

*PhCl'PSPsPIPSC- C,,C-P·PëPePs-C...C-PjC-C±PSC- PIP»P°Cl'C,PëPIPsCl'C,C-: C,PIPµCḡPrC-  
P¶PµP»P°C,PëPSPsPIC- PeP°PiCl'CrP»Pë P· PePsCḡPiCrCl'PsPj PsCḡP°PSP¶PµPIPsPiPs  
PePsP»CHbPsCḡCr C,P° PeCḡPëC€PµC±PePsCḡ PsP»PëPIPePsPIPsPiPs PePsP»CHbPsCḡCr. P'  
PeP°PiCl'CrP»C- PjC-Cl'C,CḡC,CHbCl'Cl PīPµP»PµC,Pë P±C-P»PsPiPs P°P±Ps PjP°PN₂P¶Pµ  
P±C-P»PsPiPs PePsP»CHbPsCḡCr.*

$P\hbar P \pm C\dot{I}P_s C\dot{H}P \pm C\dot{I}C - C\dot{I}I$ .  $P\ddot{Y}P^\circ P_j C\dot{I}C\dot{r}P \gg P_s P \cdot P\ddot{e}PS$   $C \in PIP\ddot{e}PrPePs$   $C, P^\circ$   $P_j P^\circ P N_{\mathbb{P}} P \P P_{\mu}$   
 $P\ddot{i}PsPIPSC - C\dot{I}C, C\dot{H}$   $PI C\dot{I}P_j PsPeC, C\dot{r}C''C, C\dot{H}C\dot{I}C\dot{I} P \cdot$   $PeP\ddot{e}C\dot{e}P_{\mu} C\dot{I}PSP\ddot{e}PeC\dot{r}$ .  
 $P' C\dot{I}P_j PsPeC, C\dot{r}PIP^\circ PSPSC\dot{I}$   $C\dot{r}P\ddot{i}PsPIC - P \gg C\dot{H}PSC\dot{H}C''C, C\dot{H}C\dot{I}C\dot{I}$ ,  $C\dot{I}PeC\%Ps$   $P\ddot{i}C\dot{H}P_{\mu} P\ddot{i}P^\circ C\dot{H}P^\circ C,$   
 $P \cdot P^\circ C\dot{I}C, PsC\dot{I}PsPIC\dot{r}PIP^\circ P \gg P\ddot{e}$   $P\ddot{i}C - C\dot{I}P \gg C\dot{I} C - PrP\ddot{e}$ .  $P$   $C - PIPSPsP_j C - C\dot{H}PSPsPiPs$   
 $PIP\ddot{e}PIC - P \gg C\dot{H}PSP_{\mu}PSPSC\dot{I}$   $PrC - C\dot{H}C\dot{I}PsC -$   $C\dot{H}P_{\mu} C\dot{I}PsPIP\ddot{e}PSP\ddot{e}$   $P_j PsP \P PSP^\circ$   $PrPsC\dot{I}C\dot{I}PiC, P\ddot{e},$   
 $C\dot{I}PeC\%Ps$   $P\ddot{i}C\dot{H}P\ddot{e}PN_{\mathbb{P}} P_j P^\circ C, P\ddot{e}$   $C, P^\circ P_j C\dot{I}C\dot{r}P \gg P_s P \cdot P\ddot{e}PS$   $PI$   $PsPrP\ddot{e}PS$   $C -$   $C, PsPN_{\mathbb{P}}$   $C\dot{I}P^\circ P_j P\ddot{e}PN_{\mathbb{P}}$   
 $C\dot{I}P^\circ C\dot{I}$   $P\ddot{i}C - C\dot{I}P \gg C\dot{I}$   $P\ddot{i}C\dot{H}P\ddot{e}PN_{\mathbb{P}} PsP_j C\dot{r} C - P \P C -$ .  $P\ddot{Y}P^\circ P_j C\dot{I}C\dot{r}P \gg P_s P \cdot P\ddot{e}PSC\dot{r}$   $PIP \gg P^\circ C\dot{I}C, P\ddot{e}PIP^\circ$   
 $P \gg C - PSC - PN_{\mathbb{P}}PSP^\circ$   $PeC - PSP_{\mu} C, P\ddot{e}PeP^\circ$ .  $P_{\mu} C - PePsPIC -$   $C\dot{H}C - PIPSC -$   $PI$   $P\ddot{i}P \gg P^\circ P \cdot P_j C -$   $PeC\dot{H}PsPIC -$

PrPsCfCfPiP°CfC,ChCfCfCf PiCfPëP±P»PëP·PSPs C±PµCfPµP· 6 PiPsPrPëPS PiC-CfP»CfC  
 PsPrPSPsCfP°P·PsPIPsC— PrPsP·Pë C,P°PjCfCfP»PsP·PëPSCf, CfPeCf PiCfPëPNëPSCfP»Pë  
 PiC-CfP»CfC C—PrPë. PŸC,P°P±C-P»ChPSP° PePsPSCfPµPSC,CfP°CfC-CfC  
 PrPsCfCfPiP°C”C,ChCfCfCf PSP° 5 PrPsP±Cf PiC-CfP»CfC C%PsPrPµPSPSPsPiPs  
 PiCfPëPNëPsPjCf PiCfPµPiP°CfP°C,Cf, PiCfPë CfChPsPjCf PjP°PeCfPëPjP°P»ChPSP°  
 PePsPSCfPµPSC,CfP°CfC-CfC PI PiP»P°P·PjC- PeCfPµPIC- PSP° <sup>2</sup>/<sub>3</sub> PIPëC%P°, PSC-P  
 PiC-CfP»CfC PiCfPëPNëPsPjCf CfP°P·PsPIPsC— PrPsP·Pë. PfPsC±P° CfC- PrP°PSC-  
 CfC,PsCfCfCfC,ChCfCfCf C,C-P»ChPePë PiP°CfC-C”PSC,C-PI P»C-C,PSChPsPiPs PIC-PeCf,  
 PjPsP¶PSP° PsC±C-PeCfPIP°C,Pë P°PSP°P»PsPiC-C±PSPëPNë CfPµP·CfP»ChC,P°C, Cf  
 PiP°CfC-C”PSC,C-PI PjPsP»PsPrCfPsPiPs PIC-PeCf.

P C-PIPSC- C,P°PjCfCfP»PsP·PëPSCf PI PiP»P°P·PjC- PeCfPµPIC- Cf CfC-P·PSPëC...  
 PiP°CfC-C”PSC,C-PI PjPsP¶CfC,Ch CfCfC,C,C”PIPs PIC-PrCfC-P·PSCfC,PëCfCfC CfPe  
 PiC-CfP»CfC CfP°P·PsPIPsC— PrPsP·Pë, C,P°Pe C- PiC-CfP»CfC P±P°PiP°C,PsCfP°P·PsPIPsPiPs  
 P·P°CfC,PsCfCfPIP°PSPSCfC.

*P PsP·PiPsPrC-P».* P—PIbŸ™ CfP·CfPIP°PSPSCfC P· P±C-P»PeP°PjPë PiP»P°P·PjPë PeCfPµPIC- bŸ”  
 99 %. PhP±bŸ™ C”Pj CfPµP·PiPsPrC-P»Cf PSPµP·PSP°C±PSPëPNë (PiCfPëP±P»PëP·PSPs  
 0,2 P»/PePi).

*PµPµC,P°P±PsP»C-P·Pj.* PŸP°PjCfCfP»PsP·PëPS PSPµ PiC-PrPrP°C”C,ChCfCfCf PµC,,PµPeC,Cf  
 PiPµCfCfPsPiPs PiCfPµC...PsPrP¶PµPSPSCfC C- PiPsPIC-P»ChPSPs  
 PjPµC,P°P±PsP»C-P·CfC”C,ChCfCfCf PI PiPµC±C-PSCfC- P· CfC,PIPsCfPµPSPSCfCfC  
 C,,P°CfPjP°PePsP»PsPiC-C±PSPs P°PeC,PëPIPSPëC... PjPµC,P°P±PsP»C-C,C-PI, C%Ps  
 P·P±PµCfC-PiP°CfC,Ch PIPëCfPµPeCf CfPµP»PµPeC,PëPIPSC-CfC,Ch PrPs α<sub>1</sub>-  
 P°PrCfPµPSPsCfPµCfPµPiC,PsCfC-PI. P’C-P»ChCfP° C±P°CfC,PëPSP° P°PeC,PëPIPSPsC—  
 CfPµC±PsPIPëPSPë PiCfPëCfCfC,PSCfC PI PeCfPµPIC- Cf PSPµP·PjC-PSPµPSPsPjCf  
 PIPëPiP»CfPrC-.

