμC,P°C†PSPëPjPë, C%P°P° PjP°C,Pë P·PSP°C†PμPSPSCḡ.

P—P°PiP°P»CHbPSC-. PkP°P±CḡCḡPe PsP±P»PëC†C†Cḡ, CḡPμP°PeC†C-C—
C,,PsC,P°C†C†C,P»PëPIPsC†C,C-, C€PsPe, P°C†C,PμPSC-Cḡ, P±C-P»CHb, CḡP°PiC,P°PIPμ
PīP°PrC-PSPSCḡ, P±C-P»CHb C† P¶PëPIPsC,C-, CḡP°PiC,P°PIPμ PīPsC€PePsPrP¶PμPSPSCḡ.

P— P±PsPeCž C†PμCḡC†PμPIPs-C†CžPrPëPSPSPsC— C†PëC†C,PμPjPë:
C†C,PμPSPsPeP°CḡPrC-Cḡ, PḡV-P±P»PsPeP°PrP°, PjC-PiCḡPμPSCḡ, PīPsC†C,C†CḡP°P»CHbPSP°
PiC-PīPsC,PμPSP·C-Cḡ, C-C€PμPjC-Cḡ PjC-PsPeP°CḡPrP°, C,CḡPsPjP±PsP· C†C†PrPëPS
PīPsP»PsPIPsPīPs PjPsP·PeC†, CḡP°PiC,P°PIP° P·C†PīPëPSPeP° C†PμCḡC†Cḡ,
PIC-PrC...PëP»PμPSPSCḡ PīPsPeP°P·PSPëPeC-PI P·PḡP", PeP°CḡPrC-PsPjC-PsPīP°C,C-Cḡ.

P— P±PsPeCž C€P»CžPSPePsPIPs-PePëC€PePsPIPsPiPs C,CḡP°PeC,Cž: PiP»PsC†PëC,, PePsP»C-C,,
PrPëC†C,,P°PiC-Cḡ, PīP°C†C,CḡPëC,, PīP°C†C,CḡPsPμPSC,PμCḡPëC,, PμP·PsC,,P°PiC-C,,
C†C,P°PjP°C,PëC,, PīPsCḡCfC€PμPSPSCḡ CḡPμP·C†P»CHbC,P°C,C-PI PīPμC†C-PSPePsPIPëC...
PīCḡPsP±, CḡPμPeC,P°P»CHbPSP° PeCḡPsPIPsC,PμC†P°, PiC-PSPiC-PIC-C,.

P— P±PsPeCž C†PëC†C,PμPjPë PeCḡPsPIC- C,P° P»C-PjC,,P°C,PëC†PSPsC— C†PëC†C,PμPjPë:
P°PSPμPjC-Cḡ, P»PμPNëPePsPīPμPSC-Cḡ.

P PsP·P»P°PrPë PjPμC,P°P±PsP»C-P·PjCž C,P° C...P°CḡC†CžPIP°PSPSCḡ: C†PīCḡP°PiP°,
PSP°P±CḡCḡPe, PīPsPrP°PiCḡP°, PSPμC†C,P°P±C-P»CHbPSPëPNë PrC-P°P±PμC,,
PiC-PīPμCḡPiP»C-PePμPjC-Cḡ, PīPμCḡPëC,,PμCḡPëC†PSC- PSP°P±CḡCḡPePë,
PiC-PīPμCḡCfCḡPëPePμPjC-Cḡ, PiC-PīPsPiP»C-PePμPjC-Cḡ, PiC-PīPμCḡPSP°C,CḡC-C"°PjC-Cḡ.

P— P±PsPeCž C†PePμP»PμC,PSPs-Pjēḡ™ CḡP·PsPIPsC— C†PëC†C,PμPjPë: P°CḡC,CḡPëC,,
P°CḡC,CḡPsP·, CḡPsP·CḡPëPI C†C†C...PsP¶PëP»P»Cḡ, C,PμPSPsC†PëPSPsPIC-C,, P±C-P»CHb C†
PeC-C†C,PeP°C..., PjC-P°C†C,PμPSC-Cḡ, C†PëPSPsPIC-C,.

