



PhPSC, PëPiC-PiPµCḡC, PµPSP·PëPIPSC- P·P°CÍPsP±Pë. PhPSC, PëP°PrCḡPµPSPµCḡPiC-C±PSC- P·P°CÍPsP±Pë P· PiPµCḡPëC,, PµCḡPëC±PSPëPj PjPµC...P°PSC-P·PjPsPj PrC-C—. P·P»PsPeP°C, PsCḡPë α-P°PrCḡPµPSPsCḡPµC†PµPiC, PsCḡC-PI. P·P·P·P· PhPÿX Pÿ02Pÿ Ph04.

***PαP°CḡPjP°PePsP»PsPiC-C±PSC- PIP»P°CÍC, PëPIPsCÍC, C-.***

*PαP°CḡPjP°PePsP·PëPSP°PjC-PeP°.*

P·PµC...P°PSC-P·Pj PrC-C—.

P°PsPeCÍP°P·PsP·PëPS C” PiPsC, CÍP¶PSPëPj C, P° CÍPµP»PµPeC, PëPIPSPëPj P°PSC, P°PiPsPSC-CÍC, PsPj PiPsCÍC, CÍPëPSP°PiC, PëC±PSPëC... α₁- P°PrCḡPµPSPsCḡPµC†PµPiC, PsCḡC-PI. P·P»PsPeCÍP°PSPSCÍ C†PëC... CḡPµC†PµPiC, PsCḡC-PI PiCḡPëP·PIPsPrPëC, Ch PrPs P·PSPëP¶PµPSPSCÍ CÍPëCÍC, PµPjPSPsPiPs P°CḡC, PµCḡC-P°P»ChPSPsPiPs C, PëCÍP°CÍ. PµCḡPµPiP°CḡP°C, P·P°CḡPrCÍCḡP°<sup>B®</sup> PiCḡPëP·PSP°C†PµPSPëPµ PrP»CÍ PiPµCḡPsCḡP°P»ChPSPsPiPs P·P°CÍC, PsCÍCÍP°PSPSCÍ 1 CḡP°P·PSP° PrPsP±CÍ PiP°C†C-C”PSC, P°Pj, C...PIPsCḡPëPj PSP° PµCÍCÍPµPSC†C-P°P»ChPSCÍ P°CḡC, PµCḡC-P°P»ChPSCÍ PiC-PiPµCḡC, PµPSP·C-Cḡ.

PαP°CḡPjP°PePsP·PëPSP°PjC-C±PSC- PµC,, PµPeC, Pë.

P·CÍP»Ps PiCḡPsPrPµPjPsPSCÍC, CḡPsPIP°PSPs, C%Ps PiCḡPµPiP°CḡP°C, P·P°CḡPrCÍCḡP°<sup>B®</sup> PSPµ CÍPiCḡPëC±PëPSCÍC” PSPµP±P°P¶P°PSPëC... PjPµC, P°P±PsP»C-C±PSPëC... PµC,, PµPeC, C-PI C- PµPsPiPs PjPsP¶PSP° P·P°CÍC, PsCÍPsPICÍP°C, Pë PiP°C†C-C”PSC, P°Pj C-P· C†CÍPeCḡPsPIPëPj PrC-P°P±PµC, PsPj, PiPsPrP°PiCḡPsCḡ P°P±Ps C-PSCÍCÍP»C-PSPsCḡPµP·PëCÍC, PµPSC, PSC-CÍC, Cḡ.

PµCḡPµPiP°CḡP°C, P·P°CḡPrCÍCḡP°<sup>B®</sup> C, P°PePsP¶ PjPsP¶PSP° PiCḡPëP·PSP°C†P°C, Pë PiP°C†C-C”PSC, P°Pj C-P· P±CḡPsPSC...C-P°P»ChPSPsCḡ P°CÍC, PjPsCḡ, PiC-PiPµCḡC, CḡPsC,, C-C”Cḡ P»C-PIPsPiPs CëP»CÍPSPsC±PeP° C, P° PiP°C†C-C”PSC, P°Pj P»C-C, PSCḡPsPiPs PIC-PeCÍ. P—P°CÍC, PsCÍCÍP°PSPSCÍ PiCḡPµPiP°CḡP°C, CÍ CÍPiCḡPëCÍC” P·PjPµPSCëPµPSPSCḡ PiC-PiPµCḡC, CḡPsC,, C-C— P»C-PIPsPiPs CëP»CÍPSPsC±PeP°, PiCḡPëPiPSC-C†CÍC” P°PiCḡPµPiP°C†C-Cḡ C, CḡPsPjP±PsC†PëC, C-PI C, P° PiC-PrCÍPëP»CḡC” P°PeC, PëPIPSC-CÍC, Ch C, PeP°PSPëPSPSPsPiPs P°PeC, PëPIP°C, PsCḡP° PiP»P°P·PjC-PSPsPiPµPSCÍ. P·CḡC-Pj C, PsPiPs, P·P°CÍC, PsCÍCÍP°PSPSCÍ PiCḡPµPiP°CḡP°C, CÍ P·P°CḡPrCÍCḡP°<sup>B®</sup> PiC-PrPIPëC%CÍC” C†CÍC, P»PëPIC-CÍC, Ch PrPs C-PSCÍCÍP»C-PSCÍ CÍ C, PëC... PiP°C†C-C”PSC, C-PI, CÍ CÍPePëC... C, P°PeP° C†CÍC, P»PëPIC-CÍC, Ch C” PiPsCḡCÍCëPµPSPsCḡ.

PÿP°PePsP¶ CÍ PrPsPIPisCÍC, CḡPsPePsPIPëC... PrPsCÍP»C-PrP¶PµPSPSCÍC... P±CÍP»Ps PiCḡPsPrPµPjPsPSCÍC, CḡPsPIP°PSPs, C%Ps PSP° PrPsPrP°C, PsPe PrPs P°PSC, PëPiC-PiPµCḡC, PµPSP·PëPIPSPsC— PrC-C— P·P°CÍC, PsCÍCÍP°PSPSCÍ PiCḡPµPiP°CḡP°C, CÍ P·P°CḡPrCÍCḡP°<sup>B®</sup> CÍPiCḡPëC±PëPSCÍC” PiPsPjC-CḡPSPµ P·PSPëP¶PµPSPSCÍ PePsPSC†PµPSC, CḡP°C†C-C— P·P°PiP°P»ChPSPsPiPs C...PsP»PµCÍC, PµCḡPëPSCÍ, P»C-PiPsPiCḡPsC, PµC—PrC-PI PSPëP·ChPePsC— C%C-P»ChPSPsCÍC, C- C, P° C, CḡPëPiP»C-C†PµCḡPëPrC-PI CÍ PiP»P°P·PjC- PeCḡPsPIC-, P° PsC, P¶Pµ Bḡ” PrP°PSPëPµ PiCḡPµPiP°CḡP°C, PjPsP¶Pµ P±CÍC, Pë PsCÍPsP±P»PëPIPs PePsCḡPëCÍPSPëPµ PiP°C†C-C”PSC, P°Pj, C...PIPsCḡPëPj PSP° P°CḡC, PµCḡC-P°P»ChPSCÍ PiC-PiPµCḡC, PµPSP·C-Cḡ C, P° PiC-PiPµCḡP»C-PiC-PrPµPjC-Cḡ.

P—P°CÍC,PsCÍCfPIP°PSPSCЦ PíCĤPμPíP°CĤP°C,Cf PъP°CĤPpCfCĤP°<sup>B®</sup> PíP°C†C-C”PSC,P°Pj, C...PIPsCĤPëPj PSP° CÍPëPjPíC,PsPjPSCf PrPsP±CĤPpsCЦPеC-CÍPSCf PíC-PíPμCĤPíP»P°P-C-CĤ PíPμCĤPμPrPjC-C...CfCĤPpsPIPsC— P·P°P»PsP·Pë (P”P“PμP—), PíCĤPëP·PIPsPrPëC,Ch PrPs P·PSP°C†PSPsPiPs PíPsPeCĤP°C%oPμPSPSCЦ CÍCĤPpsPrPëPSP°PjC-PePë C,P° P·PjPμPSCePμPSPSCЦ CÍPëPjPíC,PsPjC-PI. P’PIP°P¶P°C”C,ChCÍCЦ, C%oPs PμC,,PμPeC, PíCĤPμPíP°CĤP°C,Cf PíCĤPë P”P“PμP— PrPsCÍCЦPíP°C”C,ChCÍCЦ P·P° CĤP°C...CfPSPsPe CÍPμP»PμPeC,PëPIPsC— P±P»PsPeP°PrPë α<sub>1</sub>-P°PrCĤPμPSPsCĤPμC†PμPíC,PsCĤC-PI, CĤPpsP·C,P°C€PsPIP°PSPëC... Cf PjвĤ™ CЦP·PsPIC-PN<sub>0</sub> CÍC,CĤPpsPjC- C,P° PeP°PíCÍCfP»C- PíPμCĤPμPrPjC-C...CfCĤPpsPIPsC— P·P°P»PsP·Pë, P° C,P°PePsP¶ Cf C€PëPN<sub>0</sub>C†C- CÍPμC†PsPIPsPiPs PjC-C...CfCĤP°.

*PαP°CĤPjP°PePsPeC-PSPμC,PëPeP°.*

P’CÍPjPsPeC,CfPIP°PSPSCЦ. PμCĤPë PíPμCĤPpsCĤP°P»ChPSPsPjCf P·P°CÍC,PsCÍCfPIP°PSPSC- Cf P»CĤPrPëPSPë (C†PsP»PsPIC-PePë PjPsP»PsPrPsPiPs PIC-PeCf P°P±Ps P»CĤPrPë P»C-C,PSCĤPsPiPs PIC-PeCf P±CfPrCh-CЦPеPsC— CÍC,P°C,C-) PrPsPeCÍP°P·PsP·PëPS C€PIPëPrPePs PICÍPjPsPeC,CfC”C,ChCÍCЦ P· P±C-PsPrPsCÍC,CfPíPSC-CÍC,CĤ PSP° CĤC-PIPsC- PíCĤPëP±P»PëP·PSPs 2/3 PrPsP·Pë.

P’C-PsC,CĤP°PSCÍC,,PsCĤPjP°C†C-CЦ/PμP»C-PjC-PSP°C†C-CЦ. PμCĤPëP±P»PëP·PSPs 98 % PrPsPeCÍP°P·PsP·PëPSCf P·PIвĤ™ CЦP·CfC”C,ChCÍCЦ P· P±C-P»PeP°PjPë PíP»P°P·PjPë PeCĤPpsPIC-. P’CfP»Ps PíPsPeP°P·P°PSPs, C%oPs PrPsPeCÍP°P·PsP·PëPS PμPeCÍC,PμPSCÍPëPIPsPjPμC,P°P±PsP»C-P·CfC”C,ChCÍCЦ PI PsCĤPíP°PSC-P·PjC- P»CĤPrPëPSPë C,P° Cf PrPsCÍP»C-PrP¶CfPIP°PSPëC... PμPeCÍPíPμCĤPëPjPμPSC,P°P»ChPSPëC... C,PIP°CĤPëPS, C- PIPëPIPsPrPëC,ChCÍCЦ P· PsCĤPíP°PSC-P·PjCf PíPμCĤPμPIP°P¶PSPs P· PeP°P»PsPj.