*P’PëPIPµPrPµPSPSCfC.* PŸP°PjCfCfP»PsP·PëPS C- PNëPsPiPs PjPµC,P°P±PsP»C-C,Pë  
 PIPëPIPsPrCfC,ChCfCfCf PiPsP»PsPIPSPëPj C±PëPSPsPj C-P· CfPµC±PµCf, PiCfPëC±PsPjCf  
 PiCfPëP±P»PëP·PSPs 9 % PrPsP·Pë PIPëPIPsPrPëC,ChCfCfCf PI PSPµP·PjC-PSPµPSPsPjCf  
 PIPëPiP»CfPrC-.

PµC-CfP»CfC CfP°P·PsPIPsPiPs PiCfPëPNëPsPjCf PrPsP·Pë C,P°PjCfCfP»PsP·PëPSCf PiC-CfP»CfC  
 C—PrPë C,P° PiCfPë CfC,P°P±C-P»ChPSC-PNë PePsPSCfPµPSC,CfP°CfC-C— Cf PiP»P°P·PjC-  
 PeCfPµPIC- PiPµCfC-PsPrPë PSP°PiC-PIIPëPIPµPrPµPSPSCfC PIC-PrPiPsPIC-PrPSPs  
 CfC,P°PSPsPIP»CfC,Ch PiCfPëP±P»PëP·PSPs 10 C,P° 13 PiPsPrPëPS.

**PµP»C-PSC-C±PSC- C...P°CfP°PeC,PµCfPëCfC,PëPePë.**

**PµPsPeP°P·P°PSPSCfC.**

P>C-PeCfPIP°PSPSCfC C,,CfPSPeCfC-PsPSP°P»ChPSPëC... CfPµP·P»P°PrC-PI P· P±PsPeCf  
 PSPëP¶PSC-C... CfPµC±PsPIPëPIC-PrPSPëC... CfP»CfC...C-PI PiCfPë  
 PrPsP±CfPµCfPeC-CfPSC-PNë PiC-PiPµCfPiP»P°P·C-C— PiCfPµCfC,P°C,Pë.

**PµCfPµC,PëPiPsPeP°P·P°PSPSCfC.**

B.  $P^{\circ}C - \text{PiP}_{\mu}C\bar{H}C\bar{C}rC, P \gg \text{P}\bar{e}PIC - C\bar{r}C, CH \text{ PrPs } C, P^{\circ}PjC\bar{r}C\bar{r}P \gg \text{PsP} \cdot \text{P}\bar{e}PSC\bar{r}$   
 $PiC - \text{Pr}C\bar{H}PsC...P \gg \text{Ps}C\bar{H}P\bar{e}PrC\bar{r}, \text{PIPeP} \gg C\bar{H}C\bar{P}^{\circ}C\bar{H}C\bar{P}\bar{e} PjP_{\mu}PrP\bar{e}PeP^{\circ}PjP_{\mu}PSC, \text{PsP} \cdot \text{PSPS-}$   
 $C - \text{PSPrC}\bar{r}PePsPIP^{\circ}PSP\bar{e}PN_{\bar{e}} P^{\circ}PSPiC - \text{PsPSP}_{\mu}PIC\bar{H}PsC, P\bar{e}C\bar{P}SP\bar{e}PN_{\bar{e}} \text{PSP}^{\circ}P \pm C\bar{H}C\bar{I}Pe, P^{\circ}P \pm \text{Ps PrPs}$   
 $P \pm C\bar{r}PrCH - C\bar{I}PePsC - P \cdot \text{PrPsPiPsPjC} - P\bar{P}SP\bar{e}C... C\bar{H}P_{\mu}C\bar{P}PsPIP\bar{e}PS;$

B.  $\text{Ps}C\bar{H}C, \text{Ps}C\bar{r}C, P^{\circ}C, P\bar{e}C\bar{P}SP^{\circ} PiC - \text{PiPsC}, P_{\mu}PSP \cdot C - C\bar{I} PI P^{\circ}PSP^{\circ}PjPSP_{\mu}P \cdot C-;$

B.  $C, C\bar{I}P\bar{P}PeP^{\circ} \text{PiP}_{\mu}C\bar{P}C - \text{PSPePsPIP}^{\circ} \text{PSP}_{\mu}PrPsC\bar{r}C, P^{\circ}C, \text{PSC} - C\bar{r}C, CH.$

**$P^{\circ}P \cdot P^{\circ}C^{\circ}PjPsPrC - C\bar{I} P \cdot C - \text{PSC}\bar{e}P\bar{e}PjP\bar{e} P \gg C - \text{PeP}^{\circ}C\bar{H}C\bar{r}CHPeP\bar{e}PjP\bar{e} P \cdot P^{\circ}C\bar{r}PsP \pm P^{\circ}PjP\bar{e} C, P^{\circ}$**   
 **$C - \text{PSC}\bar{e}C - \text{PIP}\bar{e}PrP\bar{e} PIP \cdot P^{\circ}C^{\circ}PjPsPrC - PN_{\bar{e}}.$**

$P^{\circ}PsC\bar{r}P \gg C - \text{PrP}\bar{P}_{\mu}PSPSC\bar{I} PIP \cdot P^{\circ}C^{\circ}PjPsPrC - C - \text{Pi}C\bar{H}PsPIP\bar{e}PrP\bar{e}P \gg \text{P}\bar{e}C\bar{r}C\bar{I} C, C - P \gg CHPeP\bar{e} P \cdot P^{\circ}$   
 $C\bar{r}C\bar{P}^{\circ}C\bar{r}C, CH \text{ PrPs}C\bar{H}PsC\bar{r}P \gg \text{P}\bar{e}C...$

$P_{\mu}C\bar{H}P\bar{e} \text{PsPrPSPsC}\bar{P}^{\circ}C\bar{r}PSPsPjC\bar{r} P \cdot P^{\circ}C\bar{r}C, \text{Ps}C\bar{r}C\bar{r}PIP^{\circ}PSPSC - C, P^{\circ}PjC\bar{r}C\bar{r}P \gg \text{PsP} \cdot \text{P}\bar{e}PSC\bar{r}$   
 $PiC - \text{Pr}C\bar{H}PsC...P \gg \text{Ps}C\bar{H}P\bar{e}PrC\bar{r} P \cdot P^{\circ}C, P_{\mu}PSPsP \gg \text{PsP} \gg \text{PsPj}, P_{\mu}PSP^{\circ}P \gg P^{\circ}PiC\bar{H}P\bar{e}P \gg \text{PsPj},$   
 $PSC - C, P_{\mu}PrP\bar{e}PiC - \text{PSPsPj}, PrP\bar{e}PiPsPeC\bar{r}P\bar{e}PSPsPj P^{\circ}P \pm \text{Ps } C, P_{\mu}PsC, C - P \gg C - \text{PSPsPj}$   
 $P \gg C - \text{PeP}^{\circ}C\bar{H}C\bar{r}CHPePsC - PIP \cdot P^{\circ}C^{\circ}PjPsPrC - C - \text{PSP}_{\mu} PIC - \text{PrP} \cdot \text{PSP}^{\circ}C\bar{P}^{\circ}P \gg \text{Ps}C\bar{r}C\bar{I}.$   
 $PhPrPSPsC\bar{P}^{\circ}C\bar{r}PSP_{\mu} P \cdot P^{\circ}C\bar{r}C, \text{Ps}C\bar{r}C\bar{r}PIP^{\circ}PSPSC\bar{I} P \cdot C\bar{r}P\bar{e}PjP_{\mu}C, P\bar{e}PrP\bar{e}PSPsPj$   
 $PiC - \text{PrPIP}\bar{e}C\bar{r}C^{\circ}, P^{\circ} P \cdot C, C\bar{r}C\bar{H}PsC\bar{r}P_{\mu}PjC - \text{PrPsPj} - P \cdot \text{PSP}\bar{e}P\bar{P}C\bar{r}C^{\circ}$   
 $PePsPSC\bar{r}P_{\mu}PSC, C\bar{H}P^{\circ}C\bar{r}C - CH C, P^{\circ}PjC\bar{r}C\bar{r}P \gg \text{PsP} \cdot \text{P}\bar{e}PSC\bar{r} PI \text{PiP} \gg P^{\circ}P \cdot PjC - \text{Pe}C\bar{H}PsPIC-, P^{\circ}P \gg P_{\mu}$   
 $\text{Ps}C\bar{r}PeC - P \gg CHPeP\bar{e} C\bar{r}C - C\bar{H}C - \text{PIPSC} - P \cdot P^{\circ}P \gg \text{P}\bar{e}C\bar{e}P^{\circ}CHC, CHC\bar{r}C\bar{I} PI PjP_{\mu}P\bar{P}^{\circ}C... PSPsC\bar{H}PjP\bar{e},$   
 $C\bar{r} C\bar{r}PiP_{\mu}C\bar{r}C - P^{\circ}P \gg CHPSC - PN_{\bar{e}} \text{PePs}C\bar{H}P_{\mu}PeC\bar{r}C - C - \text{PrPsP} \cdot C\bar{r}PIP^{\circ}PSPSC\bar{I}$   
 $C, P^{\circ}PjC\bar{r}C\bar{r}P \gg \text{PsP} \cdot \text{P}\bar{e}PSC\bar{r} \text{PSP}_{\mu}PjP^{\circ}C^{\circ} \text{PiPsC}, C\bar{H}P_{\mu}P \pm P\bar{e}.$