P— P±PsPeCž PSPμCḡPIPsPIPsC— C†PëC†C,PμPjPë: P°C,P°PeC†C-Cḡ, PSPμPICḡP°P»PiC-Cḡ,
PSPμPNëCḡPsPīP°C,C-Cḡ, PīP°CḡPμC†C,PμP·C-Cḡ, C,CḡPμPjPsCḡ, PIPμCḡC,PëPiPs,
PrPμPiCḡPμC†C-Cḡ, P±PμP·C†PsPSPSCḡ, P°PSPsPjP°P»CHbPSC- C†PSPsPIPëPrC-PSPSCḡ,
P·PSPëP¶PμPSPSCḡ CḡPμC,,P»PμPeC†C-PI.

P— P±PsPeCž PrPëC...P°P»CHbPSPsC— C†PëC†C,PμPjPë: P°C†C,PjP°, PrPëC†PīPSPsPμ,
P»P°CḡPëPSPiC-C,, C,,P°CḡPëPSPiC-C,, C†PëPSC†C†PëC,, P±CḡPsPSC...C-C,, PīPsC†PëP»PμPSPμ
C†P»PëPSPsPIPëPrC-P»PμPSPSCḡ, PīPsC†PëP»PμPSPSCḡ PeP°C€P»Cḡ.

P PsP·P»P°PrPë P· P±PsPeCž C€PeC-CḡPë: PeCḡPsPīPëPIbḡ™ CḡPSPeP°, PiPμCḡPiPμC†,
C†PIPμCḡP±C-P¶, PīC-PrPIPëC%PμPSP° PīC-C,P»PëPIC-C†C,CHb, PIPëCḡP°P·PePë C€PeC-CḡPë,
PePsPSC,P°PeC,PSPëPNë PrPμCḡPjP°C,PëC,, PμPeC†C,,PsP»C-P°C,PëPIPsPëPNë PrPμCḡPjP°C,PëC,.

PŸPīPμC†PëC,,C-C†PSC- PIC-PrC†CžC,C,Cḡ: CḡP°PiC,P°PIPμ P·PSPëP¶PμPSPSCḡ C†Pë

PIC,C₇H₅P^oC,P^o C₇P>C₇C...C₇, P±C-P»CH C₇ PIC₇C...P^oC..., PeC₇PsPIPsPIPëP»PëPI C₇ PsPePs, PeP^oC,P^oC₇P^oPeC,P^o, C₇C₇C...C-C₇C,CH PI PsC±P^oC...

P— P±PsPeC₇ C₇C₇PsPiP_μPSC-C,P^oP»CH₅PSPsC— C₇PëC₇C,P_μPjPë: C₇PëC₇C,PëC,, PSC-PeC,C₇C₇C-C₇, P₇C-PrPIPëC%P_μPSP^o C±P^oC₇C,PsC,P^o C₇P_μC±PsPIPëP₇C₇C₇P^oPSCH₅, P·P±C-P»CH₅C₇P_μPSPSC₇ PjPsP»PsC±PSPëC... P·P^oP»PsP·, PSP_μC,C₇PëPjP^oPSPSC₇ C₇P_μC±C-, P₇PsC₇C₇C₇P_μPSPSC₇ P_μC₇PeC₇P»C₇C₇C-C-, PSP^oP±C₇C₇Pe C₇C,P^oC,P_μPIPëC... PsC₇PiP^oPSC-PI, P^oPSPsC₇PiP^oP·PjC-C₇.