PŸPμCĤPμPrPSC-PN<sub>0</sub> PíPμCĤC-PsPr PSP°PíC-PIPIPëPIPμPrPμPSPSCЦ PíCĤPμPíP°CĤP°C,Cf P· PíP»P°P·PjPë CÍC,P°PSPsPIPëC,Ch 22 PíPsPrPëPSPë, C%oPs PrP°C” PjPsP¶P»PëPIC-CÍC,Ch PíCĤPëPN<sub>0</sub>PjP°C,Pë PíCĤPμPíP°CĤP°C, 1 CĤP°P· PSP° PrPsP±Cf.

PμCĤPë PíPμCĤPpsCĤP°P»ChPSPsPjCf P·P°CÍC,PsCÍCfPIP°PSPSC- PrPsPeCÍP°P·PsP·PëPSCf PePsPSC†PμPSC,CĤP°C†C-CЦ PjPμC,P°P±PsP»C-C,C-PI Cf PíP»P°P·PjC- PeCĤPpsPIC- C” PSPëP·ChPePsCĤ. PъPsPSC†PμPSC,CĤP°C†C-CЦ Cf PíP»P°P·PjC- PeCĤPpsPIC- PSP°PN<sub>0</sub>P°PeC,PëPIPsC-C€PsPiPs (6'-PiC-PrCĤPpsPeCÍPë) PjPμC,P°P±PsP»C-C,Cf Cf P»CĤPrPëPSPë PI 40 CĤP°P·C-PI PSPëP¶C†P° P·P° PíP»P°P·PjPsPICf PePsPSC†PμPSC,CĤP°C†C-CĤ PíPμCĤPIPëPSPSPsC— CÍPíPsP»CfPePë, C%oPs CÍPIC-PrC†PëC,Ch PíCĤPps C,Pμ, C%oPs P°PSC,PëPiC-PíPμCĤC,PμPSP·PëPIPsP° PrC-CЦ PíCĤPμPíP°CĤP°C,Cf P·CfPjPsPIP»PμPSP° PíPμCĤPμPIP°P¶PSPs PrPsPeCÍP°P·PsP·PëPSPsPj.

PќP° PrP°PSPëPN<sub>0</sub> C†P°CÍ C-CÍPSCfCĤC,Ch P»PëC€Pμ PsP±PjPμP¶PμPSC- PrP°PSC- C%oPsPrPs P·P°CÍC,PsCÍCfPIP°PSPSCЦ PíCĤPμPíP°CĤP°C,Cf PíP°C†C-C”PSC,P°Pj C-P· PíPsCĤCfC€PμPSPSCЦPjPë C,,CfPSPeC†C-C— PíPμC†C-PSPePë C,P° C%oPsPrPs PíPíP»PëPICf PíCĤPμPíP°CĤP°C,C-PI, P·PrP°C,PSPëC... P·PjC-PSCĤPIP°C,Pë PíPμC†C-PSPePsPIPëPN<sub>0</sub> PjPμC,P°P±PsP»C-P·Pj (PSP°PíCĤPëPeP»P°Pr, C†PëPjPμC,PëPrPëPSCf). Pj PeP»C-PSC-C†PSPsPjCf PrPsCÍP»C-PrP¶PμPSPSC- P· CfC†P°CÍC,CĤ 12 PíP°C†C-C”PSC,C-PI C-P· PíPsPjC-CĤPSPsCĤ PíPμC†C-PSPePsPIPsCĤ PrPëCÍC,,CfPSPeC†C-C”CĤ PsPrPSPsPeCĤP°C,PSPμ P·P°CÍC,PsCÍCfPIP°PSPSCЦ PrPsPeCÍP°P·PsP·PëPSCf PíCĤPëP·PIPμP»Ps PrPs PíC-PrPIPëC%oPμPSPSCЦ AUC PSP° 43 % C,P° P·PjPμPSCePμPSPSCЦ PIPëPrPëPjPsPiPs PeP»C-CĤPμPSCÍCf PíCĤPë PíPμCĤPpsCĤP°P»ChPSPsPjCf P·P°CÍC,PsCÍCfPIP°PSPSC- PSP° 40 %. PíPe C- PíCĤPë P·P°CÍC,PsCÍCfPIP°PSPSC- C-PSC€PëC... PíCĤPμPíP°CĤP°C,C-PI, C%oPs PíPsPIPsC-CÍC,CĤ PjPμC,P°P±PsP»C-P·CfCĤC,ChCÍCЦ PíPμC†C-PSPePsCĤ,

PiP°C†C-C" PSC, P°Pj P· PsP·PSP°PeP°PjPë PiPsC†C†C€PµPSC†C C,,C†PSPeC†C-C—  
PiPµC†C-PSPePë PrPsPeC†P°P·PsP·PëPS C†P»C-Pr P·P°C†C,PsC†PsPIC†PIP°C,Pë P·  
PsC†PsP±P»PëPIPsC† PsP±PµC†PµP¶PSC-C†C,C†.

P"PsPeC†P°P·PsP·PëPS PµPeC†C,PµPSC†PëPIPsPs PjPµC,P°P±PsP»C-P·C†C" C,CHC†C†C†C PI  
PiPµC†C-PSC†C-. P"PsC†P»C-PrP¶PµPSPSC†C in vitro PIPeP°P·C†C†C,C† PSP° C,Pµ, C%oPs  
PiPsP»PsPIPsPëPj C†PëPSPsPj C†Pµ P·PrC-PNëC†PSC†C" C,CHC†C†C†C P·P° PrPsPiPsPjPsPiPsC†  
C,,PµC†PjPµPSC,C† CYP 3A4, C,P° PjPµPSC€PsC† PjC-C†PµC†C†C в†" CYP 2D6 C- CYP 2C9.

**РљР»С-РSC-С†РSC- С...Р°С†Р°РёС,РµС†РёС†С,РёРёРё.**

**РµРsРёР°Р°Р°РSPSC†C.**

Р†С†С,РµС†С-Р°Р»С†РSP° PiC-PiPµC†С,РµPSP·C-C†.

РµС†СРµPiP°C†P°C, PiPsPeP°P·P°PSPëPNë PrP»C†C P»C-PeC†PIP°PSPSC†C  
P°C†С,РµС†С-Р°Р»С†РSPsC— PiC-PiPµC†С,РµPSP·C-C— C- PrP»C†C P±C-P»С†C€PsC†C,C-  
PiP°C†C-C" PSC,C-PI PNëPsPiPs PjPsP¶PSP° P·P°C†C,PsC†PsPIC†PIP°C,Pë PrP»C†C  
PePsPSC,C†PµP»C† P°C†С,РµС†С-Р°Р»С†РSPsPiPs C,PëC†PeC† C† C†PePsC†C,C-  
PjPsPSPsC,РµС†С°PiC-C—. Pj PIPëPiP°PrPeC† PSPµPµC,,PµPeC,PëPIPsPsC†C,C-  
PjPsPSPsC,РµС†С°PiC-C— PrP»C†C P»C-PeC†PIP°PSPSC†C P°C†С,РµС†С-Р°Р»С†РSPsC—  
PiC-PiPµC†С,РµPSP·C-C— PiC†PµPiP°C†P°C, PjPsP¶PSP° P·P°C†C,PsC†PsPIC†PIP°C,Pë C†  
PePsPjP±C-PSP°C†C-C— P· C,C-P°P·PëPrPSPëPjPë PrC-C†C†PµC,PëPeP°PjPë,  
P±P»PsPeP°C,PsC†P°PjPë β-P°PrC†PµPSPsC†PµC†PµPiC,PsC†C-PI, P±P»PsPeP°C,PsC†P°PjPë  
PeP°P»С†C†C-C" PIPëC... PeP°PSP°P»C-PI C,P° C-PSPiC-P±C-C,PsC†P°PjPë  
P°PSPiC-PsC,РµPSP·PëPSPiPµC†PµC,PIPsC†C†PIP°P»С†РSPsPiPs C,,PµC†PjPµPSC,C†.

P"PsP±C†PµC†C†C-PeC-C†PSP° PiC-PiPµC†PiP»P°P·C-C† PiPµC†PµPrPjC-C...C†C†PsPIPsC—  
P·P°P»PsP·Pë.

РµС†СРµPiP°C†P°C, PiPsPeP°P·P°PSPëPNë PrP»C†C P»C-PeC†PIP°PSPSC†C  
PsP±C†C,C†C†PeC†C-C— C†PµC†PsPIPëPIC-PrPSPëC... C€P»C†C...C-PI, P° C,P°PePsP¶  
C†PëPjPiC,PsPjC-PI, PiPsPiB†™ C†P·P°PSPëC... C-P· PrPsP±C†PµC†C†C-PeC-C†PSPsC†  
PiC-PiPµC†PiP»P°P·C-C" C† PiPµC†PµPrPjC-C...C†C†PsPIPsC— P·P°P»PsP·Pë (P"P"PµP—).  
РµС†СРµPiP°C†P°C, PjPsP¶PSP° PiC†PëP·PSP°C†P°C,Pë PiP°C†C-C" PSC,P°Pj C-P· P"P"PµP— C†Pe  
PiC†Pë PSP°C†PIPsPsC†C,C- P°C†С,РµС†С-Р°Р»С†РSPsC— PiC-PiPµC†С,РµPSP·C-C—, C,P°Pe  
C- PiC†Pë PSPsC†PjP°P»С†РSPsPjC† C†С-PIPsC- P°C†С,РµС†С-Р°Р»С†РSPsPiPs C,PëC†PeC†.