$Pj \text{PrPs}C\bar{r}P \gg C - \text{PrP}\bar{P}_{\mu}PSPSC\bar{I}C... \text{in vitro } PrC - P^{\circ}P \cdot P_{\mu}PiP^{\circ}Pj, PiC\bar{H}PsPiC\bar{H}P^{\circ}PSPsP \gg \text{PsP} \gg,$   
 $C, C\bar{H}P\bar{e}C...P \gg \text{Ps}C\bar{H}PjP_{\mu}C, C - P^{\circ}P \cdot P\bar{e}Pr, C...P \gg \text{Ps}C\bar{H}PjP^{\circ}PrP\bar{e}PSPsPS, P^{\circ}PjC - C, C\bar{H}P\bar{e}PiC, P\bar{e}P \gg C - \text{PS},$   
 $PrP\bar{e}PeP \gg \text{PsC}, P_{\mu}PSP^{\circ}Pe, PiP \gg C - P \pm P_{\mu}PSPeP \gg P^{\circ}PjC - \text{Pr}, C\bar{r}P\bar{e}PjPIP^{\circ}C\bar{r}C, P^{\circ}C, P\bar{e}PS C, P^{\circ}$   
 $PIP^{\circ}C\bar{H}C, P^{\circ}C\bar{H}P\bar{e}PS \text{PSP}_{\mu} PIPiP \gg P\bar{e}PIP^{\circ}P \gg \text{P}\bar{e} \text{PSP}^{\circ} PIC - P \gg CHPSC\bar{r} C, C\bar{H}P^{\circ}PeC\bar{r}C - CH$   
 $C, P^{\circ}PjC\bar{r}C\bar{r}P \gg \text{PsP} \cdot \text{P}\bar{e}PSC\bar{r} PI \text{PiP} \gg P^{\circ}P \cdot PjC - \text{Pe}C\bar{H}PsPIC - P \gg CHPrP\bar{e}PSP\bar{e}.$   
 $P_{\mu}PsPrC - P \pm \text{PSP}\bar{e}Pj C\bar{P}\bar{e}PSPsPj C, P^{\circ}PjC\bar{r}C\bar{r}P \gg \text{PsP} \cdot \text{P}\bar{e}PS \text{PSP}_{\mu} P \cdot PjC - \text{PSC}\bar{H}PIP^{\circ}PI$   
 $C\bar{H}C - \text{PIP}_{\mu}PSC\bar{H} PIC - P \gg CHPSP\bar{e}C... C, C\bar{H}P^{\circ}PeC\bar{r}C - PN_{\bar{e}} PrC - P^{\circ}P \cdot P_{\mu}PiP^{\circ}PjC\bar{r},$   
 $PiC\bar{H}PsPiC\bar{H}P^{\circ}PSPsP \gg \text{PsP} \gg C\bar{r}, C, C\bar{H}P\bar{e}C...P \gg \text{Ps}C\bar{H}PjP_{\mu}C, C - P^{\circ}P \cdot P\bar{e}PrC\bar{r} C, P^{\circ}$   
 $C...P \gg \text{Ps}C\bar{H}PjP^{\circ}PrP\bar{e}PSPsPSC\bar{r} PI \text{PiP} \gg P^{\circ}P \cdot PjC - \text{Pe}C\bar{H}PsPIC - P \gg CHPrP\bar{e}PSP\bar{e}. P_{\mu}C\bar{H}PsC, P_{\mu}$   
 $PrP\bar{e}PeP \gg \text{PsC}, P_{\mu}PSP^{\circ}Pe C, P^{\circ} PIP^{\circ}C\bar{H}C, P^{\circ}C\bar{H}P\bar{e}PS PjPsP\bar{P}C\bar{r}C, CH$   
 $PiC\bar{H}P\bar{e}C\bar{r}PePsC\bar{H}PIP^{\circ}C, P\bar{e} P_{\mu}P \gg C - PjC - \text{PSP}^{\circ}C\bar{r}C - CH C, P^{\circ}PjC\bar{r}C\bar{r}P \gg \text{PsP} \cdot \text{P}\bar{e}PSC\bar{r} P \cdot$   
 $PiP \gg P^{\circ}P \cdot PjP\bar{e} \text{Pe}C\bar{H}PsPIC-.$

$PhPrPSPsC\bar{P}^{\circ}C\bar{r}PSP_{\mu} P \cdot P^{\circ}C\bar{r}C, \text{Ps}C\bar{r}C\bar{r}PIP^{\circ}PSPSC\bar{I} P \cdot C\bar{r}P\bar{e}P \gg CHPSP\bar{e}PjP\bar{e}$   
 $C - \text{PSPiC} - P \pm C - C, \text{Ps}C\bar{H}P^{\circ}PjP\bar{e} \text{CYP3A4 } PjPsP\bar{P}_{\mu} PiC\bar{H}P\bar{e}P \cdot \text{PIP}_{\mu}C\bar{r}C, P\bar{e} \text{PrPs}$   
 $P \cdot P \pm C - P \gg CHC\bar{e}P_{\mu}PSPSC\bar{I} PIPiP \gg P\bar{e}PIC\bar{r} C, P^{\circ}PjC\bar{r}C\bar{r}P \gg \text{PsP} \cdot \text{P}\bar{e}PSC\bar{r} PiC - \text{Pr}C\bar{H}PsC...P \gg \text{Ps}C\bar{H}P\bar{e}PrC\bar{r}.$   
 $P - P^{\circ}C\bar{r}C, \text{Ps}C\bar{r}C\bar{r}PIP^{\circ}PSPSC\bar{I} P \cdot \text{PeP}_{\mu}C, \text{PsPePsPSP}^{\circ}P \cdot \text{PsP} \gg \text{PsPj} (\text{PIC} - \text{PrPsPjP\bar{e}Pj } C\bar{r}P\bar{e}P \gg CHPSP\bar{e}Pj$   
 $C - \text{PSPiC} - P \pm C - C, \text{Ps}C\bar{H}PsPj \text{CYP3A4}) PiC\bar{H}P\bar{e}P \cdot \text{PIPsPrP\bar{e}C}, CH \text{ PrPs } P \cdot P \pm C - P \gg CHC\bar{e}P_{\mu}PSPSC\bar{I} \text{AUC}$   
 $C - C_{\max} C, P^{\circ}PjC\bar{r}C\bar{r}P \gg \text{PsP} \cdot \text{P}\bar{e}PSC\bar{r} PiC - \text{Pr}C\bar{H}PsC...P \gg \text{Ps}C\bar{H}P\bar{e}PrC\bar{r} PI 2,8 C, P^{\circ} 2,2 C\bar{H}P^{\circ}P \cdot P^{\circ}$   
 $PIC - \text{PrPiPsPIC} - \text{PrPSPs}.$

$P\bar{y}P^{\circ}PjC\bar{r}C\bar{r}P \gg \text{PsP} \cdot \text{P}\bar{e}PSC\bar{r} PiC - \text{Pr}C\bar{H}PsC...P \gg \text{Ps}C\bar{H}P\bar{e}Pr C\bar{r}P \gg C - \text{Pr } P \cdot P^{\circ}C\bar{r}C, \text{Ps}C\bar{r}PsPIC\bar{r}PIP^{\circ}C, P\bar{e} P \cdot$   
 $\text{PsP} \pm P_{\mu}C\bar{H}P_{\mu}P\bar{P}PSC - C\bar{r}C, CH C\bar{r} \text{PePsPjP} \pm C - \text{PSP}^{\circ}C\bar{r}C - C - P \cdot C\bar{r}P\bar{e}P \gg CHPSP\bar{e}PjP\bar{e}$

(PSP°PīCḡPēPeP»P°Pr PePμC,PsPePsPSP°P·PsP»PsPj, C-C,CḡP°PePsPSP°P·PsP»PsPj, PIPsCḡPēPePsPSP°P·PsP»PsPj, PeP»P°CḡPēC,CḡPsPjC-C†PēPSPsPj, C-PSPrPēPSP°PIC-CḡPsPj, PSPμP»C,,C-PSP°PIC-CḡPsPj, CḡPēC,PsPSP°PIC-CḡPsPj, C†P°PePIC-PSP°PIC-CḡPsPj) C-PīPsPjC-CḡPSPēPjPē (PSP°PīCḡPēPeP»P°Pr PμCḡPēC,CḡPsPjC-C†PēPSPsPj) C-PSPiC-P±C-C,PsCḡP°PjPē CYP3A4. Pḡ PīP°C†C-C"°PSC,P°Pj P·PSPēP·CHPePēPj PjPμC,P°P±PsP»C-P·PjPsPj CYP2D6 C,P°PjC†C†P»PsP·PēPSC† PiC-PrCḡPsC...P»PsCḡPēPr PSPμ C†P»C-Pr PīCḡPēP·PSP°C†P°C,Pē C† PīPsC"°PrPSP°PSPSC- P·C†PēP»CHPSPēPjPē C-PSPiC-P±C-C,PsCḡP°PjPē CYP3A4.

