

Р†РќРŸРŸР PJPЛЬP|P†Pİ

**PrP»Cİ PjPµPrPëC†PSPsPiPs P•P°CÍC,PsCÍCíPİP°PSPSCİ P»C-PeP°CĤCÍCĤPePsPiPs
P•P°CÍPsP±Cí**

Р’Р†Р’РĥP™ P Pĥ

РŸРeP»P°Pr:

PrC-CĤC†P° CĤPµC†PsPIPëPSP°: CÍPëP»PrPµPSP°C,,C-P»Cí C†PëC,CĤP°C,,;

1 C,P°P±P»PµC,PeP° P¶CíPİP°P»CĤPSP° PjC-CÍC,PëC,CĤ CÍPëP»PrPµPSP°C,,C-P»Cí 50 PjPi Cí C,,PsCĤPjC- CÍPëP»PrPµPSP°C,,C-P»Cí C†PëC,CĤP°C,Cí 70,24 PjPi P°P±Ps

1 C,P°P±P»PµC,PeP° P¶CíPİP°P»CĤPSP° PjC-CÍC,PëC,CĤ CÍPëP»PrPµPSP°C,,C-P»Cí 100 PjPi Cí C,,PsCĤPjC- CÍPëP»PrPµPSP°C,,C-P»Cí C†PëC,CĤP°C,Cí 140,48 PjPi;

PrPsPiPsPjC-P¶PSC- CĤPµC†PsPIPëPSPë: PeP°P»C-CĤ PİPsP»P°PeCĤPëP»C-PS; PjP°PiPSC-CĤ CÍC,PµP°CĤP°C,; PeCĤPµPjPSC-CĤ PrC-PsPeCÍPëPr PePsP»PsC—PrPSPëP№ P±PµP•PIPsPrPSPëP№; P°CÍPİP°CĤC,P°Pj (P• 951); PSP°C,CĤC-CĤ PeCĤPsCÍPeP°CĤPjPµP»PsP•P°; P°CĤPsPjP°C,PëP•P°C,PsCĤ PjBĤ™ CİC,P° PİPµCĤC†PµPIP°; P»P°PeC,PsP•P°, PjPsPSPsPiC-PrCĤP°C,; PİPsPIC-PrPsPS Pль-30.

P»C-PeP°CĤCÍCĤPeP° C,,PsCĤPjP°. PŸP°P±P»PµC,PePë P¶CíPİP°P»CĤPSC-.

PĥCÍPSPsPIPSC- C,,C-P•PëPePs-C...C-PjC-C†PSC- PİP»P°CÍC,PëPIPsCÍC,C-: C,CĤPëPeCíC,PSC- PrPIPsPsPiCíPeP»C- C,P°P±P»PµC,PePë P±C-P»PsPiPs PePsP»CĤPsCĤCí P• C,PëCÍPSPµPSPSCİPj B«50B» P°P±Ps B«100B» P• PsPrPSPsPiPs P±PsPeCí.

РќР°Р°CĤPjP°PePsC,РµCĤP°PİPµPIC,PëC†PSP° PiCĤCíPİP°. P—P°CÍPsP±Pë, C%Ps P•P°CÍC,PsCÍPsPICíCĤC,CĤCÍCİ PİCĤPë PµCĤPµPeC,PëP»CĤPSC-P№ PrPëCÍC,,CíPSPeC†C-C—. PŸPëP»PrPµPSP°C,,C-P». PльPsPr PĥPŸPİ G04B E03.

РќР°Р°CĤPjP°PePsP»PsPiC-C†PSC- PİP»P°CÍC,PëPIPsCÍC,C-.

РќР°Р°CĤPjP°PePsPrPëPSP°PjC-PeP°.

РŸPëP»PrPµPSP°C,,C-P» BĤ“ PİCĤPµPİP°CĤP°C, PrP»Cİ PİPµCĤPsCĤP°P»CĤPSPsPiPs P•P°CÍC,PsCÍCíPİP°PSPSCİ, PİCĤPëP•PSP°C†PµPSPëP№ PrP»Cİ P»C-PeCíPİP°PSPSCİ PµCĤPµPeC,PëP»CĤPSPsC— PrPëCÍC,,CíPSPeC†C-C—. PµCĤPë CÍPµPeCÍCíP°P»CĤPSPsPjCí P•P±CíPrP¶PµPSPSC- PİCĤPµPİP°CĤP°C, PIC-PrPSPsPIP»CĤC” P•PSPëP¶PµPSCí PµCĤPµPeC,PëP»CĤPSCí C,,CíPSPeC†C-CĤ CëP»CİC...PsPj PİPsCÍPëP»PµPSPSCİ PİCĤPëC,PsPeCí PeCĤPsPIC- PrPs CÍC,P°C,PµPIPsPiPs C†P»PµPSP°.

PαC-P·C-PsP»PsPiC-C‡PSPëPNë PjPμC...P°PSC-P·Pj, C%oPs P·CfPjPsPIP»CfC” PμCfPμPeC‡C-Cf, PIPeP»CfC‡P°C” PIPëPIC-P»CHbPSPμPSPSCf PsPeCfPëPrCf P°P·PsC,Cf (NO) Cf PeP°PIPμCfPSPsP·PSPëC... C,C-P»P°C... Pïc-Pr C‡P°Cf CfPμPeCfCfP°P»CHbPSPsPiPs P·P±CfPrPμPμPSPSCf. P’PëPIC-P»CHbPSPμPSPëPNë NO P°PeC,PëPICfC” C,,PμCfPjPμPSC, PiCfP°PSC-P»P°C,C‡PëPeP»P°P·Cf, C%oPs CfC,PëPjCfP»CfC” Pïc-PrPIPëC%oPμPSPSCf CfC-PIPSCf C‡PëPeP»C-C‡PSPsPiPs PiCfP°PSPsP·PëPSPjPsPSPsC,,PsCfC,,P°C,Cf (C‡P”PbPα), PIPëPeP»PëPeP°C” CfPsP·CfP»P°P±P»PμPSPSCf PiP»P°PrPePsC— PjCfCfPeCfP»P°C,CfCfPë PeP°PIPμCfPSPsP·PSPëC... C,C-P» C,P° CfPïcCfPëCfC” PïcCfPëC,PsPeCf PeCfPμPIC-.

PÿPëP»PrPμPSP°C,,C-P» C” PïPsC,CfPμPSPëPj C,P° CfPμP»PμPeC,PëPIPSPëPj C-PSPIC-P±C-C,PsCfPμPj C‡P”PbPα-CfPïPμC‡PëC,,C-C‡PSPsC— C,,PsCfC,,PsPrC-PμCfC,PμCfP°P·PëB B B B B B B B 5 (PαP”P·5) Cf PeP°PIPμCfPSPsP·PSPëC... C,C-P»P°C..., PrPμ PαP”P·5 PIC-PrPïPsPIC-PrP°C” P·P° CfPsP·PïP°Pr C‡P”PbPα. P·C,,PμPeC,Pë CfPëP»PrPμPSP°C,,C-P»Cf PSP° PμCfPμPeC‡C-Cf PjP°CfC,CH PïPμCfPëC,,PμCfPëC‡PSPëPNë C...P°CfP°PeC,PμCf. PÿPëP»PrPμPSP°C,,C-P» PSPμ C‡PëPSPëC,CH P±PμP·PïPsCfPμCfPμPrPSCbPsC— CfPμP»P°PeCfP°C‡C-PNëPSPsC— PrC-C— PSP° C-P·PsP»CHbPsPIP°PSC- PeP°PIPμCfPSPsP·PSC- C,C-P»P° P»CfPrPëPSPë, P°P»Pμ PïPsC,CfPμPSPs PïPsCfPëP»CfC” CfPsP·CfP»P°P±P»CfPμP°P»CHbPSCf PrC-Cf NO PSP° C‡Cf C,PeP°PSPëPSCf. PμCfPë P°PeC,PëPIP°C‡C-C— PjPμC,P°P±PsP»C-C‡PSPsPiPs CëP»CfC...Cf NO/C‡P”PbPα, C%oPs PIC-PrP±CfPIP°C”C,CHCfCf PïcCfPë CfPμPeCfCfP°P»CHbPSC-PNë CfC,PëPjCfP»CfC‡C-C—, C-PSPIC-P±CfPIP°PSPSCf CfPëP»PrPμPSP°C,,C-P»PsPj PαP”P·5 CfPïcCfPëC‡PëPSCfC” Pïc-PrPIPëC%oPμPSPSCf CfC-PIPSCf C‡P”PbPα Cf PeP°PIPμCfPSPsP·PSPëC... C,C-P»P°C.... PÿP°PePëPj C‡PëPSPsPj, PrP»Cf C,PsPiPs, C%oPsP± Pïc-CfP»Cf P·P°CfC,PsCfCfPIP°PSPSCf CfPëP»PrPμPSP°C,,C-P»Cf PIPëPSPëP PSPμPsP±C...C-PrPSPëPNë C,,P°CfPjP°PePsP»PsPiC-C‡PSPëPNë PμC,,PμPeC,, PSPμPsP±C...C-PrPSPμ CfPμPeCfCfP°P»CHbPSPμ P·P±CfPrPμPμPSPSCf.

P”PsCfP»C-PrPμPμPSPSCf *in vitro* PïcCfPsPrPμPjPsPSCfC,CfCfPIP°P»Pë, C%oPs CfPëP»PrPμPSP°C,,C-P» C” CfPμP»PμPeC,PëPIPSPëPj PrPs PαP”P·5, C%oPs P±PμCfPμ P°PeC,PëPIPSCf CfC‡P°CfC,CH Cf PïcCfPsC‡PμCfC- PμCfPμPeC‡C-C—. P·C,,PμPeC, CfPëP»PrPμPSP°C,,C-P»Cf PSP° PαP”P·5 PïPsC,CfPμPSC-CëPëPNë, PSC-PμPSP° C-PSCëC- PIC-PrPsPjC- C,,PsCfC,,PsPrC-PμCfC,PμCfP°P·Pë. PjPμPNë PμC,,PμPeC, Cf 10 CfP°P·C-PI PïPsC,CfPμPSC-CëPëPNë, PSC-PμPμC,,PμPeC, PSP° PαP”P·6, C%oPs P±PμCfPμ CfC‡P°CfC,CH Cf PïcCfPsC‡PμCfP°C... C,,PsC,PsPïPμCfPμC,PIPsCfPμPSPSCf Cf CfC-C,PeC-PIC‡C-. PμCfPë P·P°CfC,PsCfCfPIP°PSPSC- PjP°PeCfPëPjP°P»CHbPSPëC... CfPμPePsPjPμPSPPrPsPIP°PSPëC... PrPsP· CfPμP»PμPeC,PëPIPSC-CfC,CH CfPëP»PrPμPSP°C,,C-P»Cf PrPs PαP”P·5 Cf 80 CfP°P·C-PI PïPμCfPμPIPëC%oCfC” PNëPsPiPs CfPμP»PμPeC,PëPIPSC-CfC,CH PrPs PαP”P·1, Cf 700 CfP°P·C-PI PIPëC%oP°, PSC-PμP 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—PsPeCfPμPjP°, CfPμP»PμPeC,PëPIPSC-CfC,CH CfPëP»PrPμPSP°C,,C-P»Cf PrPs PαP”P·5 Cf 4000 CfP°P·C-PI PïPμCfPμPIPëC%oCfC” PNëPsPiPs CfPμP»PμPeC,PëPIPSC-CfC,CH PrPs PαP”P·3 b” C‡PbPbPα-CfPïPμC‡PëC,,C-C‡PSPsC— C-P·PsC,,PsCfPjPë C,,PsCfC,,PsPrC-PμCfC,PμCfP°P·Pë, C%oPs P±PμCfPμ CfC‡P°CfC,CH Cf CfPμPïcCfP»CfC‡C-C— CfPμCfC‡PμPIPSC— CfPePsCfPμC,P»PëPIPSCfC,C-.

PαP°CfPjP°PePsPeC-PSPμC,PëPeP°.

PbP±CfPsCfP±C‡C-Cf. PÿPëP»PrPμPSP°C,,C-P» CëPIPëPrPePs P°P±CfPsCfP±CfC”C,CHCfCf. PbP°PeCfPëPjP°P»CHbPSC- PïP»P°P·PjPsPIC- PePsPSC‡PμPSC,CfP°C‡C-C— PïcCfPμPïP°CfP°C,Cf PrPsCfCfPïP°CfC,CHCfCf PïcCfPsC,CfPïPsPj 30b”120 C...PIPëP»PëPS (C-P· PjPμPrC-P°PSPsCf 60 C...PIPëP»PëPS) Pïc-CfP»Cf PNëPsPiPs PïPμCfPsCfP°P»CHbPSPsPiPs P·P°CfC,PsCfCfPIP°PSPSCf PSP°C,C%oPμ. PÿPμCfPμPrPSCf P°P±CfPsP»CfC,PSP° P±C-PsPrPsCfC,CfPïPSC-CfC,CH Pïc-CfP»Cf PïPμCfPsCfP°P»CHbPSPsPiPs

P·P°CÍC,PsCÍCfPIP°PSPSCÍ CÍC,P°PSPsPIPëC,CH 41 % (C-P· PrC-P°PiP°P·PsPSPsPj P·PSP°C±PµPSCĤ PIC-Pr 25 PrPs 63 %). PJ CĤPµPePsPjPµPSPPrPsPIP°PSPsPjCÍ PrC-P°PiP°P·PsPSC- PrPsP· (PIC-Pr 25 PrPs 100 PjPi) PiP»PsC%°° PiC-Pr C,,P°CĤPjP°PePsPeC-PSPµC,PëC±PSPsCĤ PeCĤPëPIPsCĤ B«PePsPSC±PµPSC,CĤP°C†C-CÍ- C±P°CÍB» (AUC) C,P° PjP°PeCÍPëPjP°P»CHPSP° PePsPSC±PµPSC,CĤP°C†C-CÍ (PŸ_{max}) CÍPëP»PrPµPSP°C,,C-P»CÍ PiC-CÍP»CÍ PNePsPiPs PiPµCĤPsCĤP°P»CHPSPsPiPs P·P°CÍC,PsCÍCfPIP°PSPSCÍ PiC-PrPIPëC%°CfCĤC,CHCÍCÍ PiCĤPsPiPsCĤC†C-PNePSPs PrPs PrPsP·Pë.