**РµС†PµC,PëPiPsPeP°P·P°PSPSC†C.**

P—P°C†C,PsC†C†PIP°PSPSC†C PiC†PµPiP°C†P°C,C† РљР°Р°С†P†C†C†P°<sup>B®</sup>  
PiC†PµC,PëPiPsPeP°P·P°PSPµ PSP°C†C,C†PiPSPëPj PeP°C,РµPiPsC†С-C†Pj PiP°C†C-C" PSC,C-PI:

в†" PiP°C†C-C" PSC,P°Pj P· PiC-PiPµC†C†C†C,P»PëPIC-C†C,C† PrPs PrC-C†C†PsC—  
C†PµC†PsPIPëPSPë C†Pë PrPs PiPsC...C-PrPSPëC... C...C-PSP°P·PsP»C-PSC†  
(PSP°PiC†PëPeP»P°Pr, PiC†P°P·PsP·PëPSC†, C,РµС†С°P·PsP·PëPSC†, PrPsPeC†P°P·PsP·PëPSC†)  
P°P±Ps PrPs P±C†PrC†C-C†PePsC— P· PrPsPiPsPjC-P¶PSPëC... C†PµC†PsPIPëPS  
PiC†PµPiP°C†P°C,C†, P·P°P·PSP°C†PµPSPëC... C† C†PµP·PrC-P»C- B«P†PeP»P°PrB»;

вґґ“ PIP°C†C-C”PSC,P°Pj C-P· PIPëPIP°PrPeP°PjPë PsCґґC,PsCґґC,P°C,PëC†PSPsC—  
PiC-PiPsC,PµPSP·C-C— PI P°PSP°PjPSPµP·C-;

вґґ“ PIP°C†C-C”PSC,P°Pj C-P· P”P”PµP— C,P° CґґC†PіC†C,PSCґґPsCґґ PsP±CґґC,CґґC†PeC†C-C”Cґґ  
PIPµCґґC...PSC-C... CґґPµC†PsPIPëPIC-PrPSPëC... CëP»CґґC...C-PI, C...CґґPsPSC-C†PSPëPjPë  
C-PSC,,PµPeC†C-CґґPjPë CґґPµC†PsPIPëPIC-PrPSPëC... CëP»CґґC...C-PI P°P±Ps  
PSP°CґґPIPSC-CґґC,Cґґ PeP°PjPµPSC-PI C† CґґPµC†PsPIPSPjC† PjC-C...C†CґґC-;

вґґ“ PIP°C†C-C”PSC,P°Pj P· P°CґґC,PµCґґC-P°P»CґґPSPsCґґ PiC-PiPsC,PµPSP·C-Cґґ  
(CґґC,PsCґґC†C”C,CґґCґґCґґ C,C-P»CґґPePë PIP°C†C-C”PSC,C-PI C-P· P”P”PµP—).

P”PsPeCґґP°P·PsP·PëPS C† CґґPePsCґґC,C- PjPsPSPsC,PµCґґP°PiC-C—  
PiCґґPsC,PëPiPsPeP°P·P°PSPëPµ PIP°C†C-C”PSC,P°Pj C-P· PiPµCґґPµPiPsPIPSPµPSPSCґґPj  
CґґPµC†PsPIPSPiPs PjC-C...C†CґґP° P°P±Ps P°PSC†CґґC-C”Cґґ C-P· PiCґґPsPiCґґPµCґґC†PsCґґ  
PSPëCґґPePsPIPSPsCґґ PSPµPrPsCґґC,P°C,PSC-CґґC,Cґґ P°P±Ps P±PµP· PSPµC—.

**P”P·P°C”PjPsPrC-Cґґ P· C-PSCëPëPjPë P»C-PeP°CґґCґґCґґPePëPjPë P·P°CґґPsP±P°PjPë C,P°  
C-PSCëC- C,,PsCґґPjPë PIP·P°C”PjPsPrC-PNë.**

P†PSPiC-P±C-C,PsCґґPë C,,PsCґґC,,PsPrC-PµCґґC,PµCґґP°P·Pë-5 (PαP”P·-5) (PSP°PiCґґPëPeP»P°Pr,  
CґґPëP»PrPµPSP°C,,C-P», C,P°PrP°P»P°C,,C-P», PIP°CґґPrPµPSP°C,,C-P»).

PґґC†PjC-CґґPSPµ P·P°CґґC,PsCґґC†PIP°PSPSCґґ PrPsPeCґґP°P·PsP·PëPSC† P·  
C-PSPiC-P±C-C,PsCґґP°PjPë PαP”P·-5 C† PrPµCґґPePëC... PIP°C†C-C”PSC,C-PI PjPsP¶Pµ  
CґґPiCґґPëC†PëPSCґґC,Pë CґґPëPjPiC,PsPjP°C,PëC†PSC† PiC-PiPsC,PµPSP·C-Cґґ (PrPëPi.  
CґґPsP·PrC-P» B«PґґCґґPsP±P»PëPIPsCґґC,C- P·P°CґґC,PsCґґC†PIP°PSPSCґґC»).

P”PsCґґP»C-PrP¶PµPSCґґ PrPsPeCґґP°P·PsP·PëPSC† C† P»C-PeP°CґґCґґCґґPePëC... C,,PsCґґPjP°C...  
C-P· C,CґґPëPIP°P»PëPj PIPëPIC-P»CґґPSPµPSPSCґґPj PrC-CґґC†PsC— CґґPµC†PsPIPëPSPë PSPµ  
PiCґґPsPIPSPPrPëP»Pë.

P”PsPeCґґP°P·PsP·PëPS P·PSP°C†PSPsCґґ PjC-CґґPsCґґ P·PiBґґ™ CґґP·C†C”C,CґґCґґCґґ P·  
P±C-P»PeP°PjPë PIP»P°P·PjPë PeCґґPsPIC- (PSP° 98 %). P PµP·C†P»CґґC,P°C,Pë  
PrPsCґґP»C-PrP¶PµPSPSCґґ in vitro P· PIPëPePsCґґPëCґґC,P°PSPSCґґPj PIP»P°P·PjPë PeCґґPsPIC-  
P»CґґPrPëPSPë CґґPIC-PrC†P°C,Cґґ, C%Ps PiCґґPµPiP°CґґP°C, PSPµ PIPIP»PëPIP°C” PSP°  
P·PiCґґP·C†PIP°PSPSCґґ PrPsCґґP»C-PrP¶C†PIP°PSPëC... PiCґґPµPiP°CґґP°C,C-PI  
(PrPëPiPsPeCґґPëPS, C,,PµPSC-C,PsC—PS, PIP°CґґC,,P°CґґPëPS P°P±Ps C-PSPrPsPjPµC,P°C†PëPS)  
P· P±C-P»PeP°PjPë.

Pj PeP»C-PSC-C†PSC-PNë PiCґґP°PeC,PëC†C- PSPµ PIC-PrP·PSP°C†P°P»PsCґґCґґ  
PSPµCґґPiCґґPëCґґC,P»PëPIPsC— PIP·P°C”PjPsPrC-C— PiCґґPë PsPrPSPsC†P°CґґPSPSPjC†  
P·P°CґґC,PsCґґC†PIP°PSPSC- PrPsPeCґґP°P·PsP·PëPSC† C,P° C,C-P°P·PëPrPSPëC...  
PrC-C†CґґPµC,PëPeC-PI, C,,C†CґґPsCґґPµPjC-PrC†, P±P»PsPeP°C,PsCґґC-PI β-  
P°PrCґґPµPSPsCґґPµC†PµPiC,PsCґґC-PI, PSPµCґґC,PµCґґPsC—PrPSPëC...  
PiCґґPsC,PëP·P°PiP°P»CґґPSPëC... P·P°CґґPsP±C-PI, P°PSC,PëP±C-PsC,PëPeC-PI,  
PiPµCґґPsCґґP°P»CґґPSPëC... C†C†PeCґґPsP·PSPëP¶C†PIP°P»CґґPSPëC... P·P°CґґPsP±C-PI,  
C†CґґPëPePsP·C†CґґPëC†PSPëC... P·P°CґґPsP±C-PI C,P° P°PSC,PëPePsP°PiC†P»CґґPSC,C-PI.  
PµCґґPsC,Pµ PrP°PSC- C,,PsCґґPjP°P»CґґPSPëC... PrPsCґґP»C-PrP¶PµPSCґґ C-P·  
PIPëPIC†PµPSPSCґґ P»C-PeP°CґґCґґCґґPePsC— PIP·P°C”PjPsPrC-C— PIC-PrCґґC†C,PSC-.

P”PsCґґP»C-PrP¶PµPSPSCґґ in vitro PIPeP°P·C†CґґC,Cґґ PSP° C,Pµ, C%Ps PrPsPeCґґP°P·PsP·PëPS

C" CÍCfP±CÍC,CĤP°C,PsPj C†PëC,PsC...CĤPsPjCf P450 3A4 (CYP 3A4). PŸP»C-Pr P·PsP±PμCĤPμP¶PSC-CÍC,CĤ PiCĤPëP·PSP°C†P°C,Pë PrPsPeCÍP°P·PsP·PëPS PsPrPSPsC†P°CÍPSPs P·PiPsC,CfP¶PSPëPjPë C-PSPiC-P±C-C,PsCĤP°PjPë CYP 3A4, PSP°PiCĤPëPeP»P°Pr PeP»P°CĤPëC,CĤPsPjC-C†PëPSPsPj, C-PSPPrPëPSP°PIC-CĤPsPj, C-C,CĤP°PePsPSP°P·PsP»PsPj, PePμC,PsPePsPSP°P·PsP»PsPj, PSPμC,,P°P·PsPrPsPSPsPj, PSPμP»C,,C-PSP°PIC-CĤPsPj, CĤPëC,PsPSP°PIC-CĤPsPj, CÍP°PePIC-PSP°PIC-CĤPsPj, C,PμP»C-C,CĤPsPjC-C†PëPSPsPj P°P±Ps PIPsCĤPëPePsPSP°P·PsP»PsPj (PrPëPI. CĤPsP·PrC-P» B«PαP°CĤPjP°PePsP»PsPiC-C†PSC-PIP»P°CÍC,PëPIPSCÍC,C-B»).

P"PsPeCÍP°P·PsP·PëPS PiPsC,PμPSC†C-CĤC" PiC-PiPsC,PμPSP·PëPIPSCf PrC-CĤ C-PSCëPëC... α-P°PrCĤPμPSPsP±P»PsPeP°C,PsCĤC-PI, P° C,P°PePsP¶ C-PSCëPëC... P°PSC,PëPiC-PiPμCĤC,PμPSP·PëPIPSPëC... P·P°CÍP±C-PI.

P—P° CĤPμP·CfP»CĤC,P°C,P°PjPë PIC-PrPeCĤPëC,PsPiPs CĤP°PSPPrPsPjC-P·PsPIP°PSPsPiPs PiP»P°C†PμP±Ps-PePsPSC,CĤPsP»CĤPsPIP°PSPsPiPs PeP»C-PSC-C†PSPsPiPs PrPsCÍP»C-PrP¶PμPSPSCĬ P· CfC†P°CÍC,CĤ 22 P·PrPsCĤPsPIPëC... PrPsP±CĤPsPIPsP»CĤC†C-PI C†PsP»PsPIC-C†PsC— CÍC,P°C,C-, PsPrPSPsPeCĤP°C,PSPμ P·P°CÍC,PsCÍCfPIP°PSPSCĬ PrPsPeCÍP°P·PsP·PëPSCf Cf PrPsP·C- 1 PjPi Cf PiPμCĤCëPëPNë PrPμPSCĤ C†PsC,PëCĤPëPrPsP±PsPIPsPiPs PeCfCĤCÍCf PiPμCĤPsCĤP°P»CĤPSPsPiPs P·P°CÍC,PsCÍCfPIP°PSPSCĬ C†PëPjPμC,PëPrPëPSCf (400 PjPi PrPIC-C†C- PSP° PrPsP±Cf) PiCĤPëP·PIPsPrPëP»Ps PrPs PiC-PrPIPëC%PμPSPSCĬ CÍPμCĤPμPrPSCĤPsPiPs CĤC-PIPSCĬ AUC PrPsPeCÍP°P·PsP·PëPSCf PSP° 10 % C- PSPμ CÍPiCĤPëC†PëPSCĬP»Ps P¶PsPrPSPëC... CÍC,P°C,PëCÍC,PëC†PSPs P·PSP°C†CfC%PëC... P·PjC-PS CÍPμCĤPμPrPSCĤPsC— C<sub>max</sub> C,P° CÍPμCĤPμPrPSCĤPsPiPs PiPμCĤC-PsPrCf PSP°PiC-PIPIPëPIPμPrPμPSPSCĬ PrPsPeCÍP°P·PsP·PëPSCf. PŸP°PePμ PiC-PrPIPëC%PμPSPSCĬ CÍPμCĤPμPrPSCĤPsPiPs CĤC-PIPSCĬ AUC PrPsPeCÍP°P·PsP·PëPSCf PSP° 10 % PSP° C,P»C- P·P°CÍC,PsCÍCfPIP°PSPSCĬ C†PëPjPμC,PëPrPëPSCf P·PSP°C...PsPrPëC,CĤCÍCĬ Cf PjPμP¶P°C... PjC-P¶C-PSPPrPëPIC-PrCfP°P»CĤPSPsC— PIP°CĤC-P°P±PμP»CĤPSPsCÍC,C- (27 %) CÍPμCĤPμPrPSC-C... CĤC-PIPSC-PI AUC PrPsPeCÍP°P·PsP·PëPSCf PiPsCĤC-PIPSCĬPSPs P·PiP»P°C†PμP±Ps.