P^oPsC₇PIC-Pr P·P^oC₇C,PsC₇C₇PiP^oPSPSC₇ P₇C-C₇P»C₇ PIPëC...PsPrC₇ PSP^o C₇PëPSPsPe.
P_μC-C₇P»C₇ PIPëC...PsPrC₇ P₇C₇P_μPiP^oC₇P^oC,C₇ PSP^o C₇PëPSPsPe P±C₇P»Pë C-PrP_μPSC,PëC,,C-PePsPIP^oPSC- PSPëP₇C₇P_μP·P^oP·PSP^oC±P_μPSC- P₇PsP±C-C±PSC- C₇P_μP^oPeC₇C-C-. P₇C₇P_μPeC-P»CH₅PePë P₇C₇Ps C,P^oPeC- C₇P_μP^oPeC₇C-C- P₇PsPIC-PrPsPjP»C₇C₇C,CH PrPsP±C₇PsPIC-P»CH₅PSPs C,P^o P₇PsPIC-PrPsPjP»P_μPSPSC₇ PSP^oPrC...PsPrC₇C,CH PIC-Pr P₇PsP₇C₇P»C₇C₇C-C- PSP_μPIC-PrPsPjPsC- C±PëC₇P_μP»CH₅PSPsC₇C-, PSP_μ P·P^oPIP₇PrPë PjPsP₇PSP^o PrPsC₇C,PsPIC-C₇PSPs PsC₇C-PSPëC,Pë C-C... C±P^oC₇C,PsC,C₇ C,P^o PIC₇C,P^oPSPsPIPëC,Pë P₇C₇PëC±PëPSPSPëP₇ P·P₇IB₇TM C₇P·PsPe C-P· P_μPeC₇P₇PsP·PëC₇C-C₇C P»C-PeP^oC₇C₇CH₅PePsPiPs P·P^oC₇PsP±C₇. P₇C- C₇PIPëC%P^o P±C₇P»Pë P·P^oP·PSP^oC±P_μPSC- C₇Pe C±P_μC₇P_μP· C-C... C₇P_μC₇P₇PsP·PSC-C₇C,CH, C±P^oC₇C,PsC,C₇ P₇PsPIC-PrPsPjP»P_μPSPSC₇, PIC-PrC₇C₇C,PSC-C₇C,CH C±C-C,PePsPiPs P^oP»CH₅C,P_μC₇PSP^oC,PëPIPSPsPiPs P·P₇IB₇TM C₇P·PeC₇, C,P^oPe C- C±P_μC₇P_μP· PePsPjP±C-PSP^oC₇C-C₇ C₇PëC... C,,P^oPeC,PsC₇C-PI.