PµCĤPë P·P°CÍC,PsCÍCfPIP°PSPSC- CÍPëP»PrPµPSP°C,,C-P»CÍ PiC-Pr C±P°CÍ PIP°PëPIP°PSPSCÍ C—P¶C- CÍC,CfPiC-PSCH P°P±CÍPsCĤP±C†C-C— P·PSPëP¶CfC”C,CHCÍCÍ P·C- CÍPµCĤPµPrPSC-Pj PiPsPrPsPIP¶PµPSPSCÍPj C±P°CÍCÍ PrPsCÍCÍPiPSPµPSPSCÍ PjP°PeCÍPëPjP°P»CHPSPsC— PePsPSC±PµPSC,CĤP°C†C-C— (T_{max}) PrPs 60 C...PIPëP»PëPS C- CÍPµCĤPµPrPSC-Pj P·PSPëP¶PµPSPSCÍPj C_{max} PSP° 29 %.

P PsP·PiPsPrC-P». PŸPµCĤPµPrPSC-PNe CĤC-PIPSPsPIP°P¶PSPëPNe PsP±BĤ™ C”Pj CĤPsP·PiPsPrC-P»CÍ (V_d) CÍC,P°PSPsPIPëC,CH 105 P»C-C,CĤC-PI, C%°Ps CÍPIC-PrC±PëC,CH PiCĤPs CĤPsP·PiPsPrC-P» PiCĤPµPiP°CĤP°C,CÍ PI C,PeP°PSPëPSP°C... PsCĤPiP°PSC-P·PjCÍ. PµC-CÍP»CÍ PsPrPSPsCĤP°P·PsPiPsPiPs PiPµCĤPsCĤP°P»CHPSPsPiPs P·P°CÍC,PsCÍCfPIP°PSPSCÍ CÍPëP»PrPµPSP°C,,C-P»CÍ PI PrPsP·C- 100 PjPi CÍPµCĤPµPrPSCÍ PjP°PeCÍPëPjP°P»CHPSP° P·P°PiP°P»CHPSP° PiP»P°P·PjPsPIP° PePsPSC±PµPSC,CĤP°C†C-CÍ CÍPëP»PrPµPSP°C,,C-P»CÍ CÍC,P°PSPsPIPëC,CH PiCĤPëP±P»PëP·PSPs 440 PSPi/PjP» (PePsPµC,,C-C†C-C”PSC, PIP°CĤC-P°C†C-C— CÍC,P°PSPsPIPëC,CH 40 %). PhCÍPeC-P»CHPePë P·PiBĤ™ CÍP·CfPIP°PSPSCÍ CÍPëP»PrPµPSP°C,,C-P»CÍ C,P° PNePsPiPs PiPsP»PsPiPSPsPiPs N- PrPµCÍPjPµC,PëP»PjPµC,P°P±PsP»C-C,CÍ P· P±C-P»PeP°PjPë PiP»P°P·PjPë PeCĤPsPIC- PrPsCÍCÍPiP°C” 96 %, CÍPµCĤPµPrPSCÍ PjP°PeCÍPëPjP°P»CHPSP° PiP»P°P·PjPsPIP° PePsPSC±PµPSC,CĤP°C†C-CÍ PIC-P»CHPSPsPiPs CÍPëP»PrPµPSP°C,,C-P»CÍ PrPsCÍCÍPiP°C” 18 PSPi/PjP» (38 PSPjPsP»CH). PŸC,CfPiC-PSCH P·PiBĤ™ CÍP·CfPIP°PSPSCÍ P· P±C-P»PeP°PjPë PiP»P°P·PjPë PeCĤPsPIC- PSPµ P·P°P»PµP¶PëC,CH PIC-Pr P·P°PiP°P»CHPSPëC... PePsPSC±PµPSC,CĤP°C†C-PNe CÍPëP»PrPµPSP°C,,C-P»CÍ.

PJ P·PrPsCĤPsPIPëC... PrPsP±CĤPsPiPsP»CHC†C-PI, CÍPeC- P·P°CÍC,PsCÍPsPICfPIP°P»Pë CÍPëP»PrPµPSP°C,,C-P» PsPrPSPsCĤP°P·PsPiPs PI PrPsP·C- 100 PjPi, C±PµCĤPµP· 90 C...PIPëP»PëPS PI PµCÍPeCfP»CÍC,C- PIPëP·PSP°C±P°P»PsCÍCÍ PjPµPSC€Pµ 0,0002 % (CÍ CÍPµCĤPµPrPSCĤPsPjCÍ 188 PSPi) P·P°CÍC,PsCÍPsPIP°PSPsC— PrPsP·Pë.

P’C-PsC,CĤP°PSCÍC,,PsCĤPjP°C†C-CÍ. PµPµC,P°P±PsP»C-P·Pj CÍPëP»PrPµPSP°C,,C-P»CÍ PIC-PrP±CfPIP°C”C,CHCÍCÍ PiPsP»PsPiPSPëPj C±PëPSPsPj P·P° CÍC±P°CÍC,CH PjC-PeCĤPsCÍPsPjP°P»CHPSPëC... C-P·PsC,,PµCĤPjPµPSC,C-PI PiPµC±C-PSPePë CYP3A4 (PiPsP»PsPiPSPëPNe C€P»CÍC...) C,P° CYP2C9 (PrCĤCfPiPsCĤCÍPrPSPëPNe C€P»CÍC...). P”PsP»PsPiPSPëPNe C†PëCĤPeCfP»CĤCĤC±PëPNe PjPµC,P°P±PsP»C-C, CÍC,PIPsCĤCĤC”C,CHCÍCÍ C€P»CÍC...PsPj N-PrPµPjPµC,PëP»CHPIP°PSPSCÍ CÍPëP»PrPµPSP°C,,C-P»CÍ. PŸPµP»PµPeC,PëPIPSC-CÍC,CH PjPµC,P°P±PsP»C-C,CÍ C%°PsPrPs PαP”P·5 PiPsCĤC-PIPSCÍPSPSP° P·C- CÍPµP»PµPeC,PëPIPSC-CÍC,CĤ CÍPëP»PrPµPSP°C,,C-P»CÍ, P° P°PeC,PëPIPSC-CÍC,CH PjPµC,P°P±PsP»C-C,CÍ C%°PsPrPs PαP”P·5 CÍC,P°PSPsPIPëC,CH PiCĤPëP±P»PëP·PSPs 50 % P°PeC,PëPIPSPsCÍC,C- PIPëC...C-PrPSPsC— CĤPµC±PsPiPëPSPë. PµP»P°P·PjPsPIC- PePsPSC±PµPSC,CĤP°C†C-C— C†CHPsPiPs PjPµC,P°P±PsP»C-C,CÍ CÍC,P°PSPsPIP»CÍC,CH PiCĤPëP±P»PëP·PSPs 40 % PePsPSC±PµPSC,CĤP°C†C-C— CÍPëP»PrPµPSP°C,,C-P»CÍ CÍ PiP»P°P·PjC- PeCĤPsPIC-. N- PrPµPjPµC,PëP»CHPsPIP°PSPëPNe PjPµC,P°P±PsP»C-C, P·P°P·PSP°C” PiPsPrP°P»CHC€PsPiPs PjPµC,P°P±PsP»C-P·PjCÍ, P° PiPµCĤC-PsPr PNePsPiPs PSP°PiC-PIPIPëPIPµPrPµPSPSCÍ

СГС,Р°PSPsPIPëC,СН ПİСГРРëP±P»PëP·PSPs 4 PiPsPrPëPSPë.

Р'PëPIPµPrPµPSPSCII. P—P°PiP°P»СНPSPëP№ PeP»C-СГРµPSCГГ ГГРëP»PrPµPSP°C,,C-P»Cр
СГС,Р°PSPsPIPëC,СН 41 P»/PiPsPr, P·CрPjPsPIP»СГСГС±Pë PµCГС-PsPr P№PsPiPs
PSP°PİC-PIPIPëPIPµPrPµPSPSCII C,СГРëPIP°P»C-СГС,СГ 3вГ"5 PiPsPrPëPS. ПİPë PİC-СГР»CII
PµCГСPsСГР°P»СНPSPsPiPs, C,Р°Pë C- PİC-СГР»CII PIPSCрC,СГС-CëPSCНPsPIPµPSPSPsPiPs
P·P°СГС,PsСГСрPIP°PSPSCII PµPëCГPëCГРµCГC-СII ГГРëP»PrPµPSP°C,,C-P»Cр Cр
PIPëPiP»CIIPrC- PjPµC,Р°P±PsP»C-C,C-PI P·PrC-P№CГPSCГC"С,СНCГCII PiPsP»PsPIPSPëPj
C±PëPSPsPj C-P· PëP°P»PsPj (PİСГРРëP±P»PëP·PSPs 80 % PIPµPrPµPSPsC—
PµCГСPsСГР°P»СНPSPsC— PrPsP·Pë) C,Р° PjPµPSCëPsСГ PjC-СГPsСГ P·C- СГPµC±PµCГ
(PİСГРРëP±P»PëP·PSPs 13 % PIPµPrPµPSPsC— PµCГСPsСГР°P»СНPSPsC— PrPsP·Pë).

PαP°СГРPjP°PëPsPëC-PSPµC,PëPëP° Cз PsCГPsP±P»PëPIPëC... PiСГCзPi PİP°CГC-C"PSC,C-PI.

РµP°CГC-C"PSC,Pë P»C-C,PSCHPsPiPs PIC-PëCр. PJ P·PrPsСГPsPIPëC...
PrPsP±СГPsPIPpsP»СНCГC-PI P»C-C,PSCHPsPiPs PIC-PëCр (PIC-PëPsPj PIC-Pr 65 СГPsPëC-PI)
PIC-PrP·PSP°C±P°P»PsCГCII P·PSPëPµPµPSPSCII PeP»C-СГРµPSCГГCр
CГРëP»PrPµPSP°C,,C-P»Cр, C°Ps P·CрPjPsPIP»СГPIP°P»Ps PİC-PrPIPëC°PµPSPSCII
PİP»P°P·PjPsPIPëC... PëPsPSCГPµPSC,СГP°CГC-P№ CГPëP»PrPµPSP°C,,C-P»Cр C,Р° P№PsPiPs
P°PëC,PëPIPSPsPiPs N-PrPµPjPµC,PëP»СНPsPIP°PSPsPiPs PjPµC,Р°P±PsP»C-C,Cр
PİСГРРëP±P»PëP·PSPs PSP° 90 % PİPsСГC-PIPSCII PSPs P· PIC-PrPİPsPIC-PrPSPëPjPë
PëPsPSCГPµPSC,СГP°CГC-СII PjPë Cр P·PrPsСГPsPIPëC... PrPsP±СГPsPIPpsP»СНCГC-PI
PjPsP»PsPrCëPsPiPs PIC-PëCр (18вГ"45 СГPsPëC-PI). PJ P·PİвГ™ CII P·PëCр P· PIC-PëPsPIPëPjPë
PIC-PrPjC-PSPSPsCГC,СII PjPë Cр P·PİвГ™ CII P·CрPIP°PSPSC- P· P±C-P»PëP°PjPë PİP»P°P·PjPë
PëCГPsPIC- PIC-PrPİPsPIC-PrPSPµ PİC-PrPIPëC°PµPSPSCII PİP»P°P·PjPsPIPpsC—
PëPsPSCГPµPSC,СГP°CГC-C— PIC-P»СНPSPsPiPs CГРëP»PrPµPSP°C,,C-P»Cр
СГС,Р°PSPsPIPëP»Ps PİСГРРëP±P»PëP·PSPs 40 %.

РкPëCГPëPsPIP° PSPµPrPsCГC,Р°C,PSC-СГC,СН. PJ PrPsP±СГPsPIPpsP»СНCГC-PI C-P·
PİPsСГCрCëPµPSPSCII PjPë C,,CрPSPëCГC-C— PSPëCГPsPë P»PµPiPëPsPiPs C,Р°
PİPsPjC-СГPSPsPiPs CГC,CрPµPSCII (PëP»C-СГРµPSCГГ PëCГPµP°C,PëPSC-PSCр 30вГ"80
PjP»/C...PI) C,,P°СГРPjP°PëPsPëC-PSPµC,PëPëP° CГРëP»PrPµPSP°C,,C-P»Cр
P·P°P»PëCëP°P»P°CГCII PSPµP·PjC-PSPµPSPsСГ PİC-СГР»CII P№PsPiPs
PsPrPSPsСГP°P·PsPIPpsPiPs PµCГСPsСГР°P»СНPSPsPiPs P·P°СГС,PsСГСрPIP°PSPSCII Cр
PrPsP·C- 50 PjPi. PÿPµCГPµPrPSC- AUC C,Р° C_{max} N-PrPµPjPµC,PëP»СНPsPIP°PSPsPiPs
PjPµC,Р°P±PsP»C-C,Cр PİC-PrPIPëC°CрPIP°P»PëCГCII PSP° 126 % C,Р° 73 %
PIC-PrPİPsPIC-PrPSPs PİPsСГC-PIPSCII PSPs P· C,Р°PëPëPjPë PİPsPëP°P·PSPëPëP°PjPë Cр
PrPsP±СГPsPIPpsP»СНCГC-PI C,Р°PëPsPiPs Pµ PIC-PëCр P±PµP· PİPsСГCрCëPµPSCН
C,,CрPSPëCГC-C— PSPëCГPsPë. PhPrPSP°Pë C±PµCГPµP· PIPëCГPsPëCр
C-PSPPrPëPIC-PrCрP°P»СНPSCр PIP°СГC-P°P±PµP»СНPSC-СГC,СН CГC-
PIC-PrPjC-PSPSPsCГC,C- PSPµ P±CрP»Pë CГC,Р°C,PëCГC,PëC±PSPs P·PSP°C±CрC°PëPjPë.