***PĥCÍPsP±P»PëPIPsCÍC,C- P·P°CÍC,PsCÍCáPIP°PSPSCĬ.***

*PĥCĤC,PsCÍC,P°C,PëC†PSP° PiC-PiPsC,PμPSP·C-CĬ/CÍPëPSPePsPiPμ.*

PμPsC†P°C,PsPe C,PμCĤP°PiC-C—. PiPe PSP°CÍP»C-PrPsPe PiCĤPsCĬPICf PrC-C— P±P»PsPeP°C,PsCĤC-PI α-P°PrCĤPμPSPsCĤPμC†PμPiC,PsCĤC-PI, PjPsP¶Pμ CĤPsP·PIPëPSCfC,PëCÍCĬ PsCĤC,PsCÍC,P°C,PëC†PSP° PiC-PiPsC,PμPSP·C-CĬ, PiCĤPsCĬPIP»CĬCĤC†PëCÍCĤ P·P°PiP°PjPsCĤPsC†PμPSPSCĬPj C,P° CÍP»P°P±PeC-CÍC,CĤ P°P±Ps CĤC-PrCëPμ BĤ" PIC,CĤP°C,PsCĤ CÍPIC-PrPsPjPsCÍC,C- (CÍPëPSPePsPiPμ), PsCÍPsP±P»PëPIPs PSP° PiPsC†P°C,PeCf C,PμCĤP°PiC-C— (PrPëPI. CĤPsP·PrC-P» B«PŸPiPsCÍC-P± P·P°CÍC,PsCÍCfPIP°PSPSCĬ C,P° PrPsP·PëB»). Pj P·PiBĤ™ CĬP·PeCf P· C†PëPj PSP° PiPsC†P°C,PeCf C,PμCĤP°PiC-C— PSPμPsP±C...C-PrPSPs PePsPSC,CĤPsP»CĤPIP°C,Pë CĤC-PIPμPSCĤ P°CĤC,PμCĤC-P°P»CĤPSPsPiPs C,PëCÍPeCf, C%PsP± PjC-PSC-PjC-P·CfPIP°C,Pë PjPsP¶P»PëPIC- PiPsCÍC,CfCĤP°P»CĤPSC- PμC,,PμPeC,Pë.

PμCĤPëP·PSP°C†P°CĤC†Pë C,PμCĤP°PiC-CĤ P±CfPrCĤ-CĬPePëPj PμC,,PμPeC,PëPIPSPëPj P±P»PsPeP°C,PsCĤPsPj α-P°PrCĤPμPSPsCĤPμC†PμPiC,PsCĤC-PI, CÍP»C-Pr PiCĤPsC-PSC,,PsCĤPjCfPIP°C,Pë PiP°C†C-C" PSC,P°, CĬPe CfPSPëPePSCfC,Pë CÍPëPjPiC,PsPjC-PI PsCĤC,PsCÍC,P°C,PëC†PSPsC— PiC-PiPsC,PμPSP·C-C— C,P° CĬPe

PiPsPIPsPrPëC, PëCíCñ PiCñPë C—C... PIPëPSPëPePSPµPSPSC-. PµP°C†C-C" PSC, P° C, P°PePsP¶  
PiPsC, CñC-P±PSPs PiPsPiPµCñPµPrPëC, Pë PiCñPs PSPµPsP±C... C-PrPSC-CíC, Cñ  
CíPSPëPePSPµPSPSC¶ CíPëC, CíP°C†C-Pñ, Cí C¶PePëC... C-CíPSCíC" CñPëP·PëPe  
PsC, CñPëPjP°PSPSC¶ C, CñP°PIPj, PICñP°C... PsPICíCñC†Pë PjPsP¶P»PëPIC-CíC, Cñ  
PIPëPSPëPePSPµPSPSC¶ P·P°PiP°PjPsCñPsC†PµPSPSC¶ P°P±Ps CíP»P°P±PePsCíC, C- PSP°  
PiPsC†P°C, PeCí C, PµCñP°PiC-C— PiCñPµPiP°CñP°C, PsPj PñP°CñPrCíCñP°<sup>B®</sup>.

*P—P°CíC, PsCíC¿PIP°PSPSC¶ PiCñPë PiPsCíC, CñPëC... CíPµCñC†PµPIPëC... CíC, P°PSP°C....*

PiPe C- C-PSC€C- PIP°P·PsPrPëP»P°C, CíCñC†C- P°PSC, PëPiC-PiPµCñC, PµPSP·PëPIPSC-  
P·P°CíPsP±Pë, PrPsPeCíP°P·PsP·PëPS CíP»C-Pr P· PsP±PµCñPµP¶PSC-CíC, Cñ  
P·P°CíC, PsCíPsPICíPIP°C, Pë PiP°C†C-C" PSC, P°Pj C-P· PSP°CíC, CíPiPSPëPjPë PiPsCíC, CñPëPjPë  
CíPµCñC†PµPIPëPjPë CíC, P°PSP°PjPë:

вґ“ P»PµPiPµPSPµPIPëPñ PSP°P±CñC¶Pe, CíPiCñPëC†PëPSPµPSPëPñ P°PsCñC, P°P»CñPSPëPj  
P°P±Ps PjC-C, CñP°P»CñPSPëPj CíC, PµPSPsP·PsPj;

вґ“ PiC-PiPµCñCíPëCíC, PsP»C-C†PSP° CíPµCñC†PµPIP° PSPµPrPsCíC, P°C, PSC-CíC, Cñ;

вґ“ PiCñP°PIPsC€P»CíPSPsC†PePsPIP° CíPµCñC†PµPIP° PSPµPrPsCíC, P°C, PSC-CíC, Cñ,  
P·CíPjPsPIP»PµPSP° C, CñPsPjP±PsPµPjP±PsP»C-C" Cñ P»PµPiPµPSPµPIPsC— P°CñC, PµCñC-C—  
P°P±Ps PiPµCñPëPeP°CñPrC-P°P»CñPSPëPj PIPëPiPsC, PsPj;

вґ“ P»C-PIPsC€P»CíPSPsC†PePsPIP° CíPµCñC†PµPIP° PSPµPrPsCíC, P°C, PSC-CíC, Cñ P·  
PSPëP·CñPePëPj C, PëCíPePsPj PSP°PiPsPIPSPµPSPSC¶.

*P—P°CíC, PsCíC¿PIP°PSPSC¶ PiCñPë PiPsCñC¿C€PµPSPSC- C,, C¿PSPeC†C-Pñ PiPµC†C-PSPePë.*

PiPe C- PiCñPë P·P°CíC, PsCíCíPIP°PSPSC- C-PSC€PëC... PiCñPµPiP°CñP°C, C-PI, C%oPs  
PiPsPIPSC-CíC, Cñ PjPµC, P°P±PsP»C-P·CíCñC, CñCíC¶ PiPµC†C-PSPePsCñ,  
PiP°C†C-C" PSC, P°Pj P· PsP·PSP°PeP°PjPë PiPsCñCíC€PµPSPSC¶ C,, CíPSPeC†C-Pñ  
PiPµC†C-PSPePë PiCñPµPiP°CñP°C, PñP°CñPrCíCñP°<sup>B®</sup> CíP»C-Pr PiCñPëP·PSP°C†P°C, Pë P·  
PsCíPsP±P»PëPIPsCñ PsP±PµCñPµP¶PSC-CíC, Cñ (PrPëPI. CñPsP·PrC-P» B«PñPiPsCíC-P±  
P·P°CíC, PsCíCíPIP°PSPSC¶ C, P° PrPsP·PëB»). PñPµCñPµP· PIC-PrCíCíC, PSC-CíC, Cñ  
PeP»C-PSC-C†PSPsPiPs PrPsCíPIC-PrCí P·P°CíC, PsCíCíPIP°PSPSC¶ PiCñPµPiP°CñP°C, Cí  
PiP°C†C-C" PSC, P°Pj C-P· PiPµC†C-PSPePsPIPsCñ PSPµPrPsCíC, P°C, PSC-CíC, Cñ  
C, C¶P¶PePsPiPs CíC, CíPiPµPSC¶ PiCñPëP·PSP°C†PµPSPSC¶ PiCñPµPiP°CñP°C, Cí C†C-Pñ  
PeP°C, PµPiPsCñC-C— PiP°C†C-C" PSC, C-PI PSPµ CñPµPePsPjPµPSPrCíC" C, CñCíC¶.