P₇P_μC₇C₇P_μPIPs-C₇C₇PrPëPSPSC- C,P^o C₇P_μC₇P_μP±C₇PsPIP^oC₇PeC₇P»C₇C₇PSC- C₇PIPëC%P^o. P_μPsPIC-PrPsPjP»C₇P»PsC₇C₇ P₇C₇Ps C₇P_μC₇P₇PsP·PSC- C₇P_μC₇C₇P_μPIPs- C₇C₇PrPëPSPSC-, C₇P_μC₇P_μP±C₇PsPIP^oC₇PeC₇P»C₇C₇PSC- C,P^o C₇C₇PrPëPSPSC- C₇PIPëC%P^o, PIPeP»C₇C±P^oC₇C±Pë C₇P_μC₇P_μP±C₇PsPIP^oC₇PeC₇P»C₇C₇PSC₇ PeC₇PsPIPsC,P_μC±C₇, C₇C₇P±P^oC₇P^oC...PSPsC—PrP^oP»CH₅PSC₇ C,P^o PIPSC₇C,C₇C-C₇PSCH₅PsC₇P_μC₇P_μP±C₇P^oP»CH₅PSC₇ PeC₇PsPIPsC,P_μC±C₇ C,P^o P»P_μPiP_μPSP_μPIC₇ PeC₇PsPIPsC,P_μC±C₇, C%Ps P±C₇P»Pë P₇PsP₇IB₇TM C₇P·P^oPSC- C₇ C±P^oC₇C- C-P· P·P^oC₇C,PsC₇C₇PiP^oPSPSC₇Pj C₇PëP»PrP_μPSP^oC,,C-P»C₇. Pj P±C-P»CH₅C₇PsC₇C,C- P₇P^oC₇C-C₇PSC,C-PI P±C₇P»Pë PSP^oC₇PIPSC- C,,P^oPeC,PsC₇Pë C₇P_μC₇C₇P_μPIPs- C₇C₇PrPëPSPSPsPiPs C₇PëP·PëPeC₇. P_μPsPIC-PrPsPjP»C₇P»PsC₇C₇, C%Ps P±P^oPiP^oC,Ps P· C₇PëC... C₇PIPëC% PIPëPSPëPeP»Pë P₇C-Pr C±P^oC₇ P^oP±Ps PsPrC₇P^o·C₇ P₇C-C₇P»C₇ C₇P_μPeC₇C₇P^oP»CH₅PSPsC— P^oPeC,PëPIPSPsC₇C,C- C,P^o PrP_μPeC-P»CH₅PeP^o C₇PIPëC% PIPëPSPëPeP»Pë PsPrC₇P^o·C₇ P₇C-C₇P»C₇ P·P^oC₇C,PsC₇C₇PiP^oPSPSC₇ C₇PëP»PrP_μPSP^oC,,C-P»C₇ P±P_μP· C₇P_μPeC₇C₇P^oP»CH₅PSPsC— P^oPeC,PëPIPSPsC₇C,C-. P₇PSC₇C- C₇PIPëC%P^o PIPëPSPëPeP»Pë P₇C₇PsC,C₇PiPsPj PSP^oC₇C,C₇PiPSPëC... P₇PsPrPëPS C±Pë PrPSC-PI P₇C-C₇P»C₇ P·P^oC₇C,PsC₇C₇PiP^oPSPSC₇ C₇PëP»PrP_μPSP^oC,,C-P»C₇ C,P^o C₇P_μPeC₇C₇P^oP»CH₅PSPsC— P^oPeC,PëPIPSPsC₇C,C-. P₇P_μPjPsP₇P»PëPIPs PIC₇C,P^oPSPsPIPëC,Pë, C±Pë C₇C- C₇PIPëC%P^o P₇PsP₇IB₇TM C₇P·P^oPSC- C-P· P·P^oC₇C,PsC₇C₇PiP^oPSPSC₇Pj P₇C₇P_μPiP^oC₇P^oC,C₇, C-P· C₇P_μPeC₇C₇P^oP»CH₅PSPsC₇ P^oPeC,PëPIPSC-C₇C,C₇, C-P· PSP^oC₇PIPSPëPjPë C,,P^oPeC,PsC₇P^oPjPë C₇PëP·PëPeC₇ C±Pë C-P· PePsPjP±C-PSP^oC₇C-C₇C₇ C₇PëC... C,,P^oPeC,PsC₇C-PI, C±Pë P· C-PSC₇PePjPë C,,P^oPeC,PsC₇P^oPjPë.

P₇C₇PsPIPsPSPsC₇PSP^o C,P^o P»C-PjC,,P^oC,PëC±PSP^o C₇PëC₇C,P_μPjPë:
PIP^o·PsPsPeP»CH₅P·PëPIPSPëP₇ PeC₇PëP·. Pj PSP_μPIP_μP»PëPePsPjC₇ P·P^oPIC±P^oC₇PSPs P₇C₇PëP₇PëPSP_μPSPsPjC₇ PrPsC₇P»C-PrP₇P_μPSPSC- P·P^oC₇C,PsC₇C₇PiP^oPSPSC₇ P₇C₇P_μPiP^oC₇P^oC,C₇ C₇PëP»PrP_μPSP^oC,,C-P» P·P^o C₇C±P^oC₇C,C₇ P₇P^oC₇C-C₇PSC,C-PI P· P»P_μPiP_μPSP_μPIPsC₇ P^oC₇C,P_μC₇C-P^oP»CH₅PSPsC₇ P₇C-PiP_μC₇C,P_μPSP·C-C₇C₇, PIC,PsC₇PëPSPSPsC₇ C%PsPrPs C₇P_μC₇P₇PsP₇PsPrC-P±PSPsPeP»C-C,PëPSPSPsC—