PJ PrPsP±СГPsPIPpsP»СНCГC-PI C-P· C,СII PµPëPjPë PİPsСГCрCëPµPSPSCII PjPë
C,,CрPSPëCГC-C— PSPëCГPsPë (PëP»C-СГРµPSCГГ PëCГPµP°C,PëPSC-PSCр PSPëPµC±Pµ 30
PjP»/C...PI) PëP»C-СГРµPSCГГ CГРëP»PrPµPSP°C,,C-P»Cр P·PSPëPµCрPIP°PICГCII, C°Ps
CГPİСГPëC±PëPSCII P»Ps CГPµCГPµPrPSC" PİC-PrPIPëC°PµPSPSCII AUC C,Р° C_{max} PSP° 100 %
C,Р° 88 % PIC-PrPİPsPIC-PrPSPs PİPsСГC-PIPSCII PSPs C-P· PrPsP±СГPsPIPpsP»СНCГCII PjPë
C,Р°PëPsPiPs Pµ PIC-PëCр P±PµP· PİPsСГCрCëPµPSCН C,,CрPSPëCГC-C— PSPëCГPsPë.
PлСГC-Pj C,PsPiPs, P·PSP°C±PµPSPSCII AUC C,Р° C_{max} N-PrPµPjPµC,PëP»СНPsPIP°PSPsPiPs
PjPµC,Р°P±PsP»C-C,Cр P·PSP°C±CрC°Pµ PİC-PrPIPëC°CрPIP°P»PëCГCII вГ" PSP° 200 % C,Р°
79 % PIC-PrPİPsPIC-PrPSPs.

РμРμС±С-PSPePsPIP° PSPμPrPsCГC, P°С, PSC-ГC, ЧХ. PJ PrPsP±CГPsPIPsP»ЧХC†C-PI C-P·
C†PëCГPsP·PsPj PìPμC±C-PSPePë P»PμPiPePsPiPs C, P° PìPsPjC-CГPSPsPiPs CГC, CрPìPμPSCЦ
(PeP»P°CГPë Pђ C, P° P' P·P° PeP»P°CГPëC,, C-PeP°C†C-C" Cђ P\$P°PNëP»PrP°BГ" PμBГ™ Cђ)
PeP»C-CГPμPSCГ CГPëP»PrPμPSP°C,, C-P»Cр P·PSPëP¶CрPIP°PICГЦ, C%Ps
PìCГPëP·PIPsPrPëP»Ps PrPs PìC-PrPIPëC%PμPSPSCЦ AUC (84 %) C, P° C_{max} (47 %)
PìPsCГC-PIPSCЦPSPs P· PrPsP±CГPsPIPsP»ЧХC†CЦPjPë C, P°PePsPiPs P¶ PIC-PeCр P±PμP·
PìPsCГCрCëPμPSCђ C,, CрPSPeC†C-C— PìPμC±C-PSPePë. PαP°CГPjP°PePsPeC-PSPμC, PëPeP°
CГPëP»PrPμPSP°C,, C-P»Cр Cр PìP°C†C-C" PSC, C-PI C-P· PìPsCГCрCëPμPSPSCЦPjPë
C,, CрPSPeC†C-C— PìPμC±C-PSPePë C, CЦP¶PePsPiPs CГC, CрPìPμPSCЦ PSPμ
PIPëPIC†P°P»P°CГCЦ.

РљР»C-PSC-C±PSC- C...P°CГP°PeC, PμCГPëCГC, PëPePë.

PμPsPeP°P·P°PSPSCЦ.

РμCГPμPìP°CГP°C, CГPμPePsPjPμPSPPrCрC" C, ЧХCГCЦ P·P°CГC, PsCГPsPICрPIP°C, Pë
C±PsP»PsPIC-PeP°Pj C-P· PμCГPμPeC, PëP»ЧХPSPsCђ PrPëCГC,, CрPSPeC†C-C" Cђ, CЦPeP°
PIPëP·PSP°C±P°C" C, ЧХCГCЦ CЦPe PSPμP·PrP°C, PSC-ГC, ЧХ PrPsCГCЦPiC, Pë P°P±Ps
PìC-PrC, CГPëPjP°C, Pë PμCГPμPeC†C-Cђ CГC, P°C, PμPIPsPiPs C±P»PμPSP°,
PSPμPsP±C...C-PrPSCр PrP»CЦ CрCГPìC-CëPSPsPiPs CГC, P°C, PμPIPsPiPs P°PeC, Cр.

P"Р»CЦ PμC,, PμPeC, PëPIPSPsC— PrC-C— PìCГPμPìP°CГP°C, Cр PìPsC, CГC-P±PSPμ
CГPμPeCГCрP°P»ЧХPSPμ P·P±CрPrP¶PμPSPSCЦ.

PμCГPsC, PëPìPsPeP°P·P°PSPSCЦ.

- PμC-PrPIPëC%PμPSP° C±CрC, P»PëPIC-ГC, ЧХ PrPs P°PeC, PëPIPSPsC—
CГPμC±PsPIPëPSPë P°P±Ps PrPs P±CрPrCђ-CЦPePsC— P· PrPsPìPsPjC-P¶PSPëC...
CГPμC±PsPIPëPS PìCГPμPìP°CГP°C, Cр.
- PђPrPSPsC±P°CГPSPμ P·P°CГC, PsCГCрPIP°PSPSCЦ C-P· PrPsPSPsCГP°PjPë PsPeCГPëPrCр
P°P·PsC, Cр (C, P°PePëPjPë CЦPe P°PjC-P»PSC-C, CГPëC,) P°P±Ps PSC-C, CГP°C, P°PjPë Cр
P±CрPrCђ-CЦPeC-PNë C,, PsCГPjC- PìCГPsC, PëPìPsPeP°P·P°PSPμ, PsCГPeC-P»ЧХPePë
PIC-PrPsPjPs, C%Ps CГPëP»PrPμPSP°C,, C-P» PjP°C" PIPìP»PëPI PSP° CëP»CЦC...Pë
PjPμC, P°P±PsP»C-P·PjCр NO/C†P" PђPα C, P° PìPsC, PμPSC†C-CђC" PìC-PìPsC, PμPSP·PëPIPSPëPNë
PμC,, PμPeC, PSC-C, CГP°C, C-PI.
- PђPrPSPsC±P°CГPSPμ P·P°CГC, PsCГCрPIP°PSPSCЦ C-PSPiC-P±C-C, PsCГC-PI PαP" P·5 (Cр
C, PsPjCр C±PëCГP»C- CГPëP»PrPμPSP°C,, C-P»Cр) P·C- CГC, PëPjCрP»CЦC, PsCГP°PjPë
PìCрP°PSC-P»P°C, C†PëPeP»P°P·Pë, C, P°PePëPjPë CЦPe CГC-PsC†PëPìCрP°C,,
PìCГPsC, PëPìPsPeP°P·P°PSPμ, PsCГPeC-P»ЧХPePë PjPsP¶Pμ PìCГPëP·PIPμCГC, Pë PrPs
CГPëPjPìC, PsPjP°C, PëC±PSPsC— PìC-PìPsC, PμPSP·C-C— (PrPëPI. CГPsP·PrC-P»
B«P'P·P°C" PjPsPrC-CЦ P· C-PSCëPëPjPë P»C-PeP°CГCГЧХPePëPjPë P·P°CГPsP±P°PjPë C, P°
C-PSCëC- PIPëPrPë PIP·P°C" PjPsPrC-PNëB»).
- PŸC, P°PSPë, PìCГPë CЦPePëC... PSPμ CГPμPePsPjPμPSPPrPsPIP°PSP°
CГPμPeCГCрP°P»ЧХPSP° P°PeC, PëPIPSC-ГC, ЧХ (PSP°PìCГPëPeP»P°Pr, C, CЦP¶PeC-
CГPμCГC†PμPIPs-CГCрPrPëPSPSC- CГPsP·P»P°PrPë, C, P°PeC- CЦPe PSPμCГC, P°P±C-P»ЧХPSP°
CГC, PμPSPsPeP°CГPrC-CЦ P°P±Ps CГPμCГC†PμPIP° PSPμPrPsCГC, P°C, PSC-ГC, ЧХ

C, CIP¶PePsPiPs CÍC, CíPíPµPSCÍ).

- P°C, CĤP°C, P° P·PsCĤCí PSP° PsPrPSPµ PsPePs PIPSP°CÍP»C-PrPsPe
PSPµP°CĤC, PµCĤC-P°P»CHĤPSPsC— PíPµCĤPµPrPSCĤPsC— C-C€PµPjC-C‡PSPsC—
PSPµPICĤPµPíP°C, C-C— P·PsCĤPµPIPµPiPs PSPµCĤPIP°, 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, PsCÍCíPIP°PSPSCÍPj C-PSPiC-P±C-C, PsCĤC-PI PαP”P•5 C‡Pë PSC-.

- PkP°CÍPIPSC-CÍC, CH C, P°PePëC... P·P°C...PIPScĤCĤPIP°PSCĤ: PíPsCĤCíC€PµPSPSCÍ
C,, CíPSPeC†C-C— PíPµC‡C-PSPePë C, CIP¶PePsPiPs CÍC, CíPíPµPSCÍ,
P°CĤC, PµCĤC-P°P»CHĤPSP° PiC-PíPsC, PµPSP·C-CÍ (P°CĤC, PµCĤC-P°P»CHĤPSPëP№ C, PëCÍPe
PSPëP¶C‡Pµ 90/50 PjPj CĤC,, CÍC,,), PSPµC%oPsPrP°PIPSPs PíPµCĤPµPSPµCÍPµPSPëP№
C-PSCÍCíP»CHĤC, P°P±Ps C-PSC,, P°CĤPeC, PjC-PsPeP°CĤPrP° C, P° CÍPíP°PrPePsPIC-
PrPµPíPµPSPµCĤP°C, PëPIPSC- P·P°C...PIPScĤCĤPIP°PSPSCÍ CÍC-C, PeC-PIPePë, C, P°PeC- CÍPe
PíC-PíPjPµPSC, PSPëP№ CĤPµC, PëPSC-C, (PSPµPIPµP»PëPeP° PeC-P»CHPeC-CÍC, CH
C, P°PePëC... PíP°C†C-C”PSC, C-PI PjP°C” PíPµPSPµC, PëC‡PSC- CĤPsP·P»P°PrPë
C,, PsCÍC,, PsPrC-PµCÍC, PµCĤP°P· CÍC-C, PeC-PIPePë), BĤ” PsCÍPeC-P»CHPePë P±PµP·PíPµPeP°
CÍPëP»PrPµPSP°C,, C-P»Cí PSPµ PrPsCÍP»C-PrP¶CíPIP°P»P°CÍCÍ Cí CĤP°P·C- C, P°PePëC...
P·P°C...PIPScĤCĤPIP°PSCĤ.

**P·P·P°C”PjPsPrC-CÍ P· C-PSC€PëPjPë P»C-PeP°CĤCÍCHPePëPjPë P·P°CÍPsP±P°PjPë C, P°
C-PSC€C- PIPëPrPë PIP·P°C”PjPsPrC-P№.**

P·C,, PµPeC, Pë C-PSC€PëC... P»C-PeP°CĤCÍCHPePëC... P·P°CÍPsP±C-PI PSP°
CÍPëP»PrPµPSP°C,, C-P»

P”PsCÍP»C-PrP¶PµPSPSCÍ in vitro

PĤPµC, P°P±PsP»C-P·Pj CÍPëP»PrPµPSP°C,, C-P»Cí PIC-PrP±CíPIP°C”C, CHCÍCÍ
PíPµCĤPµPIP°P¶PSPs P·P° CíC‡P°CÍC, CĤ C-P·PsC,, PsCĤPjPë 3PĤ4 (PiPsP»PsPIPSPëP№
C€P»CÍC...) C, P° C-P·PsC,, PsCĤPjPë 2PŸ9 (PrCĤCíPiPsCĤCÍPrPSPëP№ C€P»CÍC...) C†PëC,
PsC...CĤPsPjCí P 450 (CYP). PŸPsPjCí C-PSPiC-P±C-C, PsCĤPë C†PëC...
C-P·PsC,, PµCĤPjPµPSC, C-PI PjPsP¶CíC, CH P·PSPëP¶CíPIP°C, Pë PeP»C-CĤPµPSCÍ
CÍPëP»PrPµPSP°C,, C-P»Cí, P° C-PSPrCíPeC, PsCĤPë C†PëC... C-P·PsC,, PµCĤPjPµPSC, C-PI
PjPsP¶CíC, CH PiC-PrPIPëC%oCíPIP°C, Pë PeP»C-CĤPµPSCÍ CÍPëP»PrPµPSP°C,, C-P»Cí.