PḡPrPSPsC†P°C†PSPμ P·P°C†C,PsC†C†PIP°PSPSC† P·PīP°CḡPsPeC†PμC,PēPSPsPj (C†PēP»CHPSPēPNḡ C-PSPiC-P±C-C,PsCḡ PŸYP2D6) PīCḡPēP·PIPPrPēC,CH PrPs PiC-PrPIPēC%PμPSPSC† PePsPSC†PμPSC,CḡP°C†C-C— PŸ<sub>max</sub> C- AUC C,P°PjC†C†P»PsP·PēPSC† PI 1,3 C- 1,6 CḡP°P·P° PIC-PrPīPsPIC-PrPSPs, P°P»Pμ C†Pμ PiC-PrPIPēC%PμPSPSC† PSPμ PIPIP°P¶P°C"°C,CHC†C† PeP»C-PSC-C†PSPs P·PSP°C†C†C%PēPj.

P†C†PSC†C"° CḡPēP·PēPe PīPsC†PēP»PμPSPSC† PiC-PīPsC,PμPSP·PēPIPSPsPiPs PμC,,PμPeC,C† PīCḡPē P·P°C†C,PsC†C†PIP°PSPSC- C,P°PjC†C†P»PsP·PēPSC† PsPrPSPsC†P°C†PSPs P·PīCḡPμPīP°CḡP°C,P°PjPē, C†PeC- PjPsP¶C†C,CH P·PSPēP¶C†PIP°C,Pē PeCḡPsPiBḡ™ C†PSPēPNḡ C,PēC†Pe, PI C,,C†. P·P°PSPμC†C,PμC,PēPeP°PjPē, C-PSPiC-P±C-C,PsCḡP°PjPē C,,PsC†C,,PsPrC-PμC†C,PμCḡP°P·Pē-5, C-PSC€PēPjPē α<sub>1</sub>-P°PrCḡPμPSPsP±P»PsPeP°C,PsCḡP°PjPē.

P"°PSC†P»C-PrP¶PμPSPSC† in vitro P·PjC-PeCḡPsC†PsPjP°P»CHPSPēPjPē C,,CḡP°PeC†C-C†PjPē PīPμC†C-PSPePē PSPμ PīPsPeP°P·P°P»Pē PIP·P°C"°PjPsPrC-C— P·P°PjC-C,CḡPēPīC,PēP»C-PSPsPj, C†P°P»CHP±C†C,P°PjPsP»PsPj, PiP»C-P±PμPSPeP»P°PjC-PrPsPj C,P° C,,C-PSP°C†C,PμCḡPēPrPsPj.

### **PḡC†PsP±P»PēPIPSC†C,C- P·P°C†C,PsC†C†PIP°PSPSC†.**

PīPe C- PīCḡPē P·P°C†C,PsC†C†PIP°PSPSC- C-PSC€PēC... α<sub>1</sub>- P°PrCḡPμPSPsP±P»PsPeP°C,PsCḡC-PI, PI PsPeCḡPμPjPēC... PIPēPīP°PrPeP°C... PīCḡPē P·P°C†C,PsC†C†PIP°PSPSC- PŸP°PjC†C†P»C-PrC† PjPsP¶P»PēPIPμ P·PSPēP¶PμPSPSC† P°CḡC,PμCḡC-P°P»CHPSPsPiPs C,PēC†PeC†, C%Ps PjPsP¶Pμ C-PSPsPrC- PīCḡPēP·PIPμC†C,Pē PrPs PIC,CḡP°C,Pē C†PIC-PrPsPjPsC†C,C-. PḡCḡPē PīPsC†PIC- PīPμCḡC€PēC... PsP·PSP°Pe PsCḡC,PsC†C,P°C,PēC†PSPsC— PiC-PīPsC,PμPSP·C-C— (P·P°PīP°PjPsCḡPsC†PμPSPSC†, C†P»P°P±PeC-C†C,CH) PīP°C†C-C"°PSC, PīPsPIPēPSPμPS PīCḡPēPNḡPSC†C,Pē PīPsCḡPēP·PsPSC,P°P»CHPSPμ PīPsP»PsP¶PμPSPSC† PrPs P·PSPēPePSPμPSPSC† PIPēC%PμP·PīP°PrP°PSPēC... C†PēPjPiC,PsPjC-PI.

PḡPμCḡPμPr C,PēPj C†Pe CḡPsP·PīPsC†P°C,Pē P»C-PeC†PIP°PSPSC† PīCḡPμPīP°CḡP°C,PsPj PŸP°PjC†C†P»C-Pr, C†P»C-Pr PīCḡPsPNḡC,Pē PjPμPrPēC†PSPμ PsP±C†C,PμP¶PμPSPSC† P·PjPμC,PsCḡ PIPēC†PIP»PμPSPSC† C-PSC€PēC... C†C†PīC†C,PSC-C... P·P°C...PIPSCḡCḡPIP°PSCH, C%Ps PjPsP¶C†C,CH C†PīCḡPēC†PēPSPēC,Pē C,P°PeC- P¶C†P°PjC- C†PēPjPīC,PsPjPē, C†Pe PrPsP±CḡPsC†PeC-C†PSP° PiC-PīPμCḡPīP»P°P·C-C† PīPμCḡPμPrPjC-C...C†CḡPsPIPSC— P·P°P»PsP·Pē.

PḡPμCḡPμPr PīPsC†P°C,PePsPj P»C-PeC†PIP°PSPSC† PSPμPsP±C...C-PrPSPs PīCḡPsPIPμC†C,Pē CḡPμPeC,P°P»CHPSPμ PsP±C†C,PμP¶PμPSPSC† PīPμCḡPμPrPjC-C...C†CḡPsPIPSC— P·P°P»PsP·Pē C,P° P·P° PSPμPsP±C...C-PrPSPsC†C,C- PrPs PīPsC†P°C,PeC† C,P° CḡPμPiC†P»C†CḡPSPs PīC-Pr C†P°C† P»C-PeC†PIP°PSPSC† PePsPSC,CḡPsP»CḡPIP°C,Pē

$\text{C}\ddot{\text{T}}\text{J}\text{C}-\text{PIp}_{\mu}\text{PSC}\ddot{\text{H}}\text{ C}\acute{\text{I}}\text{P}\ddot{\text{i}}\text{p}_{\mu}\text{C}\ddagger\text{P}\ddot{\text{e}}\text{C}_{\text{„C}}-\text{C}\ddagger\text{PSPsPiPs P}^{\circ}\text{PSC},\text{P}\ddot{\text{e}}\text{PiPi}_{\mu}\text{PSC}\acute{\text{r}}$   
 $\text{PiPi}_{\mu}\text{C}\ddot{\text{T}}\text{Jp}_{\mu}\text{PrPjC}-\text{C}\dots\text{Cr}\text{C}\ddot{\text{T}}\text{JPsPiPsC}-\text{P}\cdot\text{P}^{\circ}\text{P}\gg\text{PsP}\cdot\text{P}\ddot{\text{e}}$  (PSA).

$\text{P}\ddot{\text{K}}\text{P}\mu\text{P}\text{S}\text{P}\pm\text{C}\dots\text{C}-\text{PrPSPs P}\cdot\text{PsCl}\dot{\text{I}}\text{PsP}\pm\text{P}\gg\text{P}\ddot{\text{e}}\text{PIPsC}\dot{\text{H}}\text{PsP}\pm\text{P}\mu\text{C}\dot{\text{H}}\text{P}\mu\text{P}\dot{\text{I}}\text{PSC}-\text{Cl}\dot{\text{I}}\text{C},\text{C}\dot{\text{H}}$   
 $\text{P}\dot{\text{i}}\text{C}\dot{\text{H}}\text{P}\ddot{\text{e}}\text{P}\cdot\text{PSP}^\circ\text{C}\ddagger\text{P}^\circ\text{C},\text{P}\ddot{\text{e}}\text{P}\dot{\text{i}}\text{C}\dot{\text{H}}\text{P}\mu\text{P}\dot{\text{i}}\text{P}^\circ\text{C}\dot{\text{H}}\text{P}^\circ\text{C},\text{P}\dot{\text{i}}\text{P}^\circ\text{C}\ddagger\text{C}-\text{C}''\text{PSC},\text{P}^\circ\text{Pj P}\cdot\text{C},\text{C}\dot{\text{I}}\dot{\text{P}}\dot{\text{I}}\text{P}\text{ePsC}\dot{\text{H}}$   
 $\text{C}_{,,}\text{PsC}\dot{\text{H}}\text{PjPsC}\dot{\text{H}}\text{PSP}\ddot{\text{e}}\text{C}\dot{\text{H}}\text{P}\text{ePsPIP}\text{sC}-\text{PSP}\mu\text{PrPsCl}\dot{\text{I}}\text{C},\text{P}^\circ\text{C},\text{PSPsCl}\dot{\text{I}}\text{C},\text{C}-\text{(PeP}\gg\text{C}-\text{C}\dot{\text{H}}\text{P}\mu\text{PSC}\dot{\text{I}}$   
 $\text{PeC}\dot{\text{H}}\text{P}\mu\text{P}^\circ\text{C},\text{P}\ddot{\text{e}}\text{PSC}-\text{PSC}\dot{\text{I}} < 10\text{ PjP}\gg/\text{C}\dots\text{PI})\text{C}\ddagger\text{P}\mu\text{C}\dot{\text{H}}\text{P}\mu\text{P}\cdot\text{PIC}-\text{PrCl}\dot{\text{I}}\text{C}\dot{\text{r}}\text{C},\text{PSC}-\text{Cl}\dot{\text{I}}\text{C},\text{C}\dot{\text{H}}$   
 $\text{PrPsCl}\dot{\text{I}}\text{C},\text{P}^\circ\text{C},\text{PSC}\dot{\text{H}}\text{PsPiPs}\text{PeP}\gg\text{C}-\text{PSC}-\text{C}\ddagger\text{PSPsPiPs}\text{PrPsCl}\dot{\text{I}}\text{PIC}-\text{PrCr}\dot{\text{I}}.$