P°PSPμPjC—, PīCḡPē P·P°CíC,PsCíCíPIP°PSPSC— CíPēP»PrPμPSP°C,,C—P»Cí PīCḡPs
CḡPsP·PIPēC,PsPe PIP°P·PsPsPeP»CḡP·PēPIPSPēC... PeCḡPēP·C—PI, C%Ps
PīPsC,CḡPμP±CíPIP°P»Pē PīPsCíPīC—C,P°P»C—P·P°C†C—, PīPsPIC—PrPsPjP»CḡP»PsCíCḡ
C†P°CíC,C—CēPμ, PSC—Pḡ PīCḡPē P·P°CíC,PsCíCíPIP°PSPSC— PīP»P°C†PμP±Ps.
PḡP»C—PSC—C†PSPμ P·PSP°C†PμPSPSCḡ C†C—C”C— C—PSC,,PsCḡPjP°C†C— PrP»Cḡ
PīP°C†C—C”PSC,C—PI, CḡPeC— P·P°CíC,PsCíCíPICíCḡC,Cḡ CíPēP»PrPμPSP°C,,C—P» P·
PjPμC,PsCḡ P»C—PeCíPIP°PSPSCḡ PμCḡPμPeC,PēP»CḡPSPsC— PrPēCíC,,CíPSPeC†C—C—,
PSPμPIC—PrPsPjPs.

P— P±PsPeCž PSPμCḡPIPSPiPsC— CíPēCíC,PμPjPē: C,CḡPēPIPSPiP°, C,CḡP°PSP·PēC,PsCḡPSP°
PiP»PsP±P°P»CḡPSP° P°PjPSPμP·C—Cḡ.

PŸPīPμC†PēC,,C—C†PSC— PIC—PrC†CžC,C,Cḡ

PŸP»CžC.... PμC—CíP»Cḡ PIPēC...PsPrCí PīCḡPμPīP°CḡP°C,Cí PSP° CḡPēPSPsPe
PīPsPIC—PrPsPjP»CḡP»PsCíCḡ PīCḡPs PIPēPīP°PrPePē CḡP°PīC,PsPIPSPiPs P·PSPēPḡPμPSPSCḡ
C†Pē PIC,CḡP°C,Pē CíP»CíC...Cí, PīPsPIBḡ™ CḡP·P°PSC— Cí C†P°CíC— C—P·
P·P°CíC,PsCíCíPIP°PSPSCḡPj CíPēP»PrPμPSP°C,,C—P»Cí. Pj PrPμCḡPePēC... PIPēPīP°PrPeP°C...
PīPsPIC—PrPsPjP»CḡP»PsCíCḡ PīCḡPs PSP°CḡPIPSC—CíC,Cḡ PjPμPrPēC†PSPēC... CíC,P°PSC—PI
C,P° C—PSCēPēC... C,,P°PeC,PsCḡC—PI, C%Ps PjPsPiP»Pē PIC—PrC—PīCḡP°C,Pē CḡPsP»Cḡ Cí
CḡPsP·PIPēC,PeCí PīPsP±C—C†PSPēC... CḡPμP°PeC†C—Pḡ P· P±PsPeCí CíP»CíC...Cí. Pj
P±P°PiP°C,CḡPsC... PIPēPīP°PrPeP°C... C—PSC,,PsCḡPjP°C†C—Cḡ C%PsPrPs
PīPsPrP°P»CḡCēPsPiPs PjPμPrPēC†PSPSPiPs CíPīPsCíC,PμCḡPμPḡPμPSPSCḡ
PIC—PrCíCíC,PSCḡ. P·PēP·PSP°C†PēC,Pē, C†Pē C†C— CḡPIPēC%P° PīCḡCḡPjPs
PīPsPIBḡ™ CḡP·P°PSC— C—P· P·P°CíC,PsCíCíPIP°PSPSCḡPj CíPēP»PrPμPSP°C,,C—P»Cí, P·
PSP°CḡPIPSPēPjPē C,,P°PeC,PsCḡP°PjPē CḡPēP·PēPeCí PIC,CḡP°C,Pē CíP»CíC...Cí, P·
PePsPjP±C—PSP°C†C—C”Cḡ C†PēC... C,,P°PeC,PsCḡC—PI C†Pē P· C—PSCēPēPjPē
C,,P°PeC,PsCḡP°PjPē, PSPμPjPsPḡP»PēPIPSP.