P”PsCÍP»C-PrP¶PµPSPSCÍ in vivo

PµPsPíCíP»CÍC†C-P№PSPëP№ C,, P°CĤPjP°PePsPeC-PSPµC, PëC‡PSPëP№ P°PSP°P»C-P·
PrP°PSPëC... PeP»C-PSC-C‡PSPëC... PrPsCÍP»C-PrP¶PµPSCĤ
PíCĤPsPrPµPjPsPSCÍC, CĤCíPIP°PI P·PSPëP¶PµPSPSCÍ PeP»C-CĤPµPSCÍCí
CÍPëP»PrPµPSP°C,, C-P»Cí PíCĤPë P№PsPiPs PsPrPSPsC‡P°CÍPSPsPjCí
P·P°CÍC, PsCÍCíPIP°PSPSC- P· C-PSPiC-P±C-C, PsCĤP°PjPë CYP3A4 (C, P°PePëPjPë CÍPe
PePµC, PsPePsPSP°P·PsP», PµCĤPëC, CĤPsPjC-C†PëPS, C†PëPjPµC, PëPrPëPS). PÍPsC‡P° PíCĤPë
PsPrPSPsC‡P°CÍPSPsPjCí P·P°CÍC, PsCÍCíPIP°PSPSC- CÍPëP»PrPµPSP°C,, C-P»Cí C, P°
C-PSPiC-P±C-C, PsCĤC-PI CYP3A4 P·CĤPsCÍC, P°PSPSCÍ C‡P°CÍC, PsC, Pë PíPsP±C-C‡PSPëC...
CÍPIPëC%o PSPµ CÍPíPsCÍC, PµCĤC-PíP°P»PsCÍCÍ, CÍP»C-Pr CĤPsP·PíP»CÍPSCíC, Pë
PjPsP¶P»PëPIC-CÍC, CH P·P°CÍC, PsCÍCíPIP°PSPSCÍ PíPsC‡P°C, PePsPiPsC— PrPsP·Pë
CÍPëP»PrPµPSP°C,, C-P»Cí 25 PjPi.

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

PiC₇PsC, P_μP°P·Pë C₇PëC, PsPSP°PIC-C₇Cr, PrC₁P_¶P_μ PiPsC, CrP_¶PSPsPiPs
C-PSPiC-P±C-C, PsC₇P° P 450, Cr C₁C, P°PSC- C₇C-PIPSPsPIP°P_¶PSPsC—
PePsPSC†P_μPSC, C₇P°C†C-C— (500 PjPi 2 C₇P°P·PSP° PrPsP±Cr) C, P° C₁PëP»PrP_μPSP°C,, C-P»Cr
(PsPrPSPsC₇P°P·PsPIP° PrPsP·P° 100 PjPi) PiC₇PëP·PIPPrPëP»Ps PrPs PiC-PrPIPëC%P_μPSPSC_¶
C_{max} C₁PëP»PrP_μPSP°C,, C-P»Cr PSP° 300 % (Cr 4 C₇P°P·Pë) C, P° PiC-PrPIPëC%P_μPSPSC_¶
PiP»P°P·PjPsPIPSC— AUC C₁PëP»PrP_μPSP°C,, C-P»Cr PSP° 1000 % (Cr 11 C₇P°P·C-PI).
P_¶P_μC₇P_μP· 24 PiPsPrPëPSPë PiP»P°P·PjPsPIC- C₇C-PIPSC- C₁PëP»PrP_μPSP°C,, C-P»Cr PIC₁P_μ
C%P_μ C₁C, P°PSPsPIPëP»Pë PiC₇PëP±P»PëP·PSPs 200 PSPi/PjP», PiPsC₇C-PIPSC_¶PSPs P·
C₇C-PIPSP_μPj PiC₇PëP±P»PëP·PSPs 5 PSPi/PjP», C...P°C₇P°PeC, P_μC₇PSPëPj PrP»C_¶
P·P°C₁C, PsC₁CrPIP°PSPSC_¶ C, C-P»CHPePë C₁PëP»PrP_μPSP°C,, C-P»Cr, C%Ps
PIC-PrPiPsPIC-PrP°C” P·PSP°C†PSPsPjCr PIPiP»PëPIC₁ C₇PëC, PsPSP°PIC-C₇Cr PSP°
C€PëC₇PsPePëP_¶ C₁PiP_μPeC, C₇ C₁Cr±C₁C, C₇P°C, C-PI P 450. P_¶PëP»PrP_μPSP°C,, C-P» PSP_μ
PIPiP»PëPIP°C” PSP° C,, P°C₇PjP°PePsPeC-PSP_μC, PëPeCr C₇PëC, PsPSP°PIC-C₇Cr. P—
PsPiP»C_¶PrC₁ PSP° C†C- C,, P°C₇PjP°PePsPeC-PSP_μC, PëC†PSC- PrP°PSC- PsPrPSPsC†P°C₁PSP_μ
P·P°C₁C, PsC₁CrPIP°PSPSC_¶ C₁PëP»PrP_μPSP°C,, C-P»Cr C, P° C₇PëC, PsPSP°PIC-C₇Cr PSP_μ
C₇P_μPePsPjP_μPSPPrC₁C”C, CHC₁C_¶ (PrPëPI. C₇PsP·PrC-P» B«PhC₁Ps±P»PëPIPSC₁C, C-
P·P°C₁C, PsC₁CrPIP°PSPSC_¶B»); Cr PePsP_¶PSPsPjCr C₇P°P·C- PjP°PeC₁PëPjP°P»CHPSP°
PrPsP·P° C₁PëP»PrP_μPSP°C,, C-P»Cr PSP_μ PiPsPIPëPSPSP° PiP_μC₇P_μPIPëC%CrPIP°C, Pë 25 PjPi
PiC₇PsC, C_¶PiPsPj 48 PiPsPrPëPS.

PhPrPSPsC†P°C₁PSP_μ P·P°C₁C, PsC₁CrPIP°PSPSC_¶ C-PSPiC-P±C-C, PsC₇P° P’†P>-
PiC₇PsC, P_μP°P·Pë C₁P°PePIC-PSP°PIC-C₇Cr, C-PSPiC-P±C-C, PsC₇P° CYP3A4, Cr PrPsP·C-,
C%Ps P·P°P±P_μP·PiP_μC†Cr” C₇C-PIPSPsPIP°P_¶PSC₁ PePsPSC†P_μPSC, C₇P°C†C-C₇ (1200 PjPi 3
C₇P°P·Pë PSP° PrPsP±Cr), C, P° C₁PëP»PrP_μPSP°C,, C-P»Cr (100 PjPi PsPrPSPsC₇P°P·PsPIPs)
PiC₇PëP·PIPPrPëP»Ps PrPs PiC-PrPIPëC%P_μPSPSC_¶ C_{max} C₁PëP»PrP_μPSP°C,, C-P»Cr PSP°
140 % C, P° P·P±C-P»CHC€P_μPSPSC_¶ AUC C₁PëP»PrP_μPSP°C,, C-P»Cr PSP° 210 %. PkP_μ
PIPëC_¶PIP»P_μPSPs PIPiP»PëPIC₁ C₁PëP»PrP_μPSP°C,, C-P»Cr PSP°
C,, P°C₇PjP°PePsPeC-PSP_μC, PëPeCr C₁P°PePIC-PSP°PIC-C₇Cr (PrPëPI. C₇PsP·PrC-P»
B«P_¶PiPsC₁C-P± P·P°C₁C, PsC₁CrPIP°PSPSC_¶ C, P° PrPsP·PëB»).

P_μP_μC₇P_μPrP±P°C†P°C”C, CHC₁C_¶, C%Ps P±C-P»CHC€ PiPsC, CrP_¶PSC-
C-PSPiC-P±C-C, PsC₇Pë CYP3A4, C, P°PeC- C_¶Pe PeP_μC, PsPePsPSP°P·PsP» C, P°
C-C, C₇P°PePsPSP°P·PsP», P±CrPrC₁C, CH PjP°C, Pë P±C-P»CHC€ PIPëC₇P°P_¶P_μPSPëP_¶
PIPiP»PëPI.

P_μC₇Pë P·P°C₁C, PsC₁CrPIP°PSPSC- C₁PëP»PrP_μPSP°C,, C-P»Cr (100 PjPi PsPrPSPsC₇P°P·PsPIPs)
C, P° P_μC₇PëC, C₇PsPjC-C†PëPSC₁, PiPsPjC-C₇PSPsPiPs C-PSPiC-P±C-C, PsC₇P° CYP3A4, Cr
C₇C-PIPSPsPIP°P_¶PSPsPjCr C₁C, P°PSC- (500 PjPi 2 C₇P°P·Pë PSP° PrPsP±Cr PiC₇PsC, C_¶PiPsPj
5 PrPSC-PI) C₁PiPsC₁C, P_μC₇C-PiP°P»PsC₁C_¶ PiC-PrPIPëC%P_μPSPSC_¶ P_¶UC
C₁PëP»PrP_μPSP°C,, C-P»Cr PSP° 182 %. Pj P·PrPsC₇PsPIPëC... PrPsP±C₇PsPIPsP»CHC†C-PI
C†PsP»PsPIC-C†PsC— C₁C, P°C, C- PSP_μ C₁PiPsC₁C, P_μC₇C-PiP°P»PsC₁C_¶ PIPiP»PëPIC₁
P°P·PëC, C₇PsPjC-C†PëPSC₁ (500 PjPi PSP° PrPsP±Cr PiC₇PsC, C_¶PiPsPj 3 PrC-P±) PSP° AUC,
P_¶max, T_{max}, PePsPSC₁C, P°PSC, Cr C€PIPëPrPePsC₁C, C- P_μP»C-PjC-PSP°C†C-C— C, P°
PiPsPrP°P»CHC€PëP_¶ PiP_μC₇C-PsPr PSP°PiC-PIPIPëPIP_μPrP_μPSPSC_¶
C₁PëP»PrP_μPSP°C,, C-P»Cr P°P±Ps P_¶PsPiPs PiPsP»PsPIPSPsPiPs C†PëC₇PeCrP»C₇C₇C†PsPiPs
PjP_μC, P°P±PsP»C-C, Cr. P_¶PëPjP_μC, PëPrPëPS (C-PSPiC-P±C-C, PsC₇ C†PëC, PsC...C₇PsPjCr P 450
C, P° PSP_μC₁PiP_μC†PëC,, C-C†PSPëP_¶ C-PSPiC-P±C-C, PsC₇ CYP3A4) Cr PrPsP·C- 800 PjPi PiC₇Pë
PsPrPSPsC†P°C₁PSPsPjCr P·P°C₁C, PsC₁CrPIP°PSPSC- P·C- C₁PëP»PrP_μPSP°C,, C-P»PsPj Cr
PrPsP·C- 50 PjPi Cr P·PrPsC₇PsPIPëC... PrPsP±C₇PsPIPsP»CHC†C-PI PiC₇PëP·PIPPrPëPI PrPs
PiC-PrPIPëC%P_μPSPSC_¶ PiP»P°P·PjPsPIPëC... PePsPSC†P_μPSC, C₇P°C†C-P_¶
C₁PëP»PrP_μPSP°C,, C-P»Cr PSP° 56 %.

P“C₇H₁₄PM₂PC₂, C₇H₁₄CrC, PsPIP₂PM₂ C₇C-Pe C” C₇P»P°P±PeP₂Pj C-PSPiC-P±C-C, PsC₇H₁₄PsPj CYP3A4 Cr C₇C, C-PSC±C- PeP₂PC₇PM₂±PSP₂PeCr C- PjPsP₂PM₂ PIP₂PeP»P₂PeP°C, P₂ PiPsPjC-C₇H₁₄PSP₂ PiC-PrPIP₂PC₇PM₂±PSPSC₇ C₇H₁₄-PIPSC₇ C₇P₂»PrP₂PM₂PSP°C,,C-P»Cr Cr PiP»P°P·PjC- PeC₇H₁₄PsPIC-.

P₂PrPSPsC₇H₁₄P°P·PsPIP₂ P·P°C₇C, PsC₇CrPIP°PSPSC₇ P°PSC, P°C±P₂PrC-PI (PjP°PiPSC-C₇ PiC-PrC₇H₁₄PsPeC₇P₂PrC₇/P°P»C₇H₁₄PjC-PSC-C₇ PiC-PrC₇H₁₄PsPeC₇P₂PrC₇) PSP₂ PIP₂P»P₂PIP°P»Ps PSP° P±C-PsPrPsC₇C, CrPiPSC-C₇C, C₇H C₇P₂»PrP₂PM₂PSP°C,,C-P»Cr.