$P-CH_3C-PrPeP^{\circ}$   $PiPsPIC-PrPsPjP \gg ClP \gg PsCl'Cl$   $PiCH_3Ps$   $CH_3PsP \cdot PIPeC, PsPe$   
 $P^{\circ}PSPiC-PsPSP_{\mu}PICCH_3PsC, PeC \ddagger PSPsPiPs$   $PSP^{\circ}P \pm CH_3ClPeCr$   $PiC-Cl'P \gg Cl$   
 $P \cdot P^{\circ}Cl'C, PsCl'CrPIP^{\circ}PSPSCl$   $C, P^{\circ}PjCl'CrP \gg PsP \cdot PePSCr$ .  $Pj$   $C, P^{\circ}PePsPjCr$   $PIPePiP^{\circ}PrPeCr$   
 $P \gg C-PeCrPIP^{\circ}PSPSCl$   $Cl'P \gg C-Pr$   $PSP_{\mu}PiP^{\circ}PN_{\circ}PSPs$   $PiCH_3PePiPePSPeC, Pe$   $C, P^{\circ}$   
 $PePsPSC, CH_3PsP \gg CH_3PIP^{\circ}C, Pe$   $Cl'C, P^{\circ}PS$   $PiP^{\circ}C \ddagger C-C''PSC, P^{\circ}$   $PrPs$   $P \cdot PSPePePSP_{\mu}PSPSCl$   
 $PSP^{\circ}P \pm CH_3ClPeCr$ ,  $C, P^{\circ}PjCl'CrP \gg PsP \cdot PePS$   $PSP_{\mu}$   $Cl'P \gg C-Pr$   $PiCH_3PePN_{\circ}PjP^{\circ}C, Pe$   
 $PiPsPIC, PsCH_3PSPs$ .

PJ PrPμCΠPePëC... PïP°C†C-C''PSC,C-PI, CΠPeC- PïCΠPëPNëPjP°P»Pë P°P±Ps  
PïCΠPëPNëPjP°CΠC,CH C,P°PjC†C†P»PsP·PëPS, PïC-Pr C†P°C† C...C-CΠC†CΠPïC-C†PSPsPiPs  
PIC,CΠC†C†P°PSPSCΠ P· PïCΠPëPIPsPrC† PIPëPrP°P»PμPSPSCΠ PeP°C,P°CΠP°PeC,Pë C-  
PiP»P°C†PePsPjPë PIC-PrP·PSP°C†P°P»P°C†CΠ PSPμC†C,P°P±C-P»CHΨSC-C†C,CH  
CΠP°PNëPrC†P¶PSPsC— PsP±PsP»PsPSPePë PsPeP° (C†PëPSPrCΠPsPj P°C,PsPSC-C†PSPsC—  
P·C-PSPëC†C- (IFIS), PIP°CΠC-P°PSC, C†PëPSPrCΠPsPjC† P·PIC†P¶PμPSPsC— P·C-PSPëC†C-),  
PïPsPIBη™ CΠP·P°PSP° P· P±P»PsPeC†PIP°PSPSCΠPj B B B B B B B α<sub>1</sub>-CΠPμC†PμPïC,PsCΠC-PI.  
IFIS P·P±C-P»CHC€C†C'' CΠPëP·PëPe CΠPsP·PIPëC,PeC† PsC†PSPëC... C†C†PeP»P°PrPSPμPSCB  
PïC-Pr C†P°C† P°P±Ps PïC-C†P»CΠ C,P°PePsC— PsPïPμCΠP°C†C-C—.

PīPe PīCḥP°PIPēP»Ps, P·P° 1-2 C,PēP¶PSC- PīPμCḥPμPr PīCḥPsPIPμPrPμPSPSC¶Pj  
 PsPīPμCḥP°C†C-C— P· PīCḥPēPIPμPrCr PēP°C,P°CḥP°PeC,Pē C- PīP»P°C†PePsPjPē  
 CḥPμPePsPjPμPSPPrC†C”C,CHC†C¶ PīCḥPēPīPēPSPēC,Pē P»C-PeC†PIP°PSPSC¶  
 C,P°PjC†C†P»PsP·PēPSPsPj, PīCḥPsC,Pμ PePsCḥPēC†C,CH, PrPsC†C-P»CHPSC-C†C,CH C,P°  
 C,PμCḥPjC-PSPē PīCḥPēPīPēPSPμPSPSC¶ P»C-PeC†PIP°PSPSC¶ C,P°PjC†C†P»PsP·PēPSPsPj  
 PSP° C†CHPsPīPsPrPSC- C,PsC‡PSPs PSPμ PIC†C,P°PSPsPIP»PμPSC-. PμCḥPs IFIS  
 PīPsPIC-PrPsPjP»C¶P»PsC†C¶ C,P°PePsP¶ C- C† PīP°C†C-C”PSC,C-PI, C¶PeC-  
 PīCḥPēPīPēPSPēP»Pē P·P°C†C,PsC†C†PIP°PSPSC¶ C,P°PjC†C†P»PsP·PēPSC†  
 PIPīCḥPsPrPsPIP¶ P±C-P»CHC€ C,CḥPēPIP°P»PsPīPs PīPμCḥC-PsPrC† PrPs  
 PīPsC‡P°C,PeC† PsPīPμCḥP°C†C-C—.

P PsP·PiPsC‡PëPSP°C,Pë C,PµCḤP°PīC-CḤ C,P°PjCǻCǻP»PsP·PëPSPsPj  
PiC-PrCḤPsC...P»PsCḤPëPrPsPj PSPµ CḤPµPePsPjPµPSPPrCǻC”C,CHCǻCǻ PīP°C†C-C”PSC,P°Pj,  
CǻPePëPj P·P°PīP»P°PSPsPIP°PSP° PsPiPµCḤP°C†C-Cǻ P· PīCḤPëPIPsPrCǻ PeP°C,P°CḤP°PeC,Pë  
P°P±Ps PiP»P°CǻPePsPjPë. PµCḤPë PīC-PrPiPsC,PsPIC†C- PrPs PsPiPµCḤP°C†C-C—  
C...C-CḤCǻCḤPiPë-PsC,,C,P°P»CHPjPsP»PsPiPë PjP°CḤC,CH PrC-P·PSP°C,PëCǻCǻ, C‡Pë  
PiCḤPëPµPjP°PI (P°P±Ps PiCḤPëPµPjP°C”) PiP°C†C-C”PSC, C,P°PjCǻCǻP»PsP·PëPS, P·  
PjPµC,PsCḤ PiPsPiPµCḤPµPrP¶PµPSPSCǻ PjPsP¶P»PëPIPëC... CǻCǻPeP»P°PrPSPµPSCḤ,  
PiPsPiḤḤ™CǻP·P°PSPëC... P· IFIS.

PýP°PjCÍĆrP»PsP·PëPS C-PSC,PµPSCÍPëPIPSs PjPµC,P°P±PsP»C-P·CíC”C,CĤCÍCĬ,  
PiPsP»PsPIPSPëPj C†PëPSPsPj CYP3A4 C- CYP2D6. PµP°C†C-C”PSC,P°Pj P· PSPëP·CĤPePëPj  
PjPµC,P°P±PsP»C-P·PjPsPj CYP2D6 PSPµ CÍP»C-Pr PiCĤPëP·PSP°C†P°C,Pë

C,P°PjCÍCfP»PsP·PëPSCf PiC-PrCḡPsC...P»PsCḡPëPr CÍ PePsPjP±C-PSP°C†C-C— P·  
CÍPëP»CHbPSPëPjPë C-PSPiC-P±C-C,PScḡP°PjPë CYP3A4 (PSP°PíCḡPëPeP»P°Pr  
PePµC,PSPePsPSP°P·PsP»PsPj).