*P—P°CíC, PsCíC¿PIP°PSPSC¶ CñP°P·PsPj P· C-PSPiC-P±C-C, PsCñP°PjPë PαP" P·-5.*

P—P°CíC, PsCíCíPIP°PSPSC¶ Pë PrPsPeCíP°P·PsP·PëPS CñP°P·PsPj P· C-PSPiC-P±C-C, PsCñP°PjPë  
C,, PsCíC,, PsPrC-PµCíC, PµCñP°P·Pë-5 (PSP°PiCñPëPeP»P°Pr, CíPëP»PrPµPSP°C,, C-P»,  
C, P°PrP°P»P°C,, C-P» C, P° PIP°CñPrPµPSP°C,, C-P») CíP»C-Pr P· PsP±PµCñPµP¶PSC-CíC, Cñ,  
PsCíPeC-P»CñPePë PsP±PëPrPIP° P»C-PeP°CñCíCñPeC- P·P°CíPsP±Pë  
CíPiCñPëC†PëPSC¶CñC, Cñ PIP°P·PsPrPëP»P°C, P°C†C-Cñ, P° PsC, P¶Pµ вґ“ PjPsP¶CíC, Cñ  
CíPiCñPëC†PëPSPëC, Pë Cí PrPµC¶PePëC... PiP°C†C-C" PSC, C-PI CíPëPjPiC, PsPjP°C, PëC†PSCí  
PiC-PiPsC, PµPSP·C-Cñ. P" P»C¶ P·PjPµPSC€PµPSPSC¶ CñPëP·PëPeCí CñPsP·PIPëC, PeCí  
PsCñC, PsCíC, P°C, PëC†PSPsC— PiC-PiPsC, PµPSP·C-C— CñPµPePsPjPµPSPrCíC" C, CñCíC¶  
CñPsP·PiPsC†PëPSP°C, Pë C, PµCñP°PiC-Cñ PiCñPµPiP°CñP°C, P°PjPë PiCñCíPiPë  
C-PSPiC-P±C-C, PsCñC-PI C,, PsCíC,, PsPrC-PµCíC, PµCñP°P·Pë-5 C, C-P»CñPePë P·P° CíPjPsPIPë,  
C¶PeC%oPs PSP° C, P»C- P·P°CíC, PsCíCíPIP°PSPSC¶ α-P±P»PsPeP°C, PsCñC-PI PiP°C†C-C" PSC,

PjP°C" CÍC,P°P±C-P»CHPSCr PiPµPjPsPrPëPSP°PjC-PeCí. PъCĤC-Pj C,PsPiPs, CĤPµPePsPjPµPSPPrCíC"C,CHCÍCĬ CĤPsP·PíPsC±PëPSP°C,Pë C,PµCĤP°PíC-CĤ C-PSPiC-P±C-C,PsCĤP°PjPë C,,PsCÍC,,PsPrC-PµCÍC,PµCĤP°P·Pë-5 P· PSP°PNëPjPµPSCëPsC— PjPsP¶P»PëPIPsC— PrPsP·Pë C,P° PIPëC,CĤPëPjCíPIP°C,Pë 6-PiPsPrPëPSPSPëPNë C-PSC,PµCĤPIP°P» PjC-P¶ P·P°CÍC,PsCÍCíPIP°PSPSCĬPj PrPsPeCÍP°P·PsP·PëPSCr C,P° C-PSPiC-P±C-C,PsCĤC-PI C,,PsCÍC,,PsPrC-PµCÍC,PµCĤP°P·Pë-5. P"PsCÍP»C-PrP¶PµPSCĤ PrPsPeCÍP°P·PsP·PëPSCr Cí P»C-PeP°CĤCÍCHPePëC... C,,PsCĤPjP°C... C-P· C,CĤPëPIP°P»PëPj PIPëPIC-P»CHPSPµPSPSCĬPj PrC-CĤC±PsC— CĤPµC±PsPIPëPSPë PSPµ PíCĤPsPIPsPrPëP»Pë.

*P—P°CÍC,PsCÍCíPIP°PSPSCĬ PíP°C†C-C" PSC,P°Pj PíC-Pr C±P°CÍ C...C-CĤCíCĤPiC-C±PSPsPiPs PIPëPrP°P»PµPSPSCĬ PeP°C,P°CĤP°PeC,Pë.*

Pj PrPµCĬPePëC... PíP°C†C-C" PSC,C-PI, CĬPeC- PíCĤPëPNëPjP°P»Pë C,P°PjCÍCíP»PsP·PëPS PSP° C±P°CÍ PíCĤPsPIPµPrPµPSPSCĬ C...C-CĤCíCĤPiC-C±PSPsPiPs PIPëPrP°P»PµPSPSCĬ PeP°C,P°CĤP°PeC,Pë P°P±Ps PrPs PsPíPµCĤP°C†C-C—, PíC-Pr C±P°CÍ PIC,CĤCíC±P°PSPSCĬ CÍPíPsCÍC,PµCĤC-PiP°PICÍCĬ CĤPsP·PIPëC,PsPe C-PSC,CĤP°PsPíPµCĤP°C†C-PNëPSPsPiPs CÍPëPSPPrCĤPsPjCí PíBĤ™ CĬP»PsC— CĤP°PNëPrCíP¶PSPsC— PsP±PsP»PsPSPëPë (P†PŸPќP Pĥ, PIP°CĤC-P°PSC, CÍPëPSPPrCĤPsPjCí PICíP·CHPePsC— P·C-PSPëC†C-). PµPsPIC-PrPsPjP»CĬP»PsCÍCĬ PíCĤPs PsPeCĤPµPjC- PIPëPíP°PrPePë CĤPsP·PIPëC,PeCí C,P°PePsPiPs PíPsP±C-C±PSPsPiPs PµC,,PµPeC,Cr C- PíCĤPë P·P°CÍC,PsCÍCíPIP°PSPSC- C-PSCëPëC... α<sub>1</sub>-P±P»PsPeP°C,PsCĤC-PI, C,PsPjCí PSPµ PjPsP¶PSP° PIPëPeP»CĤC±P°C,Pë PjPsP¶P»PëPIC-CÍC,CH CĤPsP·PIPëC,PeCí C†CHPsPiPs PµC,,PµPeC,Cr PrP»CĬ C-PSCëPëC... PíCĤPµPíP°CĤP°C,C-PI PrP°PSPsPiPs PeP»P°CÍCí P»C-PeP°CĤCÍCHPePëC... P·P°CÍPsP±C-PI. PŞPµCĤPµP· C,Pµ, C%Ps P†PŸPќP Pĥ PjPsP¶Pµ PíCĤPëP·PIPsPrPëC,Pë PrPs PíC-PrPIPëC%PµPSPSCĬ C±P°CÍC,PsC,Pë PíCĤPsC†PµPrCíCĤPSPëC... CíCÍPeP»P°PrPSPµPSCĤ PíCĤPë PíCĤPsPIPµPrPµPSPSCĬ PsPíPµCĤP°C†C-C—, PíCĤPë PíC-PrPiPsC,PsPIC†C- PrPs PSPµC— CÍP»C-Pr PíPsPIC-PrPsPjPëC,Pë C...C-CĤCíCĤPiP°Pj-PsC,,C,P°P»CHPjPsP»PsPíP°Pj, C±Pë P·P°CÍC,PsCÍPsPICíC" P°P±Ps P·P°CÍC,PsCÍPsPICíPIP°PI PíP°C†C-C" PSC, P±P»PsPeP°C,PsCĤPë α<sub>1</sub>-P°PrCĤPµPSPsCĤPµC†PµPíC,PsCĤC-PI.

*PµCĤC-P°PíC-P·Pj.*

Pj PíPµCĤC-PsPr PíC-CÍP»CĬ PIPëC...PsPrCí PíCĤPµPíP°CĤP°C,Cr PSP° CĤPëPSPsPe PíC-Pr C±P°CÍ P·P°CÍC,PsCÍCíPIP°PSPSCĬ α<sub>1</sub>-P±P»PsPeP°C,PsCĤC-PI, Cr C,PsPjCí C±PëCÍP»C- PrPsPeCÍP°P·PsP·PëPSCr PSP°PrC...PsPrPëP»Pë PíPsPIC-PrPsPjP»PµPSPSCĬ PíCĤPs PIPëPíP°PrPePë C,CĤPëPIP°P»PsC— PµCĤPµPeC†C-C— C,P° PíCĤC-P°PíC-P·PjCí. Pj PIPëPíP°PrPeCí PµCĤPµPeC†C-C—, C%Ps C,CĤPëPIP°C" P±C-P»CHCëPµ 4 PiPsPrPëPS, PíP°C†C-C" PSC,Cr CÍP»C-Pr PSPµPíP°PNëPSPs P·PIPµCĤPSCrC,PëCÍCĬ P·P° PjPµPrPëC±PSPsCĤ PrPsPíPsPjPsPiPsCĤ. PíPeC%Ps PSPµPíP°PNëPSPs PSPµ CĤPsP·PíPsC±P°C,Pë P»C-PeCíPIP°PSPSCĬ PíCĤC-P°PíC-P·PjCí, PIPëPSPëPeP°C" CíCëPePsPrP¶PµPSPSCĬ C,PeP°PSPëPSPë CÍC,P°C,PµPIPsPiPs C±P»PµPSP°, C%Ps PjPsP¶Pµ PíCĤPëP·PIPµCÍC,Pë PrPs PSPµP·PIPsCĤPsC,PSPsC— PIC,CĤP°C,Pë PíPsC,PµPSC†C-C—.

*PŸPeCĤPëPSC-PSPi PSP° CĤP°Pe PíPµCĤPµPrPjC-C...CíCĤPsPIPsC— P·P°P»PsP·Pë.*

PъP°CĤC†PëPSPsPjP° PíPµCĤPµPrPjC-C...CíCĤPsPIPsC— P·P°P»PsP·Pë CÍPíCĤPëC±PëPSCĬC" P±P°PíP°C,Ps CÍPëPjPíC,PsPjC-PI, C%Ps P°CÍPsC†C-CĤCĤC,CHCÍCĬ C-P· P" P" PµP—, PrPs C,PsPiPs P¶ C†C- PrPIP° P·P°C...PIPsCĤCĤPIP°PSPSCĬ PjPsP¶CíC,CH CÍPíC-PIC-CÍPSCrPIP°C,Pë. PŸPsPjCí CÍP»C-Pr PIPëPeP»CĤC±PëC,Pë PSP°CĬPIPsC-CÍC,CH PeP°CĤC†PëPSPsPjPë PíPµCĤPµPrPjC-C...CíCĤPsPIPsC— P·P°P»PsP·Pë PíPµCĤCë PSC-P¶ PíPsC±PëPSP°C,Pë C,PµCĤP°PíC-CĤ PrPsPeCÍP°P·PsP·PëPSPsPj P· PíCĤPëPIPsPrCí CÍPëPjPíC,PsPjC-PI P" P" PµP—.

*P†PSC,,PsCḡPjP°C†C-Cl C%oPsPrPs PrPsPİPsPjC-P¶PSPëC... CḡPµC‡PsPIPëPS.*

PµP°C†C-C" PSC,P°Pj C-P· CḡC-PrPeC-ClPSPëPjPë ClPİP°PrPePsPIPëPjPë  
P·P°C...PIPsCḡCḡPIP°PSPSCİPjPë, C,P°PePëPjPë ClPe PSPµPİPµCḡPµPSPScİPëPjC-ClC,CH  
PİP°P»P°PeC,PsP·Pë, PİPsPIPSP° PSPµPrPsClC,P°C,PSC-ClC,CH P»P°PeC,P°P·Pë C,P°  
PjP°P»CHP°P±ClPsCḡP±C†C-Cl PİP»CḡPePsP·Pë-PİP°P»P°PeC,PsP·Pë, PSPµ ClP»C-Pr  
P·P°ClC,PsClPsPICrPIP°C,Pë C†PµPNë PİCḡPµPİP°CḡP°C,.

PİPµPNë P»C-PeP°CḡClCHPePëPNë P·P°ClC-P± PjC-ClC,PëC,CH PjPµPSC€Pµ 1 PjPjPsP»CH  
PSP°C,CḡC-Cḡ (23 PjPi) PSP° C,P°P±P»PµC,PeC†, C,PsP±C,Ps PİCḡP°PeC,PëC‡PSPs  
PIC-P»CHPSPëPNë PIC-Pr PSP°C,CḡC-Cḡ.