P₂PsC₇±P° PrPsC₇P»C-PrP₂PM₂PSC₇ C₇PiP₂PM₂±P₂C,,C-±PSPsC— PIP·P°C”PjPsPrC-C— P· CrC₇C-PjP° P»C-PeP°C₇H₁₄CrP₂PeP₂PjP₂ P·P°C₇PsP±P°PjP₂ PSP₂ PiC₇H₁₄PsPIP₂PrP₂»PsC₇C₇, P·P° PrP°PSP₂PjP₂ PiPsPiCrP»C₇C₇±C-PM₂PSPsPiPs C,,P°C₇H₁₄PjP°PePsPeC-PSP₂PM₂, P₂C±PSPsPiPs P°PSP°P»C-P·Cr, C,,P°C₇H₁₄PjP°PePsPeC-PSP₂PM₂, P₂PeP° C₇P₂»PrP₂PM₂PSP°C,,C-P»Cr PSP₂ P·PjC-PSC₇PIP°P»P°C₇C₇ PiC₇H₁₄P₂ PM₂PsPiPs PsPrPSPsC₇±P°C₇PSPsPjCr P·P°C₇C, PsC₇CrPIP°PSPSC- P· P»C-PeP°C₇H₁₄CrP₂PeP₂PjP₂ P·P°C₇PsP±P°PjP₂, C%Ps PSP°P»P₂PM₂P°C, C₇H PrPs PiC₇CrPiP₂ C-PSPiC-P±C-C, PsC₇H₁₄-PI CYP2C9 (C, PsP»P±CrC, P°PjC-Pr, PIP°C₇C,,P°C₇H₁₄P₂PS, C,,P₂PM₂PSC-C, PsC—PS), PiC₇CrPiP₂ C-PSPiC-P±C-C, PsC₇H₁₄-PI CYP2D6 (C, P°PeP₂C... C₇PM₂ C₇P₂»P₂PM₂PeC, P₂PIPSC- C-PSPiC-P±C-C, PsC₇H₁₄P₂ P·PIP₂C₇H₁₄PsC, PSPsPiPs P·P°C...PsPiP»P₂PM₂PSPSC₇ C₇P₂PM₂C₇PsC, PsPSC-PSC₇, C, C₇H₁₄P₂C±P₂PeP»C-C±PSC- P°PSC, P₂PrP₂PiC₇H₁₄PM₂C₇P°PSC, P₂), PiC₇CrPiP₂ C, C-P°P·P₂PrPSP₂C... C, P° C, C-P°P·P₂PrPsPiPsPrC-P±PSP₂C... PrC-C₇H₁₄PM₂C, P₂PeC-PI, PiP₂PM₂C, P»C₇H₁₄PsPIP₂C... C, P° PeP°P»C-PM₂·P±P₂PM₂C₇-PiP°C₇C±P₂C... PrC-C₇H₁₄PM₂C, P₂PeC-PI, C-PSPiC-P±C-C, PsC₇H₁₄-PI P°PSPiC-PsC, P₂PM₂P·P₂PSPiP₂PM₂C₇PM₂C, PIPsC₇H₁₄PIP°P»C₇H₁₄PSPsPiPs C,,P₂PM₂H₁₄PjP₂PM₂PSC, Cr (P₂PM₂α), P°PSC, P°PiPsPSC-C₇C, C-PI PeP°P»C₇H₁₄±C-C₇, P°PSC, P°PiPsPSC-C₇C, C-PI β- P°PrC₇H₁₄PM₂PSPsC₇H₁₄PM₂±P₂PM₂PsC, PsC₇H₁₄-PI P°P±Ps C-PSP₂CrPeC, PsC₇H₁₄-PI PjP₂PM₂C, P°P±PsP»C-P·PjCr CYP450 (C, P°PeP₂C... C₇PM₂ C₇H₁₄P₂C,,P°PjPiC-C±P₂PS, P±P°C₇H₁₄P±C-C, CrC₇H₁₄P°C, P₂).

Pj PiC₇H₁₄PsC₇±P₂PM₂C₇C- PrPsC₇P»C-PrP₂PM₂PSPSC₇ P·P° CrC₇±P°C₇C, C₇H P·PrPsC₇H₁₄PsPIP₂C... C±PsP»PsPIC-PeC-PI-PrPsP±C₇H₁₄PsPIP₂»C₇H₁₄±C-PI PsPrPSPsC₇±P°C₇PSP₂ P·P°C₇C, PsC₇CrPIP°PSPSC₇ P°PSC, P°PiPsPSC-C₇C, P° P₂PM₂PrPsC, P₂PM»C-PSC₇ P±PsC₇PM₂PSC, P°PSC₇ (PiPsPjC-C₇H₁₄PSP₂PM₂ C-PSP₂CrPeC, PsC₇ CYP3A4, CYP2C9 C-, PjPsP₂PM₂»P₂PIP₂, CYP2C19) Cr C₇H₁₄-PIPSPsPIP°P₂PM₂PjCr C₇C, P°PSC- (125 PjPi 2 C₇H₁₄P°P·P₂ PSP° PrPsP±Cr) C, P° C₇P₂»PrP₂PM₂PSP°C,,C-P»Cr Cr C₇H₁₄-PIPSPsPIP°P₂PM₂PjCr C₇C, P°PSC- (80 PjPi 3 C₇H₁₄P°P·P₂ PSP° PrPsP±Cr) PiC₇H₁₄P₂·PIP₂PrP₂»Ps PrPs P·PSP₂P₂PM₂PSPSC₇ AUC C- C_{max} C₇P₂»PrP₂PM₂PSP°C,,C-P»Cr PSP° 62,6 % C- 55,4 % PIC-PrPiPsPIC-PrPSPs. P₂PsPjCr PsPrPSPsC₇±P°C₇PSP₂ P·P°C₇C, PsC₇CrPIP°PSPSC₇ C, P°PeP₂C... PiPsC, CrP₂PM₂P₂C... C-PSP₂CrPeC, PsC₇H₁₄-PI CYP3A4, C₇PM₂ C₇H₁₄P₂C,,P°PjPiC-PS, PjPsP₂PM₂ PiC₇H₁₄P₂·PIP₂PrP₂»P₂ PrPs P±C-P»C₇H₁₄PeC PIP₂C₇H₁₄P°P₂PM₂PSPsPiPs P·PSP₂P₂PM₂PSPSC₇ PePsPSC±P₂PM₂PSC, C₇H₁₄P°C±C-C— C₇P₂»PrP₂PM₂PSP°C,,C-P»Cr PI PiP»P°P·PjC- PeC₇H₁₄PsPIC-.

P₂K-C-PePsC₇H₁₄P°PSP₂PrP₂» C₇PM₂IP»C₇C” C₇PsP±PsC₇ PiC-P±C₇H₁₄P₂Pr P°PeC, P₂PIP°C, PsC₇H₁₄P° PeP°P»C₇H₁₄±C-C”PIP₂C... PeP°PSP°P»C-PI C, P° PSC-C, C₇H₁₄P°C, Cr. P₂K-C, C₇H₁₄P°C, PSP₂PM₂ PePsPjPiPsPSP₂PSC, P·CrPjPsPIP»C₇C” PjPsP₂PM₂»P₂PI-C-C₇C, C₇H PM₂PsPiPs C₇PM₂C₇PM₂PsP·PSPsC— PIP·P°C”PjPsPrC-C— P·C- C₇P₂»PrP₂PM₂PSP°C,,C-P»PsPj.

P·C,,P₂PM₂PeC, P₂ C₇P₂»PrP₂PM₂PSP°C,,C-P»C₇ PSP° C-PSC₇C- P»C-PeP°C₇H₁₄CrP₂PeC- P·P°C₇PsP±P₂

P₂PsC₇P»C-PrP₂PM₂PSPSC₇ in vitro

P₂Y₂P₂»PrP₂PM₂PSP°C,,C-P» B₇” C₇P»P°P±PeP₂PM₂ C-PSPiC-P±C-C, PsC₇H₁₄ C-P·PsC,,PsC₇H₁₄Pj 1P₂, 2C9, 2C19, 2D6, 2E1 C, P° 3P₂4 C±P₂C, PsC...C₇H₁₄PsPjCr P 450 (P₂P₂50 > 150 PjPePjPsP»C₇H). P₂K₇CrPeC-P»C₇H₁₄PeP₂ PiC-PePsPIC- PiP»P°P·PjPsPIC- PePsPSC±P₂PM₂PSC, C₇H₁₄P°C±C-C— C₇P₂»PrP₂PM₂PSP°C,,C-P»Cr PrPsC₇H₁₄-PIPSC₇C₇C, C₇H PiC₇H₁₄P₂±P»P₂·PSPs B B B B B B B 1

PjPePjPsP»ЧБ, PIPiP»PëPI PíCтjPμPíP°CтjP°C,Cr PSP° PeP»C-ЧтjPμPSCí CíCfP±CíC,CтjP°C,C-PI C†PëC... C-P·PsC,,PμCтjPjPμPSC,C-PI PjP°P»PsPNëPjPsPIC-ЧтjPSPëPNë.

P'C-PrCíCfC,PSC- PrP°PSC- C%PsPrPs PIP·P°C”PjPsPrC-C— CíPëP»PrPμPSP°C,,C-P»Cf C- C,P°PePëC... PSPμCíPíPμC†PëC,,C-C†PSPëC... C-PSPiC-P±C-C,PSCтjC-PI C,,PSCíC,,PsPrC-PμCíC,PμCтjP°P·Pë, C†Pe C,PμPsC,,C-P»C-PS C,P° PrPëPiC-ЧтjPëPrP°PjPsP».

P”PsCíP»C-PrP¶PμPSPSCII in vivo

PhCíPeC-P»ЧБPePëB 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”PнPα, P±CfP»Ps PICíC,P°PSPsPIP»PμPSPs, C%Ps CíPëP»PrPμPSP°C,,C-P» PíPsC,PμPSC†C-ЧтC” PiC-PíPsC,PμPSP·PëPIPSCf PrC-Чт PSC-C,CтjP°C,C-PI, C,PSPjCf PNëPsPiPs PsPrPSPsC†P°CíPSPμ P·P°CíC,PSCíCfPIP°PSPSCII P· PrPsPSPsCтjP°PjPë PsPeCíPëPrCf P°P·PsC,Cf P°P±Ps P· PSC-C,CтjP°C,P°PjPë Cí P±CfPrЧБ-C†PeC-PNë C,,PSCтjPjC- PíCтjPsC,PëPíPsPeP°P·P°PSPμ (PrPëPI. CтjPsP·PrC-P» B«PμCтjPsC,PëPíPsPeP°P·P°PSPSCII»).

P C-PsC†PëPiCáP°C,. P”PsPeP»C-PSC-C†PSC- PrPsCíP»C-PrP¶PμPSPSCII PíCтjPsPrPμPjPsPSCíC,CтjCfPIP°P»Pë P°PrPëC,PëPIPSPëPNë CíPëCíC,PμPjPSPëPNë PμC,,PμPeC, P·PSPëP¶PμPSPSCII P°CтjC,PμCтjC-P°P»ЧБPSPsPiPs C,PëCíPeCf PíCтjPë PsPrPSPsC†P°CíPSPsPjCf P·P°CíC,PSCíCfPIP°PSPSC- C-PSPiC-P±C-C,PSCтjC-PI PαP”P·5 P· CтjC-PsC†PëPiCfP°C,PSPj. PнP»C-PSC-C†PSC- PrPsCíP»C-PrP¶PμPSPSCII PíCтjPsPrPμPjPsPSCíC,CтjCfPIP°P»Pë, C%Ps CтjC-PsC†PëPiCfP°C, PíPsCíPëP»ЧтC” PiC-PíPsC,PμPSP·PëPIPSCf PrC-Чт C-PSPiC-P±C-C,PSCтjC-PI PαP”P·5.

PJ PíP°C†C-C”PSC,C-PI, C†PeC- P±CтjP°P»Pë CíC†P°CíC,ЧБ Cí PrPsCíP»C-PrP¶PμPSPSC-, PSPμ CíPíPsCíC,PμCтjC-PiP°P»PsCíCн PíPsP·PëC,PëPIPSPsPiPs PeP»C-PSC-C†PSPsPiPs PμC,,PμPeC,Cf PIC-Pr PsPrPSPsC†P°CíPSPsPiPs P·P°CíC,PSCíCfPIP°PSPSCII C-PSPiC-P±C-C,PSCтjC-PI PαP”P·5 P· CтjC-PsC†PëPiCfP°C,PSPj. PμCтjPsC,PëPíPsPeP°P·P°PSPμ PsPrPSPsC†P°CíPSPμ P·P°CíC,PSCíCfPIP°PSPSCII CтjC-PsC†PëPiCfP°C,Cf P· C-PSPiC-P±C-C,PSCтjP°PjPë PαP”P·5 (Cf C,PSPjCf C†PëCíP»C- P·C- CíPëP»PrPμPSP°C,,C-P»PsPj) (PrPëPI. CтjPsP·PrC-P»B B«PμCтjPsC,PëPíPsPeP°P·P°PSPSCII»).