PŷP°PjCÍCfP»PsP·PëPSCf PiC-PrCḡPsC...P»PsCḡPëPr CÍP»C-Pr P·P°CÍC,PScÍPsPICfPIP°C,Pë P·  
PsP±PµCḡPµP¶PSC-CÍC,Cḡ CÍ PíPsC”PrPSP°PSPSC- P· CÍPëP»CHbPSPëPjPë  
(PSP°PíCḡPëPeP»P°Pr PePµC,PSPePsPSP°P·PsP»PsPj) C- PíPsPjC-CḡPSPëPjPë  
C-PSPiC-P±C-C,PScḡP°PjPë CYP3A4 (PSP°PíCḡPëPeP»P°Pr PµCḡPëC,CḡPsPjC-C†PëPSPsPj)  
(PrPëPI. CḡPsP·PrC-P» B«P’P·P°C”PjPsPrC-C‡ P· C-PSC€PëPjPë P»C-PeP°CḡCÍCHbPePëPjPë  
P·P°CÍPsP±P°PjPë C,P° C-PSC€C- PIPëPrPë PIP·P°C”PjPsPrC-PNëB»).

PµPsPIC-PrPsPjP»C‡P»PsCÍCH PíCḡPs PIPëPíP°PrPePë P°P»PµCḡPiC-C†PSPëC...  
CḡPµP°PeC†C-PNë PSP° C,P°PjCÍCfP»PsP·PëPS CÍ PíP°C†C-C”PSC,C-PI P· PSP°C‡PIPSC-CÍC,Cḡ  
PI P°PSP°PjPSPµP·C- P°P»PµCḡPiC-C— PSP° CÍCfP»CHbC,,P°PSC-P»P°PjC-PrPë. PŷP»C-Pr  
PrPsC,CḡPëPjCfPIP°C,PëCÍCH PsP±PµCḡPµP¶PSPScÍC,C- PíCḡPë P·P°CÍC,PScÍCfPIP°PSPSC-  
C,P°PjCÍCfP»PsP·PëPSCf PiC-PrCḡPsC...P»PsCḡPëPrCÍ PíP°C†C-C”PSC,P°Pj, CÍ C‡PePëC...  
CḡP°PSC-C€Pµ PIC-PrP·PSP°C†P°P»P°CÍCH P°P»PµCḡPiC-C‡ PSP°  
CÍCfP»CHbC,,P°PSC-P»P°PjC-PrPë.

PµCḡPµPíP°CḡP°C, CÍP»C-Pr PíCḡPëPMëPjP°C,Pë P·P° PíCḡPëP·PSP°C†PµPSPSC‡Pj  
P»C-PeP°CḡC‡.

*P—P°CÍC,PScÍCžPIP°PSPSC‡ Cž PíPµCḡC-PsPr PIP°PiC-C,PSPScÍC,C- P°P±Ps  
PíPsPrCžPIP°PSPSC‡ PiCḡCžPrPrCḡ.*

PŷP°PjCÍCfP»C-Pr PSPµ PíPsPeP°P·P°PSPëPNë PrP»C‡ P·P°CÍC,PScÍCfPIP°PSPSC‡  
P¶C-PSPeP°Pj.

PµPµCḡC,PëP»CHbPSC-CÍC,Cḡ. PµC-Pr C†P°CÍ PeP»C-PSC-C†PSPëC... PrPsCÍP»C-PrP¶PµPSCH  
C,P°PjCÍCfP»PsP·PëPSCf PíCḡPsC,C‡PiPsPj PePsCḡPsC,PePsPiPs C,P° C,CḡPëPIP°P»PsPiPs  
C†P°CÍCf PIC-PrPjC-C†P°P»PëCÍC‡ PíPsCḡCfC€PµPSPSC‡ PµC‡PeCfP»C‡C†C-C—.  
P’PëPíP°PrPePë PíPsCḡCfC€PµPSPSC‡ PµC‡PeCfP»C‡C†C-C—, CḡPµC,CḡPsPiCḡP°PrPSPsC—  
PµC‡PeCfP»C‡C†C-C— C- PSPµPrPsCÍC,P°C,PSCHbPsC— PµC‡PeCfP»C‡C†C-C—  
PIC-PrP·PSP°C†P°P»PëCÍC‡ PI PíC-CÍP»C‡CḡPµC”CÍC,CḡP°C†C-PNëPSPëPNë PíPµCḡC-PsPr  
P·P°CÍC,PScÍCfPIP°PSPSC‡ C,P°PjCÍCfP»PsP·PëPSCf.

*P—PrP°C,PSC-CÍC,Cḡ PIPíP»PëPIP°C,Pë PSP° C€PIPëPrPeC-CÍC,Cḡ CḡPµP°PeC†C-C— PíCḡPë  
PePµCḡCžPIP°PSPSC- P°PIC,PSc,CḡP°PSCÍPíPsCḡC,PSPj P°P±Ps C-PSC€PëPjPë  
PjPµC...P°PSC-P·PjP°PjPë.*

P”PsCÍP»C-PrP¶PµPSPSC‡ PIPíP»PëPICf PíCḡPµPíP°CḡP°C,CÍ PSP° P·PrP°C,PSC-CÍC,Cḡ  
PePµCḡCfPIP°C,Pë P°PIC,PSc,CḡP°PSCÍPíPsCḡC,PSPj P°P±Ps C-PSC€PëPjPë  
PjPµC...P°PSC-P·PjP°PjPë PSPµ PíCḡPsPIPSPrPëP»PëCÍC‡. PhPrPSP°Pe PíP°C†C-C”PSC,C-PI  
PSPµPsP±C...C-PrPSPs PíPsPíPµCḡPµPrPëC,Pë PíCḡPs PjPsP¶P»PëPIC-CÍC,Cḡ  
PIPëPSPëPePSPµPSPSC‡ CÍPsPSP»PëPIPsCÍC,C-, PíPsPjCfC,PSC-PSPSC‡ P·PsCḡCf,  
P·P°PíP°PjPsCḡPsC†PµPSPSC‡ C,P° PSPµPíCḡPëC,PSPjPSPScÍC,C-.

***PŸPİPsCÍC-P± P·P°CÍC,PsCÍCžPIP°PSPSCİİ C,P° PrPsP·Pë.***

P PµPePsPjPµPSPPrPsPIP°PSP° PrPsP·P° PrP»Cİİ PrPsCĤPsCÍP»PëC... бĤ“ 1  
PeP°PİCÍCÍP»P° C%PsPrPµPSPSPs, PİC-CÍP»Cİİ CÍPSC-PrP°PSPeCÍ P°P±Ps PİC-CÍP»Cİİ  
PİPµCĤC€PsPiPs PİCĤPëP№PsPjCÍ C—P¶C-. PᄂP°PİCÍCÍP»CÍ CÍP»C-Pr PePsPIC,P°C,Pë  
C†C-P»PsCĤ, CÍPëPrCİC†Pë P°P±Ps CÍC,PsCİC†Pë. PᄂP°PİCÍCÍP»CÍ PSPµ PjPsP¶PSP°  
CĤPsP·P¶PsPICÍPIP°C,Pë, CĤPsP·P»P°PjCÍPIP°C,Pë, PsCÍPeC-P»CĤPePë C†Pµ P±CÍPrPµ  
PİPµCĤPµC€PePsPrP¶P°C,Pë PjPsPrPëC,,C-PePsPIP°PSPsPjCÍ PIPëPIC-P»CĤPSPµPSPSCĤ  
P°PeC,PëIPSPsPiPs C-PSPICĤPµPrC-C”PSC,P°.

*PµP°C†C-C”PSC,Pë P»C-C,PSCHbPsPiPs PIC-PeCž PSPµ PİPsC,CĤPµP±CÍCĤC,CHb  
PePsCĤPµPeC†C-C— PrPsP·Pë. PhPrPSP°Pe CÍP»C-Pr PjP°C,Pë PSP° CÍPIP°P·C-, C%Ps CÍ  
PiPµCĤC-P°C,CĤPëC†PSPëC... PİP°C†C-C”PSC,C-PI CĤPëP·PëPe PsCĤC,PsCÍC,P°C,PëC†PSPsC—  
PiC-PİPsC,PµPSP·C-C— P·P±C-P»CHbC€PµPSPëP№.*

*PµP°C†C-C”PSC,P°Pj P· PSPëCĤPePsPIPsCĤ PSPµPrPsCÍC,P°C,PSC-CÍC,CĤ PSPµ  
PİPsC,CĤC-P±PSP° PePsCĤPµPeC†C-Cİİ PrPsP·Pë.*

*PµP°C†C-C”PSC,P°Pj P· PİPµC†C-PSPePsPIPsCĤ PSPµPrPsCÍC,P°C,PSC-CÍC,CĤ  
PİPsPjC-CĤPSPsPiPs C,P° CÍPµCĤPµPrPSCĤPsPiPs CÍC,CÍPİPµPSCİİ C,CİP¶PePsCÍC,C- PSPµ  
PİPsC,CĤC-P±PSP° PePsCĤPµPeC†C-Cİİ PrPsP·Pë (PrPëPI. C,P°PePsP¶  
B«PµCĤPsC,PëPİPsPeP°P·P°PSPSCİİB»).*

*P”C-C,Pë.*

PµCĤPµPİP°CĤP°C, PSPµ P·P°CÍC,PsCÍPsPICÍPIP°C,Pë PrC-C,CİPj. P’PµP·PİPµPeP° C,P°  
PµC,,PµPeC,PëIPSC-CÍC,CHb P·P°CÍC,PsCÍCÍPIP°PSPSCİİ C,P°PjCÍCÍP»PsP·PëPSCÍ PrC-C,CİPj  
(PrPs 18 CĤPsPeC-PI) PSPµ PsC†C-PSCĤPIP°P»P°CÍCĤ.