***P—P°ClC,PsClCžPIP°PSPSCİ Cž PİPµCḡC-PsPr PIP°PİC-C,PSPScİC,C- P°P±Ps  
PİPsPrCžPIP°PSPSCİ PİCḡCžPrPrCḡ.***

*PµP°C†C-C" PSC,Pë P· P°CḡC,PµCḡC-P°P»CHPSPsCḡ PİC-PİPµCḡC,PµPSP·C-C" Cḡ.*

P°PİC-C,PSC-ClC,CH.

PŞPµCḡPµP· PIC-PrClC†C,PSC-ClC,CH PSP° PrP°PSPëPNë C‡P°ClC PSP°P»PµP¶PSPëC... C,P°  
PrPsP±CḡPµ PePsPSC,CḡPsP»CHPsPIP°PSPëC... PrPsClP»C-PrP¶PµPSCḡ C%oPsPrPs  
PIPëPIC‡PµPSPSCİ P·P°ClC,PsClC†PIP°PSPSCİ PİCḡPµPİP°CḡP°C,Cl PIP°PİC-C,PSPëPj,  
P±PµP·PİPµC‡PSC-ClC,CH P·P°ClC,PsClC†PIP°PSPSCİ PİCḡPµPİP°CḡP°C,Cl  
PµP°CḡPrC†CḡP°<sup>B®</sup> Cl PİPµCḡC-PsPr PIP°PİC-C,PSPScİC,C- P·P°P»PëC€P°C" C,CHClCİ  
PSPµPIC†C,P°PSPsPIP»PµPSPsCḡ. PŷPsPjC† PİCḡPµPİP°CḡP°C, ClP»C-Pr  
P·P°ClC,PsClPsPICrPIP°C,Pë P»PëC€Pµ C,PsPrC-, PePsP»Pë PİPsC,PµPSC†C-PNëPSC-  
PİPµCḡPµPIP°PİPë P»C-PeC†PIP°PSPSCİ, P· C,PsC‡PePë P·PsCḡC† P»C-PeP°CḡCİ,  
PIPëPİCḡP°PIPPrPsPIC†CḡC,CH PİPsC,PµPSC†C-PNëPSPëPNë CḡPëP·PëPe. P†PsC‡P° Cl  
PrPsClP»C-PrP¶PµPSPSCİC... PSP° C,PIP°CḡPëPSP°C... PİCḡPµPİP°CḡP°C, PSPµ  
PIPëCİPIP»CİPI C,PµCḡP°C,PsPİPµPSPSPSc— PrC-C—, PNëPsPİPs P·P°ClC,PsClC†PIP°PSPSCİ  
Cl PrC†P¶Pµ PIPëClPsPePëC... PrPsP·P°C..., C%oPs PİCḡPëP±P»PëP·PSPs Pİ 300 CḡP°P·C-PI  
PİPµCḡPµPIPëC%oClPIP°P»Pë PjP°PeClPëPjP°P»CHPSC† CḡPµPePsPjPµPSPPrPsPIP°PSC†  
PrPsP·Cl PrP»Cİ P»CḡPrPëPSPë, PİCḡPëP·PİPsPrPëP»Ps PrPs P·PSPëP¶PµPSPSCİ  
C,CḡPëPIP°P»PsClC,C- P¶PëC,C,Cİ PİP»PsPrP°.

P°PsPrC†PIP°PSPSCİ PİCḡC†PrPrCḡ.

P°PsClP»C-PrP¶PµPSPSCİ PSP° C%oClCḡP°C..., ClPeC- PsC,CḡPëPjC†PIP°P»Pë  
PsPrPSPsCḡP°P·PsPIC† PİPµCḡPsCḡP°P»CHPSC† PrPsP·Cl CḡP°PrC-PsP°PeC,PëPIPSPsPİPs  
PrPsPeClP°P·PsP·PëPSC† Cl PİPµCḡC-PsPr P»P°PeC,P°C†C-C—, PIPëP·PSP°C‡PëP»Pë, C%oPs  
PrPsPeClP°P·PsP·PëPS PSP°PePsPİPëC‡C†C" C,CHClCİ Cl PjPsP»PsC†C- C%oClCḡC-PI, PİCḡPë  
C†CHPsPjC† PjP°PeClPëPjP°P»CHPSP° PePsPSC†PµPSC,CḡP°C†C-Cl Cl PjPsP»PsC†C-  
PİCḡPëP±P»PëP·PSPs Cl 20 CḡP°P·C-PI PİPµCḡPµPIPëC%oClC" PePsPSC†PµPSC,CḡP°C†C-Cḡ Cl  
PİP»P°P·PjC- PjP°C,PµCḡC-.

PκP°CİPIPSC- PrP°PSC- Cl P»CḡPrPµPNë PrC†P¶Pµ PsP±PjPµP¶PµPSC-, PİCḡPsC,Pµ,  
PİCḡPsPrPµPjPsPSC†C,CḡPsPIP°PSPs, C%oPs PIPëPrC-P»PµPSPSCİ PrPsPeClP°P·PsP·PëPSC† Cl  
PjP°C,PµCḡPëPSC†CHPePµ PjPsP»PsPePs C" PrC†P¶Pµ PSPµP·PSP°C‡PSPëPj (PIC-PrPSPsClPSP°  
PrPsP·P° PrP»Cİ PrPëC,PëPSPë ClC,P°PSPsPIPëC,CH PjPµPSC€Pµ 1 %). P PëP·PëPe PrP»Cİ  
PSPsPİPsPSP°CḡPsPrP¶PµPSPsPİPs P°P±Ps PSPµPjPsPIP»CİC,Pë PSPµ PjPsP¶PSP°

PIPëPeP»CĤC±PëC,Pë, C,PsPjCf PrPsPeCfP°P·PsP·PëPS CfP»C-Pr P·P°CfC,PsCfPsPICfPIP°C,Pë P»PëCëPµ C,PsPrC-, PePsP»Pë P·P° PrCfPjPePsCĤ P»C-PeP°CĤCf PîPsC,PµPSC†C-PMëPSP° PePsCĤPëCfC,CĤ PîPµCĤPµPIP°P¶P°C” PîPsC,PµPSC†C-PMëPSPëPMë CĤPëP·PëPe.

*PµP°C†C-C”PSC,Pë C-P· P”P”PµP—.*

P†PSC,,PsCĤPjP°C†C-Cf Cf PrP°PSPsPjCf CĤPsP·PrC-P»C- PSPµ CfC,PsCfCfC”C,CĤCfCf PîP°C†C-C”PSC,C-PI C-P· P”P”PµP—.

***P—PrP°C,PSC-CfC,CĤ PIPiP»PëPIP°C,Pë PSP° CëPIPëPrPeC-CfC,CĤ CĤPµP°PeC†C-C— PîCĤPë PePµCĤCzPIP°PSPSC- P°PIC,PsC,CĤP°PSCfPîPsCĤC,PsPj P°P±Ps C-PSCëPëPjPë PjPµC...P°PSC-P·PjP°PjPë.***

P—PrP°C,PSC-CfC,CĤ PePµCĤCfPIP°C,Pë P°PIC,PsC,CĤP°PSCfPîPsCĤC,PsPj C,P° PîCĤP°C†CĤPIP°C,Pë P· C-PSCëPëPjPë PjPµC...P°PSC-P·PjP°PjPë PjPsP¶Pµ PîPsPiC-CĤCëPëC,PëCfCf, PsCfPsP±P»PëPIP° PSP° PîPsC±P°C,PeCf P»C-PeCfPIP°PSPSCf.

***PŸPîPsCfC-P± P·P°CfC,PsCfCzPIP°PSPSCf C,P° PrPsP·Pë.***

PµCĤPµPîP°CĤP°C, PŷP°CĤPrCfCĤP°<sup>B®</sup> PjPsP¶PSP° PîCĤPëPMëPjP°C,Pë CfPe CfCĤP°PSC†C-, C,P°Pe C- PîPIPµC±PµCĤC-.

PµCĤPµPîP°CĤP°C, P·P°CfC,PsCfPsPICfCĤC,CĤ PîPµCĤPsCĤP°P»CĤPSPs.

*PĥCĤC,PµCĤC-P°P»CĤPSP° PiC-PîPµCĤC,PµPSP·C-Cf.*

PµCĤPµPîP°CĤP°C, CfP»C-Pr P·P°CfC,PsCfPsPICfPIP°C,Pë 1 CĤP°P· PSP° PrPsP±Cf. PµPsC±P°C,PePsPIP° PrPsP·P° CfC,P°PSPsPIPëC,CĤ 1 PjPi, C%oPsP± P·PIPµCfC,Pë PrPs PjC-PSC-PjCfPjCf CĤPëP·PëPe CĤPsP·PIPëC,PeCf PsCĤC,PsCfC,P°C,PëC±PSPsC— P°CĤC,PµCĤC-P°P»CĤPSPsC— PiC-PîPsC,PµPSP·C-C— C,P°/P°P±Ps CfPëPSPePsPîPµ (PrPëPI. CĤPsP·PrC-P» B«PĥCfPsP±P»PëPIP°CfC,C- P·P°CfC,PsCfCfPIP°PSPSCfB»). PµC-CfP»Cf 1-2 C,PëP¶PSC-PI PîPsC±P°C,PePsPIP°C— C,PµCĤP°PîC-C— PrPsP·Cf PjPsP¶PSP° PîC-PrPIPëC%oPëC,Pë PrPs 2 PjPi, P° PîPsC,C-Pj, CfPeC%oPs PSPµPsP±C...C-PrPSPs, C- PrPs 4 PjPi. Pj P±C-P»CĤCëPsCfC,C- PîP°C†C-C”PSC,C-PI PIC-PrPîPsPIC-PrCĤ PSP° C,PµCĤP°PîC-CĤ CfPîPsCfC,PµCĤC-PiP°C”C,CĤCfCf PîCĤPë P·P°CfC,PsCfCfPIP°PSPSC- PîCĤPµPîP°CĤP°C,Cf Cf PrPsP·C- 4 PjPi P°P±Ps PSPëP¶C±C-PMë. PîPeC%oPs PSPµPsP±C...C-PrPSPs, PrPsP·Cf PîCĤPµPîP°CĤP°C,Cf PjPsP¶PSP° PîC-PrPIPëC%oPëC,Pë PrPs 8 PjPi P°P±Ps PrPs PjP°PeCfPëPjP°P»CĤPSPsC— CĤPµPePsPjPµPSPPrPsPIP°PSPsC— PrPsP·Pë Bĥ” 16 PjPi.