PhPrPSPsC†P°CíPSPμ P·P°CíC,PSCíCfPIP°PSPSCII CíPëP»PrPμPSP°C,,C-P»Cf C,P° P±P»PsPeP°C,PSCтjC-PI α-P°PrCтjPμPSPsCтjPμC†PμPíC,PSCтjC-PI PjPsP¶Pμ PíCтjPëP·PIPμCíC,Pë PrPs CтjPsP·PIPëC,PeCf CíPëPjPíC,PSPjP°C,PëC†PSPsC— PiC-PíPsC,PμPSP·C-C— Cí PrPμC†PePëC... CíC...PëP»ЧБPSPëC... PrPs C†ЧБPsPiPs PíP°C†C-C”PSC,C-PI. PýP°PeP° CтjPμP°PeC†C-C† PSP°PNëC†P°CíC,C-C€Pμ PIPëPSPëPeP°P»P° PíCтjPsC,C†PiPsPj 4 PiPsPrPëPS PíC-CíP»C†P·P°CíC,PSCíCfPIP°PSPSCII CíPëP»PrPμPSP°C,,C-P»Cf (PrPëPI. CтjPsP·PrC-P»Pë B«PhCíPsP±P»PëPIPsCíC,C- P·P°CíC,PSCíCfPIP°PSPSCII» C,P° B«PýPíPsCíC-P± P·P°CíC,PSCíCfPIP°PSPSCII C,P° PrPsP·PëB»). PJ PíCтjPsC†PμCíC- 3 PrPsCíP»C-PrP¶PμPSCн CíPíPμC†PëC,,C-C†PSPsC— PIP·P°C”PjPsPrC-C— P»C-PeP°CтjCíCнPePëC... P·P°CíPsP±C-PI P±P»PsPeP°C,PSCтj α-P°PrCтjPμPSPsCтjPμC†PμPíC,PSCтjC-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,PSCíCfPsPICfPIP°P»Pë PsPrPSPsC†P°CíPSPs PíP°C†C-C”PSC,P°Pj C-P· PrPsP±CтjPsC†PeC-CíPSPsCн PiC-PíPμCтjPíP»P°P·C-C”Cн PíPμCтjPμPrPjC-C...CfCтjPsPiPsC— P·P°P»PsP·Pë, CíC,P°P±C-P»C-P·P°C†C-C† CíC,P°PSCf C†PePëC... P±CfP»P° PrPsCíC†PiPSCfC,P° PíCтjPë P·P°CíC,PSCíCfPIP°PSPSC- PrPsPeCíP°P·PsP·PëPSCf. PJ C†PëC... PíPsPíCfP»C†C-C†C... CíPíPsCíC,PμCтjC-PiP°P»PsCíC† CíPμCтjPμPrPSC” PrPsPrP°C,PePsPIPμ P·PSPëP¶PμPSPSCII P°CтjC,PμCтjC-P°P»ЧБPSPsPiPs C,PëCíPeCf Cí PíPsP»PsP¶PμPSPSC- P»PμP¶P°C†Pë PSP° 7/7 PjPj CтjC,. CíC,.. 9/5 PjPj CтjC,. CíC,. C,P° 8/4 PjPj CтjC,. CíC,. C,P° CíPμCтjPμPrPSC” P·PSPëP¶PμPSPSCII P°CтjC,PμCтjC-P°P»ЧБPSPsPiPs C,PëCíPeCf Cí PíPsP»PsP¶PμPSPSC-

СГС,PsCЦC±Pë PSP° 6/6 PjPj CГC,.. СГС,.. 11/4 PjPj CГC,.. СГС,.. 4/5 PjPj CГC,.. СГС..
PIC-PrPiPsPIC-PrPSPs. PμCГPë PsPrPSPsC±P°СГPSPsPjCр P·P°СГC,PsCГCрPIP°PSPSC-
CГPëP»PrPμPSP°C,,C-P»Cр C,P° PrPsPeCГP°P·PsP·PëPSCр PiP°C†C-C”PSC,P°Pj,
CГC,P°P±C-P»C-P·P°C†C-CЦ CГC,P°PSCр CЦPePëC... P±CрP»P° PrPsCГCЦPiPSCрC,P° PiCГPë
P·P°СГC,PsCГCрPIP°PSPSC- PrPsPeCГP°P·PsP·PëPSCр, C-PSPsPrC-
PiPsPIC-PrPsPjP»CЦP»PsCГCЦ PiCГPs CГPsP·PIPëC,PsPe CГPëPjPiC,PsPjP°C,PëC±PSPsC—
PsCГC,PsCГC,P°C,PëC±PSPsC— PiC-PiPsC,PμPSP·C-C—. Pj C†PëC...
PiPsPIC-PrPsPjP»PμPSPSCЦC... P№C€P»PsCГCЦ PiCГPs PIPëPiP°PrPePë
P·P°PiP°PjPsCГPsC±PμPSPSCЦ C,P° CГC,P°PS PiPμCГPμPr PSPμPiCГPëC,PsPjPSC-CГC,CГ,
P°P»Pμ P±PμP· CГPëPSPePsPiPμ.

PкPμ CГPiPsCГC,PμCГC-PiP°P»PsCГCЦ P¶PsPrPSPëC... P·PSP°C±CрC%PëC...
PIP·P°C”PjPsPrC-P№ PiCГPë PsPrPSPsC±P°СГPSPsPjCр P·P°СГC,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 PiCГPëP·PIPsPrPëPI PrPs PiPsPrPsPIP¶PμPSPSCЦ
C±P°СГCр PeCГPsPIPsC,PμC±C-, CГPiCГPëC±PëPSPμPSPsC— P·P°СГC,PsCГCрPIP°PSPSCЦPj
P°C†PμC,PëP»CГP°P»C-C†PëP»PsPIPsC— PePëCГP»PsC,Pë (150 PjPi).

PŸPëP»PrPμPSP°C,,C-P» (50 PjPi) PSPμ PiPsC,PμPSC†C-CГPIP°PI PiC-PiPsC,PμPSP·PëPIPsCр
PrC-CГ P°P»PePsPiPsP»CГ Cр P·PrPsCГPsPIPëC... PrPsP±CГPsPIPsP»CГC†C-PI PiCГPë
CГPμCГPμPrPSCГPsPjCр PjP°PeCГPëPjP°P»CГPSPsPjCр CГC-PIPsC- PμC,P°PSPsP»Cр
PeCГPsPIC- 80 PjPi/PrP».

Pj PiP°C†C-C”PSC,C-PI, CЦPeC- P·P°СГC,PsCГCрPICрPIP°P»Pë CГPëP»PrPμPSP°C,,C-P», PSPμ
CГPiPsCГC,PμCГC-PiP°P»PsCГCЦ P¶PsPrPSPëC... PIC-PrPjC-PSPSPsCГC,PμP№ PiCГPsC,,C-P»CГ
PiPsP±C-C±PSPëC... PμC,,PμPeC,C-PI PiPsCГC-PIPsCЦPSPs P· PiP»P°C†PμP±Ps PiCГPë
PsPrPSPsC±P°СГPSPsPjCр P·P°СГC,PsCГCрPIP°PSPSC- C,P°PePëC... PeP»P°СГC-PI
PiC-PiPsC,PμPSP·PëPIPsPëC... P»C-PeP°CГCГCГPePëC... P·P°СГCрP±C-PI, CЦPe
PrC-CрCГPμC,PëPePë, P±P»PsPeP°C,PsCГPë β-P°PrCГPμPSPsCГPμC†PμPiC,PsCГC-PI,
C-PSPiC-P±C-C,PsCГPë PђPμPα, P°PSC,P°PiPsPSC-CГC,Pë P°PSPiC-PsC,PμPSP·PëPSCр P†P†,
P°PSC,PëPiC-PiPμCГC,PμPSP·PëPIPsC- P»C-PeP°CГCГCГPeC- P·P°СГCрP±Pë
(CГCрPrPëPSPsCГPsP·C€PëCГCГPIP°P»CГPSC- C,P° C†PμPSC,CГP°P»CГPSPsC— PrC-C—),
P±P»PsPeP°C,PsCГPë P°PrCГPμPSPμCГPiC-C±PSPëC... PSPμP№CГPsPSC-PI,
P±P»PsPeP°C,PsCГPë PeP°P»CГC†C-C”PIPëC... PeP°PSP°P»C-PI C,P° P±P»PsPeP°C,PsCГPë α-
P°PrCГPμPSPsCГPμC†PμPiC,PsCГC-PI.

Pj CГPiPμC†C-P°P»CГPSPsPjCр PrPsCГP»C-PrP¶PμPSPSC- PIP·P°C”PjPsPrC-C— PiCГPë
PsPrPSPsC±P°СГPSPsPjCр P·P°СГC,PsCГCрPIP°PSPSC- CГPëP»PrPμPSP°C,,C-P»Cр (100 PjPi) C,P°
P°PjP»PsPrPëPiC-PSCр PiP°C†C-C”PSC,P°Pj C-P· P°CГC,PμCГC-P°P»CГPSPsCГ
PiC-PiPμCГC,PμPSP·C-C”CГ CГPiPsCГC,PμCГC-PiP°P»PsCГCЦ PrPsPrP°C,PePsPIPμ
P·PSPëP¶PμPSPSCЦ CГPëCГC,PsP»C-C±PSPsPiPs P°CГC,PμCГC-P°P»CГPSPsPiPs C,PëCГPeCр PI
PiPsP»PsP¶PμPSPSC- P»PμP¶P°C±Pë PSP° 8 PjPj CГC,.. CГC,.. P·C-PrPiPsPIC-PrPSPμ
P·PSPëP¶PμPSPSCЦ PrC-P°СГC,PsP»C-C±PSPsPiPs P°CГC,PμCГC-P°P»CГPSPsPiPs C,PëCГPeCр
CГC,P°PSPsPIPëP»Ps 7 PjPj CГC,.. CГC,.. PjC- PrPsPrP°C,PePsPIC- P·PSPëP¶PμPSPSCЦ
P°CГC,PμCГC-P°P»CГPSPsPiPs C,PëCГPeCр P±CрP»Pë PiPsCГC-PIPsCЦPSPSPëPjPë P·
C,P°PePëPjPë, C%Ps CГPiPsCГC,PμCГC-PiP°P»PëCГCЦ PiCГPë P·P°СГC,PsCГCрPIP°PSPSC-
P»PëC€Pμ CГPëP»PrPμPSP°C,,C-P»Cр Cр P·PrPsCГPsPIPëC... PrPsP±CГPsPIPsP»CГC†C-PI.

PŸPëP»PrPμPSP°C,,C-P» Cр PrPsP·C- 100 PjPi PSPμ PIPiP»PëPIP°PI PSP°
C,,P°CГPjP°PePsPeC-PSPμC,PëC±PSC- PiPsPeP°P·PSPëPePë C-PSPiC-P±C-C,PsCГC-PI P’P†P»-

PiC7PsC,PuP°P·Pë, CÍP°PePIC-PSP°PIC-C7Cf C,P° C7PëC,PsPSP°PIC-C7Cf, CIPeC- C" CÍCfP±CÍC,C7P°C,P°PjPë CYP3A4.

PJ P·PrPsC7PsPIPëC... PrPsP±C7PsPIPsP»CHC†C-PI C±PsP»PsPIC-C±PsC— CÍC,P°C,C- P·P°CÍC,PsCÍCfPIP°PSPSCÍ CÍPëP»PrPµPSP°C,,C-P»Cf Cí C7C-PIPsPIP°P¶PSPsPjCf CÍC,P°PSC- (80 PjPi 3 C7P°P·Pë PSP° PrPsP±Cf) PiC7PëP·PIPsPrPëP»Ps PrPs PiC-PrPIPëC%oPµPSPSCÍ AUC C,P° C_{max} P±PsCÍPµPSC,P°PSCf (125 PjPi 2 C7P°P·Pë PSP° PrPsP±Cf) PSP° 49,8 % C,P° 42 % PIC-PrPiPsPIC-PrPSPs.

P"PsPrP°PIP°PSPSCÍ PsPrPSPsC7P°P·PsPIPsC— PrPsP·Pë CÍPëP»PrPµPSP°C,,C-P»Cf PrPs CÍP°PeCfP±C-C,C7PëP»Cf / PIP°P»CÍP°C7C,P°PSCf Cí C7C-PIPsPIP°P¶PSPsPjCf CÍC,P°PSC- Cí PiP°C†C-C"PSC,C-PI P· P°C7C,PµC7C-P°P»CHPSPsCf PiC-PiPµC7C,PµPSP·C-C"Ch P±CfP»Ps PiPsPiCÍP·P°PSPµ P·C- P·PSP°C±PSPs P±C-P»CHC€ PIPëC7P°P¶PµPSPëPj P·PSPëP¶PµPSPSCÍPj P°C7C,PµC7C-P°P»CHPSPsPiPs C,PëCÍPeCf PiPsC7C-PIPsCÍPSPs P· PiC7PëPµPsPjPsPj C,C-P»CHPePë CÍP°PeCfP±C-C,C7PëP»Cf/PIP°P»CÍP°C7C,P°PSCf. PÿP°PePëPj C±PëPSPsPj, CÍP»C-Pr PrPsC,C7PëPjCfPIP°C,PëCÍCÍ PsP±PµC7PµP¶PSPsCÍC,C- PiC7Pë PiPsC±P°C,PeCf P·P°CÍC,PsCÍCfPIP°PSPSCÍ CÍPëP»PrPµPSP°C,,C-P»Cf Cí PiP°C†C-C"PSC,C-PI, CIPeC- PsC,C7PëPjCfC7C,Ch P»C-PeCfPIP°PSPSCÍ CÍP°PeCfP±C-C,C7PëP»PsPj/PIP°P»CÍP°C7C,P°PSPsPj.

PñCÍPsP±P»PëPIPsCÍC,C- P·P°CÍC,PsCÍCáPIP°PSPSCÍ.

P"Ps PiPsC±P°C,PeCf C,PµC7P°PiC-C— CÍP»C-Pr P·C-P±C7P°C,Pë PjPµPrPëC±PSPëPµ P°PSP°PjPSPµP· PiP°C†C-C"PSC,P° C,P° PiC7PsPIPµCÍC,Pë C,,C-P·PëPeP°P»CHPSPµ PsP±CÍC,PµP¶PµPSPSCÍ PrP»CÍ PrC-P°PiPSPsCÍC,PëPePë PµC7PµPeC,PëP»CHPSPsC— PrPëCÍC,,CfPSPeC†C-C— C,P° PIPëP·PSP°C±PµPSPSCÍ C—C— PjPsP¶P»PëPIPëC... PiC7PëC±PëPS.