***PµPµCĤPµPrPsP·CžPIP°PSPSCİİ.***

*PŸPëPjPİC,PsPjPë. PµPµCĤPµPrPsP·CÍPIP°PSPSCİİ C,P°PjCÍCÍP»PsP·PëPSCÍ  
PiC-PrCĤPsC...P»PsCĤPëPrCÍ PjPsP¶Pµ CÍPİCĤPëC†PëPSPëC,Pë PiPsCÍC,CĤC- C,P° C,CİP¶PeC-  
PiC-PİPsC,PµPSP·PëIPSC- PµC,,PµPeC,Pë, P° C,P°PePsP¶ P±P»CĤPIP°PSPSCİİ C,P°  
PİCĤPsPSPsCÍ. PŸCİP¶PeC- PiC-PİPsC,PµPSP·PëIPSC- PµC,,PµPeC,Pë  
CÍPiPsCÍC,PµCĤC-PİP°P»PëCÍCİİ PİCĤPë CĤC-P·PSPëC... CÍC,CÍPİPµPSCİİC...  
PİPµCĤPµPrPsP·CÍPIP°PSPSCİİ.*

*P»C-PeCžPIP°PSPSCİİ. Pj CĤP°P·C- PIPëPSPëPePSPµPSPSCİİ PiPsCÍC,CĤPsC—  
P°CĤC,PµCĤC-P°P»CHbPSPsC— PiC-PİPsC,PµPSP·C-C— PIPSP°CÍP»C-PrPsPe  
PİPµCĤPµPrPsP·CÍPIP°PSPSCİİ CÍP»C-Pr PİCĤPsPIPsPrPëC,Pë PİC-PrC,CĤPëPjCÍPIP°P»CHbPSCÍ  
C,PµCĤP°PİC-CĤ, CÍPİCĤCİPjPsPIP°PSCÍ PSP° PIC-PrPSPsPIP»PµPSPSCİİ  
PSPsCĤPjP°P»CHbPSPsC— C,,CÍPSPeC†C-C— CÍPµCĤC†PµPIPs-CÍCÍPrPëPSPSPsC—  
CÍPëCÍC,PµPjPë (PSP°PİCĤPëPeP»P°Pr, PİP°C†C-C”PSC, PjP°C” PİCĤPëP№PSCİİC,Pë  
PiPsCĤPëP·PsPSC,P°P»CHbPSPµ PİPsP»PsP¶PµPSPSCİİ). PİPeC%Ps C†Pµ PSPµ  
PrPsPİPsPjP°PİP°C”, CÍP»C-Pr PİCĤPsPIPµCÍC,Pë C-PSC,,CÍP·C-P№PSCÍ C,PµCĤP°PİC-CĤ C,P°  
PİCĤPëP·PSP°C†PëC,Pë PIP°P·PsPİCĤPµCÍPsCĤPSC- P·P°CÍPsP±Pë. PŸP»C-Pr  
PePsPSC,CĤPsP»CĤPIP°C,Pë C,,CÍPSPeC†C-CĤ PSPëCĤPsPe C,P° PİCĤPsPIPsPrPëC,Pë  
P·P°PİP°P»CHbPSC- PiC-PrC,CĤPëPjCÍPIP°P»CHbPSC- P·P°C...PsPrPë. PŞPµCĤPµP·*

PIPëCÍPsPePëPNë CÍC,CÍPíC-PSCĤ P·PíBĤ™ CÍP·CÍPIP°PSPSCÍ C,P°PjCÍCÍP»PsP·PëPSCÍ P·P±C-P»PeP°PjPë PíP»P°P·PjPë PeCĤPsPIC- PíCĤPsPIPµPrPµPSPSCÍ PíPµPjPsPrC-P°P»C-P·CÍPSP°PICĤCÍPr C±Pë PrPsC†C-P»CĤPSPµ.

P— PjPµC,PsCĤ PíCĤPëPíPëPSPµPSPSCÍ PíPsPrP°P»CĤC€PsPiPs PICÍPjPsPeC,CÍPIP°PSPSCÍ PíCĤPµPíP°CĤP°C,CÍ PjPsP¶PSP° C€C,CÍC±PSPs PIPëPeP»PëPeP°C,Pë P±P»CĤPIP°PSPSCÍ. PµCĤPë PíPµCĤPµPrPsP·CÍPIP°PSPSC- P·PSP°C±PSPsCĤ PeC-P»CĤPeC-CÍC,CĤ PíCĤPµPíP°CĤP°C,CÍ PíP°C†C-C”PSC,CÍ CÍP»C-Pr PíCĤPsPjPëC,Pë C€P»CÍPSPsPe, PjPsP¶PSP° P·P°CÍC,PsCÍCÍPIP°C,Pë P°PeC,PëPIPsPIP°PSPµ PICÍPíC-P»P»CÍ C,P° PSPëP·CĤPePsPsCÍPjPsC,PëC±PSC- PíCĤPsPSPsCÍPSC- P·P°CÍPsP±Pë, C,P°PeC- CÍPe CÍCÍP»CĤC,,P°C, PSP°C,CĤC-CĤ.

**PµPsP±C-C±PSC- CĤPµP°PeC†C-C—.**

PµPsP±C-C±PSC- CĤPµP°PeC†C-C— PeP»P°CÍPëC,,C-PePsPIP°PSC- P·P° PsCĤPiP°PSP°PjPë C-CÍPëCÍC,PµPjP°PjPë C,P° C±P°CÍC,PsC,PsCĤ C—C... PIPëPSPëPePSPµPSPSCÍ: C±P°CÍC,Ps (> 1/100, < 1/10); PSPµC±P°CÍC,Ps (> 1/1000, < 1/100); CĤC-PrPePs (> 1/10000, < 1/1000); PrCÍP¶Pµ CĤC-PrPePs (< 1/10000), PSPµPIC-PrPsPjPs (PSPµPjPsP¶P»PëPIPs PsC†C-PSPëC,Pë P·P° PrPsCÍC,CÍPíPSPëPjPë PrP°PSPëPjPë).

*PkPµCĤPIPsPIP° CÍPëCÍC,PµPjP°:* C±P°CÍC,Ps BĤ“ P·P°PíP°PjPsCĤPsC±PµPSPSCÍ, PSPµC±P°CÍC,Ps BĤ“ PíPsP»PsPIPsPëPNë P±C-P»CĤ, CĤC-PrPePs BĤ“ PSPµPíCĤPëC,PsPjPSC-CÍC,CĤ (CÍPëPSPePsPíPµ).

*PĥCĤPiP°PSPë P·PsCĤCž:* PSPµPIC-PrPsPjPs BĤ“ P·P°C,CÍPjP°PSPµPSPSCÍ P·PsCĤCÍ\*, PíPsCĤCÍC€PµPSPSCÍ P·PsCĤCÍ\*.

*PŸPµCĤC†PµPIPs-CÍCžPrPëPSPSP° CÍPëCÍC,PµPjP°:* PSPµC±P°CÍC,Ps BĤ“ PsCĤC,PsCÍC,P°C,PëC±PSP° PíC-PíPsC,PµPSP·C-CÍ, PIC-PrC±CÍC,C,CÍ CÍPµCĤC†PµP±PëC,C,CÍ (PíP°P»CĤPíC-C,P°C†C-CÍ).

*P”PëC...P°P»CĤPSP° CÍPëCÍC,PµPjP°:* PSPµC±P°CÍC,Ps BĤ“ CĤPëPSC-C,, PSPµPIC-PrPsPjPs BĤ“ PSPsCÍPsPIP° PeCĤPsPIPsC,PµC±P°\*.