P—C—Cḡ. PŸPēPjC†P°CíPsPIP° PIC,CḡP°C,P° P·PsCḡCí, PīPsC†PμCḡPIPSPSC—PSPSCḡ PsC†PμPḡ,
PīPμC†C—PSPSCḡ PI PsC†P°C..., PīC—PrPIPēC%PμPSPSCḡ PIPSCíC,CḡC—CēPSCḡPsPsC†PSPSPiPs
C,PēCíPeCí, PSP°P±CḡCḡPe CíC—C,PeC—PIPePē, CíCíPrPēPSPSC—
P·P°C...PIPSPCḡPIP°PSPSCḡ CíC—C,PeC—PIPePē C†Pē PeCḡPsPIPSP,C,PμC†P°,
PIC—PrCēP°CḡCíPIP°PSPSCḡ CíPeP»PēCíC,PsPiPs C,C—P»P°.

PμC—CíP»Cḡ PIPēC...PsPrCí PīCḡPμPīP°CḡP°C,Cí PSP° CḡPēPSPsPe CḡC—PrPePs
PīPsPIC—PrPsPjP»CḡP»PsCíCḡ PīCḡPs PIPēPīP°PrPePē PSPμP°CḡC,PμCḡC—P°P»CḡPSPsC—
PīPμCḡPμPrPSCḡPsC— C—CēPμPjC—C†PSPsC— PSPμPICḡPsPīP°C,C—C— P·PsCḡPsPIPSPiPs
PSPμCḡPIP°, C%Ps C” PīCḡPēC†PēPSPsCḡ P·PSPēPḡPμPSPSCḡ P·PsCḡCí,
PIPeP»CḡC†P°CḡC†Pē PīPsCíC,C—PḡPSCí PIC,CḡP°C,Cí P·PsCḡCí, CḡPeC— P±CíP»Pē
PīPsPIBḡ™ CḡP·P°PSC— Cí C†P°CíC— C—P· P·P°CíC,PsCíCíPIP°PSPSCḡPj
C—PSPiC—P±C—C,PsCḡC—PI PαP”P°5, PIPeP»CḡC†P°CḡC†Pē CíPēP»PrPμPSP°C,,C—P». Pj
P±P°PiP°C,CḡPsC...B PīP°C†C—C”PSC,C—PI, P°P»Pμ PSPμ Cí PICíC—C..., P±CíP»Pē
PSP°CḡPIPSC— P°PSP°C,PsPjC—C†PSC— P°P±Ps CíCíPrPēPSPSC— C,,P°PeC,PsCḡPē CḡPēP·PēPeCí
CḡPsP·PIPēC,PeCí PSPμP°CḡC,PμCḡC—P°P»CḡPSPsC— PīPμCḡPμPrPSCḡPsC—
C—CēPμPjC—C†PSPsC— PSPμPICḡPsPīP°C,C—C— P·PsCḡPsPIPSPiPs PSPμCḡPIP°,
PIPeP»CḡC†P°CḡC†Pē (P°P»Pμ PSPμ PsP±PjPμPḡCíCḡC†PēCíCḡ) C,P°PeC—: PSPēP·CḡPePμ
CíPīC—PIPIC—PrPSPsCēPμPSPSCḡ PrC—P°PjPμC,CḡP° PμPeCíPeP°PIP°C†C—C— C,P° PrPēCíPeP°
P·PsCḡPsPIPSPiPs PSPμCḡPIP° (P·P°CíC,C—PḡPSPēPḡ PrPēCíPe P·PsCḡPsPIPSPiPs PSPμCḡPIP°),
PIC—Pe PīPsPSP°Pr 50 CḡPsPeC—PI, P°CḡC,PμCḡC—P°P»CḡPSP° PiC—PīPμCḡC,PμPSP·C—Cḡ,
P·P°C...PIPSPCḡPIP°PSPSCḡ PePsCḡPsPSP°CḡPSPēC... P°CḡC,PμCḡC—Pḡ,
PiC—PīPμCḡP»C—PiC—PrPμPjC—Cḡ C,P° PīP°P»C—PSPSCḡ. PkPμPjPsPḡP»PēPIPSP

PIPëP·PSP°C±PëC,Pë, C±Pë C†C- C¼PIPëC‰P° PïC½C¼PjPs PïPsPIB½™ C¼P·P°PSC- C-P·
P·P°CÍC,PsCÍCfPIP°PSPSC¼Pj C-PSPiC-P±C-C,PsC½C-PI PαP”P•5 C±Pë P· PSP°C¼PIPSPëPjPë
P°PSP°C,PsPjC-C±PSPëPjPë P°P±Ps CÍCfPrPëPSPSPëPjPë C,,P°PeC,PsC½P°PjPë C½PëP·PëPeCf,
C±Pë P· PePsPjP±C-PSP°C†C-C”C½ C†PëC... PICÍC-C... C,,P°PeC,PsC½C-PI, C±Pë P· C-PSC€PëPjPë
C,,P°PeC,PsC½P°PjPë.