PαP°PeC,PsC7Pë C7PëP·PëPeCá CÍPµC7C†PµPIPs-CÍCáPrPëPSPSPëC... P·P°C...PIPsC7C7PIP°PSCñ

PñCÍPeC-P»CHPePë CÍPµPeCÍCfP°P»CHPSP° P°PeC,PëPIPsC-CÍC,Ch CÍCfPiC7PsPIPsPrP¶CfC" C,ChCÍCÍ PiPµPIPsPëPj C7PëP·PëPePsPj P· P±PsPeCf CÍPµC7C†CÍ, PrPs PiPsC±P°C,PeCf P±CfPrCh-CÍPePsPiPs P»C-PeCfPIP°PSPSCÍ PµC7PµPeC,PëP»CHPSPsC— PrPëCÍC,,CfPSPeC†C-C— P»C-PeP°C7 PjP°C" PsC†C-PSPëC,Pë CÍC,P°PS CÍPµC7C†PµPIPs- CÍCfPrPëPSPSPsC— CÍPëCÍC,PµPjPë PiP°C†C-C"PSC,P°. PÿPëP»PrPµPSP°C,,C-P» C±PëPSPëC,Ch CÍCfPrPëPSPsC7PsP·C€PëC7C7PIP°P»CHPSCf PrC-Cñ, C%oPs PiC7PsCÍPIP»CÍC" C,ChCÍCÍ P»PµPiPePëPj C,P° PePsC7PsC,PePsC±P°CÍPSPëPj P·PSPëP¶PµPSPSCÍPj P°C7C,PµC7C-P°P»CHPSPsPiPs C,PëCÍPeCf. P"Ps PiC7PëP·PSP°C±PµPSPSCÍ CÍPëP»PrPµPSP°C,,C-P»Cf P»C-PeP°C7 PjP°C" C7PµC,PµP»CHPSPs P·PIP°P¶PëC,Pë, C±Pë PjPsP¶Pµ C,P°PePëPµ PµC,,PµPeC, PSPµCÍPiC7PëCÍC,P»PëPIPs PIPiP»PëPIP°C,Pë PSP° PiP°C†C-C"PSC,C-PI C-P· PiPµPIPsPëPjPë PsCÍPSPsPIPsPëPjPë P·P°C...PIPsC7C7PIP°PSPSCÍPjPë, PsCÍPsP±P»PëPIPs Cí PePsPjP±C-PSP°C†C-C— C-P· CÍPµPeCÍCfP°P»CHPSPsCñ P°PeC,PëPIPsC-CÍC,Cñ. P"Ps PiP°C†C-C"PSC,C-PI C-P· PiC-PrPIPëC%oPµPSPsCñ C±CfC,P»PëPIC-CÍC,Cñ PrPs PIP°P·PsPrPëP»CÍC,P°C,PsC7C-PI PSP°P»PµP¶P°C,Ch PiP°C†C-C"PSC,Pë C-P· PsP±CÍC,C7CfPeC†C-C"Ch PIPëPIC-PrPSPsPiPs C,C7P°PeC,Cf P»C-PIPsPiPs C€P»CfPSPsC±PeP° (PSP°PiC7PëPeP»P°Pr, CÍC,PµPSPsP· P°PsC7C,Pë, PiC-PiPµC7C,C7PsC,,C-C±PSP° PsP±CÍC,C7CfPeC,PëPIPSP° PeP°C7PrC-PsPjC-PsPiP°C,C-CÍ) C,P° PiP°C†C-C"PSC,Pë C-P· C7C-PrPeC-CÍPSPëPj CÍPëPSPPrC7PsPjPsPj PjCfP»CHC,PëCÍPëCÍC,PµPjPSPsC— P°C,C7PsC,,C-C—, PsPrPSPëPj C-P·

P'PμP·PiPμPeP° C,P° PμC,,PμPeC,PëPIPSC-Cl'c,CH PsPrPSPsC±P°Cl'PSPsPiPs
P·P°Cl'c,PsCl'CrPIP°PSPSCl' Cl'PëP»PrPμPSP°C,,C-P»Cr P· C-PSC€PëPjPë
C-PSPIc-P±C-C,PsCtjP°PjPë PαP''P·5 P°P±Ps C-PSC€PëPjPë PiCtjPμPiP°CtjP°c,P°PjPë PrP»Cl'
P»C-PeCrPIP°PSPSCl' PiC-PiPμCtjC,PμPSP·C-C— P»PμPiPμPSPμPiPsC— P°CtjC,PμCtjC-C—,
Cl'PeC- PjC-Cl'c,Cl'c,CH Cl'PëP»PrPμPSP°C,,C-P», C±Pë P· C-PSC€PëPjPë
PiCtjPμPiP°CtjP°c,P°PjPë PrP»Cl' P»C-PeCrPIP°PSPSCl' PμCtjPμPeC,PëP»CHPSPsC—
PrPëCl'c,,CrPSPeC†C-C— PSPμ PIPëPIC±P°P»PëCl'Cl'. PÿPsPjCr P·P°Cl'c,PsCl'CrPIP°PSPSCl'
C,P°PePëC... PePsPjP±C-PSP°C†C-PNë PSPμ CtjPμPePsPjPμPSPPrPsPIP°PSPμ.

P'PiP»PëPI PSP° P·C-Ctj

PμPsPIC-PrPsPjP»Cl'P»PsCl'cH PiCtjPs PIPëPSPëPePSPμPSPSCl' PrPμC,,PμPeC,C-PI P·PsCtjCr
C,P° PIPëPiP°PrPePë PSPμP°CtjC,PμCtjC-P°P»CHPSPsC— PiPμCtjPμPrPSCHPSc—
C-C€PμPjC-C±PSPsC— PSPμPICtjPsPiP°c,C-C— P·PsCtjPsPiPs PIP° Cr
P·PIBtj™ Cl'P·PeCr C-P· P·P°Cl'c,PsCl'CrPIP°PSPSCl'Pj Cl'PëP»PrPμPSP°C,,C-P»Cr C,P°
C-PSC€PëC... C-PSPIc-P±C-C,PsCtjC-PI PαP''P·5 (PrPëPI. CtjPsP·PrC-P» B«PμPsP±C-C±PSC-
CtjPμP°PeC†C-C—B»). PμP°C†C-C''PSC,C-PI Cl'P»C-Pr PiPsPiPμCtjPμPrPëC,Pë, C%Ps Cr
CtjP°P·C- CtjP°PiC,PsPiPsPiPs PiPsCtjCrC€PμPSPSCl' P·PsCtjCr P·P°Cl'c,PsCl'CrPIP°PSPSCl'
PiCtjPμPiP°CtjP°c,Cr Cl'P»C-Pr PiCtjPëPiPëPSPëC,Pë C,P° PSPμPiP°PNëPSPs
P·PIPμCtjPSCrC,PëCl'Cl' PrPs P»C-PeP°CtjCl' (PrPëPI. CtjPsP·PrC-P»
B«PμCtjPsC,PëPiPsPeP°P·P°PSPSCl'»).

PkPrPSPsC±P°Cl'PSPμ P·P°Cl'c,PsCl'CrPIP°PSPSCl' P· CtjPëC,PsPSP°PIC-CtjPsPj

PkPrPSPsC±P°Cl'PSPμ P·P°Cl'c,PsCl'CrPIP°PSPSCl' Cl'PëP»PrPμPSP°C,,C-P»Cr C,P°
CtjPëC,PsPSP°PIC-CtjCr PSPμ CtjPμPePsPjPμPSPPrCrC''C,CHCl'Cl' (PrPëPI. CtjPsP·PrC-P»
B«P·P·C''PjPsPrC-Cl' P· C-PSC€PëPjPë P»C-PeP°CtjCl'CHPePëPjPë P·P°Cl'PsP±P°PjPë C,P°
C-PSC€C- PIPëPrPë PIP·P°C''PjPsPrC-PNëB»).

*PkPrPSPsC±P°Cl'PSPμ P·P°Cl'c,PsCl'CrPIP°PSPSCl' P· P±P»PsPeP°c,PsCtjP°PjPë α-
P°PrCtjPμPSPsCtjPμC†PμPiC,PsCtjC-PI*

PμP°C†C-C''PSC,P°Pj, Cl'PeC- P·P°Cl'c,PsCl'PsPICrCtjC,CH P±P»PsPeP°c,PsCtjPë α-
P°PrCtjPμPSPsCtjPμC†PμPiC,PsCtjC-PI, P·P°Cl'c,PsCl'PsPICrPIP°c,Pë Cl'PëP»PrPμPSP°C,,C-P»
Cl'P»C-Pr P· PsP±PμCtjPμP¶PSC-Cl'c,Ctj, PsCl'PeC-P»CHPePë C,P°PeP° PePsPjP±C-PSP°C†C-Cl'
PjPsP¶Pμ PiCtjPëP·PIPμCl'c,Pë PrPs Cl'PëPjPiC,PsPjP°c,PëC±PSPsC— PiC-PiPsC,PμPSP·C-C— Cr
PrPμCl'PePëC... Cl'c...PëP»CHPSPëC... PrPs C†CHPsPiPs PiP°C†C-C''PSC,C-PI.
PÿPëPjPiC,PsPjP°c,PëC±PSP° PiC-PiPsC,PμPSP·C-Cl' P·P°P·PIPëC±P°PNë PIPëPSPëPeP°c''
PiCtjPsC,Cl'PiPsPj 4 PiPsPrPëPS PiC-Cl'P»Cl' P·P°Cl'c,PsCl'CrPIP°PSPSCl'
Cl'PëP»PrPμPSP°C,,C-P»Cr. PÿPsPjCr PrPs PiPsC±P°c,PeCr P·P°Cl'c,PsCl'CrPIP°PSPSCl'
Cl'PëP»PrPμPSP°C,,C-P»Cr PiPμPjPsPrPëPSP°PjC-C±PSC- PiPsPeP°P·PSPëPePë PiP°C†C-C''PSC,P°
PiCtjPë C,PμCtjP°PiC-C— P±P»PsPeP°c,PsCtjP°PjPë α-P°PrCtjPμPSPsCtjPμC†PμPiC,PsCtjC-PI
PjP°CtjC,CH P±CrC,Pë Cl'c,P°P±C-P»CHPSPëPjPë. PÿP»C-Pr CtjPsP·PiP»Cl'PSCrC,Pë
PjPsP¶P»PëPIC-Cl'c,CH P·P°Cl'c,PsCl'CrPIP°PSPSCl' PiPsC±P°c,PePsPiPsC— PrPsP·Pë
Cl'PëP»PrPμPSP°C,,C-P»Cr 25 PjPi (PrPëPI. CtjPsP·PrC-P» B«PÿPiPsCl'c-P±
P·P°Cl'c,PsCl'CrPIP°PSPSCl' C,P° PrPsP·PëB»). PkCtjC-Pj C,PsPiPs, PiPsC,CtjC-P±PSPs
PiCtjPsC-PSC,,PsCtjPjCrPIP°c,Pë PiP°C†C-C''PSC,C-PI, Cl'Pe PrC-Cl'c,Pë Cr CtjP°P·C-
PiPsCl'PIPë Cl'PëPjPiC,PsPjC-PI PsCtjC,PsCl'c,P°c,PëC±PSPsC— PiC-PiPsC,PμPSP·C-C—.

P'PiP»PëPI PSP° PeCtjPsPiPsC,PμC±C'

P''PsCl'P»C-PrP¶PμPSPSCl' B C,CtjPsPjP±PsC†PëC,C-PI B P»CHPrPëPSPë

В ПiСґPsPrPμPjPsPSCГC,СґCрPIP°P»Pө, В C%Ps В in B vitro В CГPөP»PrPμPSP°C,,C-P»B
В PиPsC,PμPSCГC-СґC” P°PSC,PөP°PиCґPμPиP°CГC-PMөPSC- PμC,,PμPөC,Pө PSP°C,СґC-Сґ
PSC-C,СґPsPиCґCрCГPөPrCр. PкPμPjP°C” P¶PsPrPSPsC— C-PSC,,PsCґPjP°CГC-C— C%PsPrPs
P±PμP·PиPμPөPө P·P°CГC,PsCГCрPIP°PSPSCГ CГPөP»PrPμPSP°C,,C-P»Cр PиP°CГC-C”PSC,P°Pj
C-P· PиPsCґCрCөPμPSPSCГPjPө P·PиPsCґC,P°PSPSCГ PөCґPsPIC- P°P±Ps PиPsCГC,СґPsCґ
PиPμPиC,PөC±PSPsCґ PIPөCґP°P·PөPsCґ. PҗP°PөPөPj C±PөPSPsPj, P·P°CГC,PsCГCрPIP°PSPSCГ
CГPөP»PrPμPSP°C,,C-P»Cр PиP°CГC-C”PSC,P°Pj CГC-C”C— PиCґCрPиPө PjPsP¶P»PөPIPμ
P»PөCөPμ PиC-CГP»CГ CґPμC,PμP»СґPSPsC— PsCГC-PSPөPө CГPиC-PIPиC-PrPSPsCөPμPSPSCГ
PөPsCґPөCГC,C- C,P° CґPөP·PөPөC-PI.