*P PµPíCĤPsPrCžPeC,PëPIPsPSP° CÍPëCÍC,PµPjP°:* C±P°CÍC,Ps BĤ“ CĤPsP·P»P°PrPë PµCÍPeCÍP»CÍC†C-C—, PIPeP»CĤC±P°CĤC±Pë CĤPµC,CĤPsPiCĤP°PrPSCÍ PµCÍPeCÍP»CÍC†C-CĤ C- PSPµPrPsCÍC,P°C,PSC-CÍC,CĤ PµCÍPeCÍP»CÍC†C-C—, PrCÍP¶Pµ CĤC-PrPePs BĤ“ PíCĤC-P°PíC-P·Pj.

*PŸCĤP°PIPsPëPNë C,CĤP°PeC,:* PSPµC±P°CÍC,Ps BĤ“ P·P°PíPsCĤ, PrC-P°CĤPµCÍ, PSCÍPrPsC,P°, P±P»CĤPIP°PSPSCÍ, PSPµPIC-PrPsPjPs BĤ“ CÍCÍC...C-CÍC,CĤ CÍ CĤPsC,C-.

*PëPeC-CĤP°, PíC-PrC€PeC-CĤPSP° PeP»C-C,PePsPIPëPSP°:* PSPµC±P°CÍC,Ps BĤ“ CÍPIPµCĤP±C-P¶, PIPëCÍPëPíP°PSPSCÍ, PeCĤPsPíPëPíBĤ™ CÍPSPeP°, CĤC-PrPePs BĤ“ P°PSPíC-PsPSPµPICĤPsC,PëC±PSPëPNë PSP°P±CĤCÍPe, PrCÍP¶Pµ CĤC-PrPePs BĤ“ CÍPëPSPrCĤPsPj PŸC,C-PIPµPSCÍP°-P”P¶PsPSCÍPsPSP°, PSPµPIC-PrPsPjPs BĤ“ PjCÍP»CĤC,PëC,,PsCĤPjPSP° PµCĤPëC,PµPjP°\*, PµPeCÍC,,PsP»C-P°C,PëPIPsPëPNë PrPµCĤPjP°C,PëC,\*, CĤPµP°PeC†C-C— C,,PsC,PsC±CÍC,P»PëPIPsCÍC,C-.\*.

*P—P°PíP°P»CĤPSC- CĤPsP·P»P°PrPë:* PSPµC±P°CÍC,Ps BĤ“ P°CÍC,PµPSC-CÍ.

\*P'C-PrPjC-C†P°P»PëCíCЦ PI PíC-CíP»CЦCђPµC"CíC,CђP°C†C-PNëPSPëPNë PíPµCђC-PsPr P·P°CíC,PscíCfPIP°PSPSCЦ C,P°PjCíCfP»PsP·PëPSCf.

PлCђC-Pj PIPëC%PµP·P°P·PSP°C†PµPSPëC... PíPsP±C-C†PSPëC... CђPµP°PeC†C-PNë, Cí PíC-CíP»CЦCђPµC"CíC,CђP°C†C-PNëPSPëPNë PíPµCђC-PsPr PSP°PrC...PsPrPëP»Pë CíPíPsPSC,P°PSPSC- PíPsPIC-PrPsPjP»PµPSPSCЦ PíCђPs PIPëPíP°PrPePë P°C,CђC-P°P»CHPSPSc— C,,C-P±CђPëP»CЦC†C-C—, C,P°C...C-PeP°CђPrC-C—, P°CђPëC,PjC-C—, PrPëCíPíPSPsPµ. PhCíPeC-P»CHPePë PíCђPs P·P°P·PSP°C†PµPSC- PIPëPíP°PrPePë PíPsPIC-PrPsPjP»CЦP»PsCíCЦ CíPíPsPSC,P°PSPSPs, C†P°CíC,Psc,P° CђPµP°PeC†C-PNë C,P° C—C... P·PIbђ™ CЦP·PsPe P· C,P°PjCíCfP»PsP·PëPSPsPj PSPµ PjPsP¶CíC,CH P±CíC,Pë PrPsCíC,PsPIC-CђPSPs PIPëP·PSP°C†PµPSC-.

PíPe C- PíCђPë PíCђPëPNëPsPjC- C-PSCëPëC... P°P»CHC,,P°-P±P»PsPeP°C,PscђC-PI, PjPsP¶Pµ CíPíPsCíC,PµCђC-PiP°C,PëCíCЦ CíPsPSP»PëPIC-CíC,CH, CíCíC...C-CíC,CH Cí CђPsC,C-, PSP°P±CђCЦPePë.

PhPíPëCíP°PSC- PIPëPíP°PrPePë C-PSC,CђP°PsPíPµCђP°C†C-PNëPSPSc— PSPµCíC,P°P±C-P»CHPSPsCíC,C- CђP°PNëPrCfP¶PSPsC— PsP±PsP»PsSPPePë PsPeP° (PIP°CђC-P°PSC, CíPëPSPrCђPsPjCf PICfP·CHPePsC— P·C-PSPëC†C-, PIC-PrPsPjPëPNë CЦPe IFIS) PíC-Pr C†P°Cí PsPíPµCђP°C†C-C— P· PíCђPëPIPsPrCf PeP°C,P°CђP°PeC,Pë C,P° PíP»P°CfPePsPjPë Cí PíP°C†C-C"PSC,C-PI, CЦPeC- C,CђPëPIP°P»PëPNë C†P°Cí PíCђPëPNëPjP°P»Pë C,P°PjCíCfP»PsP·PëPS.

***PýPµCђPjC-PS PíCђPëPrP°C,PSPsCíC,C-.*** 3 CђPsPePë.

PкPµ P·P°CíC,PscíCfPsPICfPIP°C,Pë PíC-CíP»CЦ P·P°PeC-PSC†PµPSPSCЦ C,PµCђPjC-PSCf PíCђPëPrP°C,PSPsCíC,C-, PIPEP°P·P°PSPsPíPs PSP° CíPíP°PePsPIC†C-.

***PJPjPsPIPë P·P±PµCђC-PiP°PSPSCЦ.***

P—P±PµCђC-PiP°C,Pë PI PsCђPëPíC-PSP°P»CHPSC-PNë CíPíP°PePsPIC†C- PíCђPë C,PµPjPíPµCђP°C,CíCђC- PSPµ PIPëC%Pµ 25 B°Pý.

P—P±PµCђC-PiP°C,Pë Cí PSPµPrPsCíC,CíPíPSPsPjCf PrP»CЦ PrC-C,PµPNë PjC-CíC†C-.

***PJPíP°PePsPIPEP°.***

PµPs 10 PeP°PíCíCfP» PI P±P»C-CíC,PµCђC-, PíPs 3 P±P»C-CíC,PµCђPë PI PíP°C†C†C-.

***PлP°C,PµPíPsCђC-CЦ PIC-PrPíCfCíPeCf.*** P—P° CђPµC†PµPíC,PsPj.

***P'PëCђPsP±PSPëPe.***

$P_{\mu}C^{\dagger}P \pm P \rangle C - C \dagger PSP_{\mu} \quad P^{\circ}PeC^{\dagger}C - PsPSP_{\mu}C_{\eta}PSP_{\mu} \quad C, PsPIP^{\circ}C_{\eta}P\ddot{e}C^{\dagger}C, PIPs \quad B \langle P^{\circ}C^{\dagger}PePsPIPs -$   
 $PIP\ddot{e}C_{\eta}PsP \pm PSP\ddot{e}C \dagger P\ddot{e}PN_{\mathbb{N}} \quad C \dagger P_{\mu}PSC, C_{\eta} \quad B \langle P^{\circ}PsC_{\eta}C \% P^{\circ}PiC - PIC^{\dagger}CHPeP\ddot{e}PN_{\mathbb{N}} \quad C \dots C - PjC - PePs -$   
 $C, P^{\circ}C_{\eta}PjP^{\circ}C \dagger P_{\mu}PIC, P\ddot{e}C \dagger PSP\ddot{e}PN_{\mathbb{N}} \quad P \cdot P^{\circ}PIPsPrB \rangle.$

**$P_{\mathbb{H}}C - C^{\dagger}C \dagger P_{\mu}P \cdot PSP^{\circ}C \dots PsPrP \P P_{\mu}PSPSC_{\eta} \quad PIP\ddot{e}C_{\eta}PsP \pm PSP\ddot{e}PeP^{\circ} \quad C, P^{\circ} \quad PN_{\mathbb{N}}PsPiPs$**   
 **$P^{\circ}PrC_{\eta}P_{\mu}C^{\dagger}P^{\circ} \quad PjC - C^{\dagger}C \dagger C_{\eta} \quad PiC_{\eta}PsPIP^{\circ}PrP \P P_{\mu}PSPSC_{\eta} \quad PrC - C_{\eta}P \rangle CHPsPsC^{\dagger}C, C -.$**

$PjPeC_{\eta}P^{\circ}C - PSP^{\circ}, \quad 03134, \quad Pj. \quad P_{\mathbb{H}}P\ddot{e}C - PI, \quad PIC^{\dagger}P \rangle. \quad P_{\mathbb{H}}P\ddot{e}C_{\eta}C^{\dagger}, \quad 17.$