*P”PsP±CĤPsCfPeC-CfPSP° PiC-PîPµCĤPîP»P°P·C-Cf PîPµCĤPµPrPjC-C...CzCĤPsPIP°C— P·P°P»PsP·Pë.*

P PµPePsPjPµPSPPrPsPIP°PSP° PîPsC±P°C,PePsPIP° PrPsP·P° PîCĤPµPîP°CĤP°C,Cf PŷP°CĤPrCfCĤP°<sup>B®</sup> CfC,P°PSPsPIPëC,CĤ 1 PjPi 1 CĤP°P· PSP° PrPsP±Cf, C%oPsP± P·PIPµCfC,Pë PrPs PjC-PSC-PjCfPjCf CĤPëP·PëPe CĤPsP·PIPëC,PeCf PsCĤC,PsCfC,P°C,PëC±PSPsC— P°CĤC,PµCĤC-P°P»CĤPSPsC— PiC-PîPsC,PµPSP·C-C— C,P°/P°P±Ps CfPëPSPePsPîPµ (PrPëPI. CĤPsP·PrC-P» B«PĥCfPsP±P»PëPIP°CfC,C- P·P°CfC,PsCfCfPIP°PSPSCfB»). P—P°P»PµP¶PSPs PIC-Pr C-PSPPrPëPIC-PrCfP°P»CĤPSPëC...

PsCfPsP±P»PëPIPsCfC, PµPNë CfcHjPsPrPëPSP°PjC-PePë PïP°CfC-C"psc, P° C, P°  
CfPëPjPïc, PsPjC-PI P"P"PµP—, PrPsP·Cf PjPsP¶PSP° Pïc-PrPIPëC%PëC, Pë PrPs 2 PjPi, PïPsC, C-Pj  
PrPs 4 PjPi C- PrPs PjP°PeCfPëPjP°P»CHbPSPsC— CjPµPePsPjPµPSPPrPsPIP°PSPsC— PrPsP·Pë Bfj"  
8 PjPi. P PµPePsPjPµPSPPrPsPIP°PSPëPNë C-PSC, PµCjPIP°P» Pïc-PrP±PsCjCf PrPsP·Pë  
CfC, P°PSPsPIPëC, CH 1-2 C, PëP¶PSC-. P—PIPëCfP°PNëPSP° CjPµPePsPjPµPSPPrPsPIP°PSP°  
PrPsP·P° PïcHjPµPïP°CjP°C, Cf CfC, P°PSPsPIPëC, CH 2-4 PjPi PSP° PrPsP±Cf.

*PµP°CfC-C"psc, P°Pj P»C-C, pschbPsPiPs PIC-PeCz CfP»C-Pr P·P°CfC, PsCfPsPICfPIP°C, Pë  
P·PIPëCfP°PNëPSC- PrPsP·Pë PrP»Cf PrPsCjPsCfP»PëC....*

*PµP°CfC-C"psc, P°Pj C-P. PïPsCjCzCëPµPSPSCfPj C,, CzPSPeCfC-C— PSPëCjPsPe CfP»C-Pr  
P·P°CfC, PsCfPsPICfPIP°C, Pë P·PIPëCfP°PNëPSC- PrPsP·Pë PrP»Cf PrPsCjPsCfP»PëC...,  
PsCfPeC-P»CHbPePë C,, P°CjPjP°PePsPeC-PSPµC, PëCfPSC- PïP°CjP°PjPµC, CjPë  
PïcHjPµPïP°CjP°C, Cf PïcHjPë PïPsCjCfCëPµPSPSC- C,, CfPSPeCfC-C— PSPëCjPsPe PSPµ  
P·PjC-PSCfCfC, CHCfCfC.*

*PµP°CjPrCfCjP°<sup>B</sup> PSPµ PIPëPIPsPrPëC, CHCfCfC P· PsCjPiP°PSC-P·PjCf P·P°  
PrPsPïPsPjPsPiPsCf PiPµPjPsPrC-P°P»C-P·Cf.*

*PµP°CfC-C"psc, Pë P· PïPsCjCzCëPµPSPSCfPjPë C,, CzPSPeCfC-C— PïPµCfC-PSPePë. PkP°  
PrP°PSPëPNë CfP°Cf C-PSC,, PsCjPjP°CfC-Cf C%PsPrPs P·P°CfC, PsCfCfPIP°PSPSCf  
PïcHjPµPïP°CjP°C, Cf PïP°CfC-C"psc, P°Pj C-P. PïPsCjCfCëPµPSPSCfPjPë C,, CfPSPeCfC-C—  
PïPµCfC-PSPePë C, P° C%PsPrPs PIPïP»PëPICf PïcHjPµPïP°CjP°C, C-PI, P·PrP°C, PSPëC...  
P·PjC-PSCfPIP°C, Pë PïPµCfC-PSPePsPIPëPNë PjPµC, P°P±PsP»C-P·Pj (PSP°PïcHjPëPeP»P°Pr,  
CfPëPjPµC, PëPrPëPSCf), C" PsP±PjPµP¶PµPSPsCf. PïPe C- PïcHjPë P·P°CfC, PsCfCfPIP°PSPSC-  
C-PSCëPëC... PïcHjPµPïP°CjP°C, C-PI, C%Ps PïPsPIPSC-CfC, Cf  
PjPµC, P°P±PsP»C-P·CfCfC, CHCfCfC PïPµCfC-PSPePsCf, PïP°CfC-C"psc, P°Pj C-P.  
PsP·PSP°PeP°PjPë PïPsCjCfCëPµPSCf C,, CfPSPeCfC-C— PïPµCfC-PSPePë PïcHjPµPïP°CjP°C,  
CfP»C-Pr PïcHjPëP·PSP°CfP°C, Pë P· PsP±PµCjPµP¶PSC-CfC, Cf (PrPëPI. CjPsP·PrC-P»  
B«PfcfPsP±P»PëPIPsCfC, C- P·P°CfC, PsCfCfPIP°PSPSCfB» C, P°  
B«PαP°CjPjP°PePsPeC-PSPµC, PëPeP°B»).*

*P"C-C, Pë.*

*P·PµP·PïPµPeP° C, P° PµC,, PµPeC, PëPIPSC-CfC, CH P·P°CfC, PsCfCfPIP°PSPSCf  
P»C-PeP°CjCfCHbPePsPiPs P·P°CfPsP±Cf PµP°CjPrCfCjP°<sup>B</sup> PrC-C, CfPj PSPµ  
PrPsCfP»C-PrP¶CfPIP°P»PëCfCf.*

***PµPµCjPµPrPsP·CzPIP°PSPSCf.***

*PïPeC%Ps PïPµCjPµPrPsP·CfPIP°PSPSCf PïcHjPµPïP°CjP°C, Cf PïcHjPëP·PIPµP»Ps PrPs  
P°CjC, PµCjC-P°P»CHbPSPsC— Pïc-PïPsC, PµPSP·C-C—, PïP°CfC-C"psc, P° CfP»C-Pr  
C, PµCjPjC-PSPsPIPs PïPsPeP»P°CfC, Pë PSP° CfPïPëPSCf, PsPïcCfCf, PëPICëPë PiPsP»PsPICf  
PIPSPëP· P' PsPeCjPµPjPëC... PIPëPïP°PrPeP°C... PjPsP¶PSP° PIP¶PëC, Pë C-PSCëC-  
CfPëPjPïc, PsPjP°C, PëCfPSC- P·P°C...PsPrPë.*

*PïPeC%Ps CfPëPjPïc, PsPjP°C, PëCfPSPëC... P·P°C...PsPrC-PI PSPµPrPsCfC, P°C, pschbPs, PrP»Cf  
P»C-PeCfPIP°PSPSCf CëPsPeCf PïPµCjCëPsCfPµCjPiPsPIPs CfP»C-Pr P·P°CfC, PsCfCfPIP°C, Pë  
PïP»P°P·PjPsP·P°PjC-PSPSPëPePë. PµC-CfP»Cf CfCHbPsPiPs PïcHjPë*

PSPμPsP±C...C-PrPSPsCfC,C- Cfp»C-Pr P·P°CfC,PsCfCfPIP°C,Pë  
CfCfPrPëPSPsP·PICfP¶CfPIP°P»CHbPSC- PïCfPμPïP°CfP°C,Pë. PŸP»C-Pr  
CfPïPsCfC,PμCfC-PiP°C,Pë P·P° CfC,P°PSPsPj C,,CfPSPeCfC-C— PSPëCfPpsPe C,P° PïCfPë  
PSPμPsP±C...C-PrPSPsCfC,C- P·P°CfC,PsCfPpsPICfPIP°C,Pë PïC-PrC,CfPëPjCfCfCfC-  
P·P°C...PsPrPë.

P“PμPjPsPrC-P°P»C-P· PSPμ PïPsPeP°P·P°PSPëP№, PsCfPeC-P»CHbPePë PrPsPeCfP°P·PsP·PëPS  
P·PSP°CfPSPsCf PjC-CfPpsCf P·PiBf™ C¶P·CfC”C,CHbCfC¶ P· P±C-P»PeP°PjPë PïP»P°P·PjPë  
PeCfPpsPIC-.

**PμPsP±C-CfPSC- CfPμP°PeCfC-C—.**

PfCfC,PμCfC-P°P»CHbPSP° PiC-PiPμCfC,PμPSP·C-C¶.

Pj PeP»C-PSC-CfPSPëC... PrPsCfP»C-PrP¶PμPSPSC¶C... P· CfCfP°CfC,Cf PïP°CfC-C”PSC,C-PI  
P· P°CfC,PμCfC-P°P»CHbPSPsCf PiC-PiPμCfC,PμPSP·C-Cf PSP°P№CfP°CfC,C-C¶Pμ  
PIPëPSPëPeP°P»Pë PïPsP±C-CfPSC- CfPμP°PeCfC-C— PïPsCfC,CfCfP°P»CHbPSPsPiPs C,PëPïCf  
(C¶PeC- Cf CfC-PrPeC-CfPSPëC... PIPëPïP°PrPeP°C... CfCfPïCfPpsPIPsPrP¶CfPIP°P»PëCfC¶  
PIC,CfP°C,PsCf CfPIC-PrPsPjPsCfC,C-), P°P±Ps PSPμCfPïPμCfPëC,,C-CfPSC- PïPsP±C-CfPSC-  
CfPμP°PeCfC-C—.

P”PsP±CfPpsC¶PeC-CfPSP° PiC-PiPμCfPïP»P°P·C-C¶ PiPμCfPμPrPjC-C...CfCfPpsPIPsC—  
P·P°P»PsP·Pë.

P—P° PrP°PSPëPjPë PePsPSC,CfPpsP»CHbPsPIP°PSPëC... PeP»C-PSC-CfPSPëC...  
PrPsCfP»C-PrP¶PμPSCb, Cf PïP°CfC-C”PSC,C-PI C-P· P”P“PμP— CfPïPsCfC,PμCfC-PiP°PICfC¶  
C,P°PePëP№ CfP°PjPëP№ PïCfPpsC,,C-P»CHb PïPsP±C-CfPSPëC... CfPμP°PeCfC-P№, C¶Pe C- Cf  
PïP°CfC-C”PSC,C-PI P· P°CfC,PμCfC-P°P»CHbPSPsCf PiC-PiPμCfC,PμPSP·C-Cf.