P—PIC-C,C¿PIP°PSPSC¼ PïC½Ps PïC-PrPsP·C½C½PIP°PSC- PïPsP±C-C±PSC- C½PµP°PeC†C-C—.
P—PIC-C,CfPIP°PSPSC¼ PïC½Ps PïC-PrPsP·C½C½PIP°PSC- PïPsP±C-C±PSC- C½PµP°PeC†C-C—
PïC-CÍP»C¼ C½PµC”CÍC,C½P°C†C-C— P»C-PeP°C½CÍC½PePsPiPs P·P°CÍPsP±Cf PjP°C”
PIP°P¶P»PëPIPµ P·PSP°C±PµPSPSC¼. P¡Pµ PrP°C” P·PjPsPiCf PïC½PsPIPsPrPëC,Pë
P±PµP·PïPµC½PµC½PIPSPëPNë PjPsPSC-C,PsC½PëPSPi CÍPïC-PIPIC-PrPSPsC€PµPSPSC¼
PePsC½PëCÍC,C- C- C½PëP·PëPeC-PI, PïPsPIB½™ C¼P·P°PSPëC... C-P·
P·P°CÍC,PsCÍCfPIP°PSPSC¼Pj C†C½PsPiPs P»C-PeP°C½CÍC½PePsPiPs P·P°CÍPsP±Cf.
P>C-PeP°C½C¼Pj CÍP»C-Pr P·PIC-C,CfPIP°C,Pë PïC½Ps P±CfPrC½-C¼PeC-
PïC-PrPsP·C½C½PIP°PSC- PïPsP±C-C±PSC- C½PµP°PeC†C-C— PIC-PrPïPsPIC-PrPSPs PrPs
PIPëPjPsPi P·P°PePsPSPsPrP°PICÍC,PIP°.

PŷPµC½PjC-PS PïC½PëPrP°C,PSPsCÍC,C-. 3 C½PsPePë.

PJPjPsPIPë P·P±PµC½C-PiP°PSPSC¼.

P—P±PµC½C-PiP°C,Pë PïC½Pë C,PµPjPïPµC½P°C,CfC½C- PSPµ PIPëC‰Pµ 25 B°Pŷ.

P—P±PµC½C-PiP°C,Pë Cf PSPµPrPsCÍC,CfPïPSPsPjCf PrP»C¼ PrC-C,PµPNë PjC-CÍC†C-.

PJPïP°PePsPIPeP°.

PµPs 1 P°P±Ps 4 C,P°P±P»PµC,PePë Cf P±P»C-CÍC,PµC½C-. PµPs PsPrPSPsPjCf
P±P»C-CÍC,PµC½Cf Cf PeP°C½C,PsPSPSC-PNë PïP°C†C†C-.

PљP°C,PµPiPsC½C-C¼ PIC-PrPïCfCÍPeCf.

P—P° C½PµC†PµPïC,PsPj.

P’PëC½PsP±PSPëPe.

P’PµP»CfPïPs, P»C-PePë C,P° PePsCÍPjPµC,PëPeP°, Pr.Pr.

**PњC-CÍC†PµP·PSP°C...PsPrP¶PµPSPSC¼ PIPëC½PsP±PSPëPeP° C,P° P°PrC½PµCÍP°
PjC-CÍC†C¼ PïC½PsPIP°PrP¶PµPSPSC¼ PNëPsPiPs PrC-C¼P»C½PSPsCÍC,C-.**

P’CfP». P”P°PSC-C†P° 5, 48000 PљPsPïC½PëPIPSPëC†C¼, PÍPsC½PIP°C,C-C¼.