P”P»C¶ PsCfC-PSPePë CfP°CfC,PsC,Pë PIPëPSPëPePSPμPSPSC¶ PïPsP±C-CfPSPëC... PrC-P№  
PIPëPePsCfPëCfC,PsPICfC”C,CHbCfC¶ PSP°CfC,CfPïPSP° PeP»P°CfPëC,,C-PeP°CfC-C¶:  
PrCfP¶Pμ CfP°CfC,Ps (≥ 1/10), CfP°CfC,Ps (≥ 1/100 C- < 1/10), PSPμCfP°CfC,Ps (≥ 1/1000 C-  
< 1/100), CfC-PrPePs (≥ 1/10000 C- < 1/1000), PrCfP¶Pμ CfC-PrPePs (< 1/10000), C,P°  
CfP°CfC,PsC,P° PSPμPIC-PrPsPjP° (PSPμPjPsP¶P»PëPIPs PsCfC-PSPëC,Pë PSP°  
PïC-PrCfC,P°PIC- C-CfPSCfCfCfPëC... PrP°PSPëC...).

PfPSC,,PμPeCfC-C— C,P° C-PSPIP°P·C-C—.

PfP°CfC,Ps: C-PSC,,PμPeCfC-C— PrPëC...P°P»CHbPSPëC... C¶P»C¶C...C-PI, C-PSC,,PμPeCfC-C—  
CfPμCfPpsPIPëPIC-PrPSPëC... C¶P»C¶C...C-PI.

P— P±PsPeCf CfPëCfC,PμPjPë PeCfPpsPIC- C,P° P»C-PjC,,P°C,PëCfPSPsC— CfPëCfC,PμPjPë.

P”CfP¶Pμ CfC-PrPePs: P»PμP№PePsPïPμPSC-C¶, C,CfPpsPjP±PsCfPëC,PsPïPμPSC-C¶.

P— P±PsPeCf C-PjCfPSPSPsC— CfPëCfC,PμPjPë.

PkPμCfP°CfC,Ps: P°P»PμCfPïC-CfPSC- CfPμP°PeCfC-C—.

PbPμC,P°P±PsP»C-CfPSC- C,P° P°P»C-PjPμPSC,P°CfPSC- CfPpsP·P»P°PrPë.



P— P±PsPeCf C€P»CfPSPePsPIPs-PeP€C€PePsPIPsPiPs C,CfP°PeC,Cf.

PŞP°CfC,Ps: P±C-P»CH Cf P¶P€PIPsC,C-, PrP€CfPipµPifCfC-Cf, CfCfC...C-CfC,CH Cf CfPsC,C-, PSCfPrPsC,P°.

PkPµC±P°CfC,Ps: P·P°PipScf, PjPµC,PµPsCfP€P·Pj, P±P»CfPIP°PSPSCf, Pip°CfC,CfPsPµPSC,PµCfP€C,, PrC-P°CfPµCf.

P“PµPip°C,PsP±C-P»C-P°CfPSC- CfPsP·P»P°PrP€.

PkPµC±P°CfC,Ps: PIC-PrC...P€P»PµPSPSCf PIC-Pr PSPScfPjP€ PipSpP°P·PSP€PeC-PI C,,CfPSPeC†C-C— PipµC±C-PSPeP€.

P“CfP¶Pµ CfC-PrPePs: C...PsP»PµCfC,P°P·, PiPµPip°C,P€C,, P¶PsPIC,CfPSP€C†Cf.

P— P±PsPeCf C€PeC-CfP€ C,P° Pic-PrC€PeC-CfPSPSc— C,PeP°PSP€PSP€.

PŞP°CfC,Ps: CfPIPµCfP±C-P¶.

PkPµC±P°CfC,Ps: PIP€CfP€Pip°PSPSCf PSP° C€PeC-CfC-.

P“CfP¶Pµ CfC-PrPePs: PeCfPsPip€Pibf™ CfPSPeP°, P°P»PsPipµC†C-Cf, PicfCfPifCfCfP°.

P— P±PsPeCf PsPipScfPSPSc-CfCfC...PsPIPsPiPs P°Pip°CfP°C,Cf C,P° CfPipSp»CfC±PSPSc— C,PeP°PSP€PSP€.

PŞP°CfC,Ps: P±C-P»CH Cf CfPip€PSC-, PjC-P°P»Pic-Cf.

PkPµC±P°CfC,Ps: P°CfC,CfP°P»Pic-Cf.

P C-PrPePs: Pjbf™ CfP·PsPIC- CfPip°P·PjP€, Pjbf™ CfP·PsPIP° CfP»P°P±PeC-CfC,CH.

P— P±PsPeCf PSP€CfPsPe C,P° CfPµC±PsPIP€PIC-PrPSPsPiPs C,CfP°PeC,Cf.

PŞP°CfC,Ps: C†P€CfC,P€C,, PSPµC,CfP€PjP°PSPSCf CfPµC±C-.

PkPµC±P°CfC,Ps: PrP€P·CfCfC-Cf, C±P°CfC,Pµ CfPµC±PsPIP€PifCfCfPeP°PSPSCf, PiPµPjP°C,CfCfC-Cf.

P C-PrPePs: PipSp»C-CfCfC-Cf.

P“CfP¶Pµ CfC-PrPePs: Pic-PrPIP€C%PµPSP€PN€ PrC-CfCfPµP·, CfPsP·P»P°PrP€ CfPµC±PsPIP€PifCfCfPeP°PSPSCf, PSC-PeC,CfCfC-Cf.

P— P±PsPeCf CfPµPifCfPsPrCfPeC,P€PIPSPSc— CfP€CfC,PµPjP€ C,P° PjPsP»PsC±PSP€C... P·P°P»PsP·.

PkPµC±P°CfC,Ps: C-PjPipSc,PµPSC†C-Cf.

P“CfP¶Pµ CfC-PrPePs: Pic-PSPµPePsPjP°CfC,C-Cf, PifCfC-P°Pic-P·Pj.

PŞP°CfC,PsC,P° PSPµPIC-PrPsPjP°: CfPµC,CfPsPifCfP°PrPSP° PµCfP€CfP»CfC†C-Cf.

P— P°Pip°P»CHPSC- CfPsP·P»P°PrP€ C,P° CfPµP°PeC†C-C— Cf PjC-CfC†C- PIPIPµPrPµPSPSCf.

PŞP°CÍC,Ps: P°CÍC,PµPSC-Cİ, P±C-P»CH CÍ PiCĤCÍPrPSC-PNē PeP»C-C,C†C-,  
PiCĤPëPiPsPiPsPrC-P±PSC- CÍPëPjPiC,PsPjPë, PiPµCĤPëC,,PµCĤPëC†PSC- PSP°P±CĤCİPePë.

PkPµC†P°CÍC,Ps: P±C-P»CH CÍ C,C-P»C-, PSP°P±CĤCİPe PsP±P»PëC†Cİ.

P”CÍP¶Pµ CĤC-PrPePs: Pïc-PrPIPëC%oPµPSP° PIC,PsPjP»CĤPIP°PSC-CÍC,CH,  
P·P°PiP°P»CHPSPµ PSPµP·PrCÍP¶P°PSPSCİ.

P PµP·CÍP»CHC,P°C,Pë PrPsCÍP»C-PrP¶PµPSCĤ.

PkPµC†P°CÍC,Ps: P·P±C-P»CHC€PµPSPSCİ PjP°CÍPë C,C-P»P°.

P—PIC-C,C¿PIP°PSPSCİ PiCĤPs Pïc-PrPsP·CĤCĤPIP°PSC- PiPsP±C-C†PSC- CĤPµP°PeC†C-C—.

PµC-CÍP»Cİ CĤPµC”CÍC,CĤP°C†C-C— P»C-PeP°CĤCÍCHPePsPiPs P·P°CÍPsP±CÍ  
PIP°P¶P»PëPIPs PiPsPIC-PrPsPjP»CİC,Pë PiCĤPs Pïc-PrPsP·CĤCĤPIP°PSC- PiPsP±C-C†PSC-  
CĤPµP°PeC†C-C—. PjPµ PrP°C” P·PjPsPiCÍ P·PrC-PNēCÍPSCĤPIP°C,Pë  
P±PµP·PiPµCĤPµCĤPIPSPëPNē PjPsPSC-C,PsCĤPëPSPi CÍPïc-PIPIC-PrPSPsC€PµPSPSCİ  
PePsCĤPëCÍC,C- C,P° CĤPëP·PëPeC-PI, PiPsPiBĤ™ CİP·P°PSPëC... C-P·  
P·P°CÍC,PsCÍCÍPIP°PSPSCİPj P»C-PeP°CĤCÍCHPePsPiPs P·P°CÍPsP±CÍ. PĤPµPrPëC†PSPëPj  
PiCĤP°C†C-PIPSPëPeP°Pj PSPµPsP±C...C-PrPSPs PiPsPIC-PrPsPjP»CİC,Pë PiCĤPs P±CÍPrCH-  
CİPeC- PiPsP±C-C†PSC- CĤPµP°PeC†C-C— P·P° PrPsPiPsPjPsPiPsCĤ  
PSP°C†C-PsPSP°P»CHPSPsC— CÍPëCÍC,PµPjPë PiPµCĤPµPrP°C†C- C-PSC,,PsCĤPjP°C†C-C—.

**PjPµCĤPjC-PS PïcĤPëPrP°C,PSPsCÍC,C-.** 4 CĤPsPePë.

**PjPjPsPIPë P·P±PµCĤC-PiP°PSPSCİ.** P—P±PµCĤC-PiP°C,Pë CÍ PSPµPrPsCÍC,CÍPiPSPsPjCÍ  
PrP»Cİ PrC-C,PµPNē PjC-CÍC†C- PïcĤPë C,PµPjPiPµCĤP°C,CÍCĤC- PSPµ PIPëC%oPµ 30 °C.

**PjPiP°PePsPIPëP°.** PµPs 10 C,P°P±P»PµC,PsPe CÍ P±P»C-CÍC,PµCĤC-, PiPs 3  
P±P»C-CÍC,PµCĤPë CÍ PeP°CĤC,PsPSPSC-PNē CÍPiP°PePsPIC†C-.

**PĤP°C,PµPiPsCĤC-Cİ PIC-PrPïcÍCÍPeCÍ.** P—P° CĤPµC†PµPïc,PsPj.

**P’PëCĤPsP±PSPëPe.** PµC,,P°PNēP·PµCĤ PĤPµPSCĤC,,PµPeC†CÍCĤPëPSPi P”PsPNēC†P»PµPSPr  
P”PjP±Pİ/

Pfizer Manufacturing Deutschland GmbH.

**PĤC-CÍC†PµP·PSP°C...PsPrP¶PµPSPSCİ PIPëCĤPsP±PSPëPeP° C,P° P°PrCĤPµCÍP°  
PjC-CÍC†Cİ PïcĤPsPIP°PrP¶PµPSPSCİ PNēPsPiPs PrC-CİP»CHPSPsCÍC,C-.**

РѢСІСІРІР°Р»СѢРГР°Р»СѢ 1, 79108 РѢСѢР°Р№Р±СІСѢРі Р†Рј Р'СѢР°Р№СІРіР°СІ,  
РќС-РјРµС‡С‡РёРSP°/

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