P’C,СґP°C,P° CГP»CзC...Cз

P>C-PөP°CґCГPj CГP»C-Pr PиPsCґP°PrPөC,Pө PиP°CГC-C”PSC,P°Pj PиCґPөPиPөPSPөC,Pө
P·P°CГC,PsCГCрPIP°PSPSCГ C-PSPиC-P±C-C,PsCґC-PI PαP”P•5, PIPөP»СґC±P°СґC±Pө
PиCґPμPиP°CґP°C, P’C-PIP°PMөCґP°, C,P° PSPμPиP°PMөPSPs P·PIPμCґPSCрC,PөCГCГ P·P°
PjPμPrPөC±PSPsCґ PrPsPиPsPjPsPиPsCґ Cр CґP°P·C- CґP°PиC,PsPIPсPиPs P·PSPөP¶PμPSPSCГ
C±Pө PIC,СґP°C,Pө CГP»CрC...Cр. P¶CґPs CГC- CГPIPөC%P°, CГPөC- C,P°PөPsP¶
PjPsP¶CрC,Сґ CГCрPиCґPsPIPсPrP¶CрPIP°C,PөCГCГ PrP·PIPсPSPsPj Cр PICрC...P°C... C,P°
P·P°PиP°PjPsCґPsC±PμPSPSCГPj, PиPsPIC-PrPsPjP»CГP»PsCГCГ P· P°CГPsCГC-P°CГC-C”Cґ Cр
C±P°CГC- C-P· P·P°CГC,PsCГCрPIP°PSPSCГPj C-PSPиC-P±C-C,PsCґC-PI PαP”P•5,
PIPөP»СґC±P°СґC±Pө PиCґPμPиP°CґP°C, P’C-PIP°PMөCґP°. P’PөP·PSP°C±PөC,Pө, C±Pө CГC-
CГPIPөC%P° PиCґCГPjPs PиPsPиBґ™ CГP·P°PSC- C-P· P·P°CГC,PsCГCрPIP°PSPSCГPj
C-PSPиC-P±C-C,PsCґC-PI PαP”P•5, C±Pө P· C-PSCөPөPjPө C,,P°PөC,PsCґP°PjPө,
PSPμPjPsP¶P»PөPIPs.

*PҗPrPSPsC±P°CГPSPμ P·P°CГC,PsCГCзPIP°PSPSCГ C-P· PиC-PиPsC,PμPSP·PөPIPSPөPjPө
PиCґPμPиP°CґP°C,P°PjPө*

P’C-PIP°PMөCґP° C±PөPSPөC,Сґ CГPөCГC,PμPjPSCр
CГCрPrPөPSPsCґPsP·CөPөCґCґPиP°P»СґPSCр PrC-Сґ C,P° PjPsP¶Pμ PrP°P»C-
P·PSPөP¶CрPIP°C,Pө P°CґC,PμCґC-P°P»СґPSPөPMө C,PөCГPө Cр PиP°CГC-C”PSC,C-PI, CГPөC-
P·P°CГC,PsCГCрPICрCґC,Сґ PиC-PиPsC,PμPSP·PөPIPSC- P>C-PөP°CґCГCґPөC- P·P°CГPsP±Pө.
Pj PsPөCґPμPjPsPjCр PrPsCГP»C-PrP¶PμPSPSC- P»C-PөP°CґCГCґPөPsC— PIP·P°C”PjPsPrC-C—
PиCґPө PsPrPSPsC±P°CГPSPsPjCр P·P°CГC,PsCГCрPIP°PSPSC- P°PjP»PsPrPөPиC-PSCр (5 PjPi
P°P±Ps 10 PjPi) C,P° PиCґPμPиP°CґP°C,Cр P’C-PIP°PMөCґP° (100 PjPi) PиPμCґPsCґP°P»СґPSPs
CГPиPsCГC,PμCґC-PиP°P»PsCГCГ CГPμCґPμPrPSC” PrPsPrP°C,PөPsPIPμ P·PSPөP¶PμPSPSCГ
CГPөCГC,PsP»C-C±PSPsPиPs C,PөCГPөCр PSP° 8 PjPj CґC,.. CГC,.. PrC-P°CГC,PsP»C-C±PSPsPиPs
Bґ” PSP° 7 PjPj CґC,.. CГC..

*P—P°C...PIPсCґCґPиP°PSPSCГ, C%Ps PиPμCґPμPrP°CґC,СґCГCГ CГC,P°C,PμPIPөPj
CөP»CГC...PsPj*

P—P°CГC,PsCГCрPIP°PSPSCГ PиCґPμPиP°CґP°C,Cр P’C-PIP°PMөCґP° PSPμ P·P°C...PөC%P°C”
PIC-Pr P·P°C...PIPсCґCґPиP°PSCґ, C%Ps PиPμCґPμPrP°CґC,СґCГCГ CГC,P°C,PμPIPөPj
CөP»CГC...PsPj. PҗP»C-Pr PиCґPsC-PSCГC,СґCрPөC,CрPIP°C,Pө PиP°CГC-C”PSC,C-PI C%PsPrPs
PSPμPsP±C...C-PrPSPөC... P·P°PиPsP±C-P¶PSPөC... P·P°CГPsP±C-PI PrP»CГ P·P°C...PөCГC,Cр
PIC-Pr P·P°C...PIPсCґCґPиP°PSCґ, C%Ps PиPμCґPμPrP°CґC,СґCГCГ CГC,P°C,PμPIPөPj
CөP»CГC...PsPj, PIPөP»СґC±P°СґC±Pө PIC-CґCрCГ C-PjCрPSPsPrPμC,,C-C±PөC,Cр
P»CґPrPөPSPө.

PαPμCґC,PөP»СґPSC-CГC,Сґ

PrPsC-ClP»Cl P·P°ClC,PsClCfPIP°PSPSCl PrPsP·Pë 100 PjPi P·PrPsCtjPsPIPëPjPë
PrPsP±CtjPsPIPsP»CHCtClPjPë PSPµ ClPìPsClC,PµCtjC-PiP°P»PsClCl PIPìP»PëPICr PSP°
PjPsCtjC,,PsP»PsPiC-Ct C±Pë CtjCrC...PsPjC-ClC,Ch ClPìPµCtjPjP°C,PsP·PsC—PrC-PI.

P"Ps ClPeP»P°PrCr P'C-PIP°PMcCtjP° PIC...PsPrPëC,Ch P»P°PeC,PsP·P°, ClPeC%Ps Cr P'P°Cl
PIClC,P°PSPsPIP»PµPSPs PSPµPìPµCtjPµPSPsClPëPjC-ClC,Ch PrPµClPePëC... CtCrPeCtjC-PI,
PìCtjPsPePsPSClCfP»CHC,CfPNc,PµClCl P· P»C-PeP°CtjPµPj, PìPµCtjC€ PSC-P¶
PìCtjPëPNcPjP°C,Pë CtPµPNc P»C-PeP°CtjClChPePëPNc P·P°ClC-P±. PkClPìP°CtjC,P°Pj (P• 951)
PI ClPeP»P°PrC- P»C-PeP°CtjClChPePsPiPs P·P°ClPsP±Cr C" PìPsC...C-PrPSPëPj
C,,PµPSC-P»P°P»P°PSC-PSCr, C%Ps ClPIP»ClC" PSPµP±PµP·PìPµPeCr PrP»Cl
C...PIPsCtjPëC... PSP° C,,PµPSC-P»PePµC,PsPSCrCtjC-Ct.

*P—P°ClC,PsClCzPIP°PSPSCl Cz PìPµCtjC-PsPr PIP°PiC-C,PSPsClC,C- P°P±Ps
PiPsPrCzPIP°PSPSCl PiCtjCzPrPrCt.*

PrCtjPµPìP°CtjP°C, PSPµ PìCtjPëP·PSP°C±PµPSPëPNc PrP»Cl P·P°ClC,PsClCfPIP°PSPSCl
P¶C-PSPeP°PjPë.

*P—PrP°C,PSC-ClC,Ch PIPìP»PëPIP°C,Pë PSP° C€PIPëPrPeC-ClC,Ch CtjPµP°PeCtC-C— PìCtjPë
PePµCtjCzPIP°PSPSC- P°PIC,PsC,CtjP°PSClPìPsCtjC,PsPj P°P±Ps C-PSC€PëPjPë
PjPµC...P°PSC-P·PjP°PjPë.*

P"PsClP»C-PrP¶PµPSCt PIPìP»PëPICr PìCtjPµPìP°CtjP°C,Cr PSP° P·PrP°C,PSC-ClC,Ch
PePµCtjCrPIP°C,Pë C,CtjP°PSClPìPsCtjC,PSPëPjPë P·P°ClPsP±P°PjPë C,P° PìCtjP°CtChPIP°C,Pë
P· C-PSC€PëPjPë PjPµC...P°PSC-P·PjP°PjPë PSPµ PìCtjPsPIPsPrPëP»PsClCl.

PkClPeC-P»CHPePë Cr PìCtjPsCtPµClC- PeP»C-PSC-C±PSPëC... PrPsClP»C-PrP¶PµPSCt
P·P°ClC,PsClCfPIP°PSPSCl ClPëP»PrPµPSP°C,,C-P»Cr PìPsPIC-PrPsPjP»ClP»PsClCl PìCtjPs
PIPëPìP°PrPePë P·P°PìP°PjPsCtjPsC±PµPSPSCl C,P° PìPsCtjCrC€PµPSPSCl P· P±PsPeCr
PsCtjPìP°PSC-PI P·PsCtjCr, PìPµCtjPµPr C,PëPj, ClPe ClC-PrP°C,Pë P·P° PePµCtjPjPs
C,CtjP°PSClPìPsCtjC,PSPsPiPs P·P°ClPsP±Cr P°P±Ps PìCtjP°CtChPIP°C,Pë P·
PjPµC...P°PSC-P·PjP°PjPë, PìP°CtC-C" PSC,P°Pj PSPµPsP±C...C-PrPSPs P·Bt™ ClClCfPIP°C,Pë,
ClPePsCt C" C—C...PSCl C-PSPPrPëPIC-PrCrP°P»CHPSP° CtjPµP°PeCtC-Cl PSP°
P·P°ClC,PsClCfPIP°PSPSCl PìCtjPµPìP°CtjP°C,Cr.

PŸPìPsClC-P± P·P°ClC,PsClCzPIP°PSPSCl C,P° PrPsP·Pë.

PrCtjPµPìP°CtjP°C, P·P°ClC,PsClPsPICrCtC,Ch PìPµCtjPsCtjP°P»CHPSPs.

PŸP°P±P»PµC,PeCr ClP»C-Pr PìPsPIPSC-ClC,Ct CtjPsP·P¶CrPIP°C,Pë C,P°
PìCtjPsPePsPIC,PSCrC,Pë.

P"PsCtjPsClP»C-

PrCtjPë PSPµPsP±C...C-PrPSPsClC,C- PìCtjPµPìP°CtjP°C, P'C-PIP°PMcCtjP° ClP»C-Pr

P·P°CÍC,PsCÍPsPICfPIP°C,Pë P·P° 1 PiPsPrPëPSCf PrPs CÍPµPeCÍCfP°P»CHPSPSc—
P°PeC,PëPIPSPScÍC,C-. P PµPePsPjPµPSPPrPsPIP°PSP° PrPsP·P° CÍC,P°PSPsPIPëC,CH 50 PjPi.
PhCÍPeC-P»CHPePë PiCtjPë PsPrPSPScfP°CÍPSPsPjCf P·P°CÍC,PsCÍCfPIP°PSPSC-
PiCtjPµPiP°CtjP°C,Cf P·C—P¶PµCt PIC-PrP±CfPIP°C”C,CHCÍCt P·P°C,CtjPëPjPeP°
P°P±CÍPsCtjP±CtC-C— C,P° CÍPiPsPIC-P»CHPSCtC”C,CHCÍCt PµC,,PµPeC,
PiCtjPµPiP°CtjP°C,Cf (PrPëPI. CtjPsP·PrC-P» B«PαP°CtjPjP°PePsPeC-PSPµC,PëPeP°B»),
CtjPµPePsPjPµPSPPrPsPIP°PSPs P·P°CÍC,PsCÍPsPICfPIP°C,Pë PiCtjPµPiP°CtjP°C, PSP°C,C%µPµ.

P—P°P»PµP¶PSPs PIC-Pr PµC,,PµPeC,PëPIPSPScÍC,C- C,P° PiPµCtjPµPSPScÍPëPjPsCÍC,C-
PrPsP·Cf PiCtjPµPiP°CtjP°C,Cf PjPsP¶PSP° P·P±C-P»CHCεPëC,Pë PrPs 100 PjPi P°P±Ps
P·PjPµPSCεPëC,Pë PrPs 25 PjPi*. PµP°PeCÍPëPjP°P»CHPSP° CtjPµPePsPjPµPSPPrPsPIP°PSP°
PrPsP·P° CÍC,P°PSPsPIPëC,CH 100 PjPi.

PµP°PeCÍPëPjP°P»CHPSP° CtjPµPePsPjPµPSPPrPsPIP°PSP° CfP°CÍC,PsC,P°
P·P°CÍC,PsCÍCfPIP°PSPSCtj PiCtjPµPiP°CtjP°C,Cf CÍC,P°PSPsPIPëC,CH 1 CtjP°P· PSP°
PrPsP±Cf.

PµP°CtC-C”PSC,Pë P»C-C,PSCHPsPiPs PIC-PeCé

PkPµPsP±C...C-PrPSC-CÍC,CH Cf PePsCtjPµPeCtC-C— PrPsP·Pë PiP°CtC-C”PSC,P°Pj
P»C-C,PSCHPsPiPs PIC-PeCf (PIC-PePsPj PIC-Pr 65 CtjPsPeC-PI) PIC-PrCÍCfC,PSCtj.

PµP°CtC-C”PSC,Pë P·PSPëCtjPePsPIPsCtj PSPµPrPsCÍC,P°C,PSC-CÍC,CH

P”P»Ctj PiP°CtC-C”PSC,C-PI C-P·PSPëCtjPePsPIPsCtj PSPµPrPsCÍC,P°C,PSC-CÍC,CH
P»PµPiPePsPiPs C,P° PiPsPjC-CtjPSPsPiPs CÍC,CfPiPµPSCtj C,CtjP¶PePsCÍC,C-
(PeP»C-CtjPµPSCÍ PeCtjPµP°C,PëPSC-PSCf 30Btj”80 PjP»/C...PI) CtjPµPePsPjPµPSPPrPsPIP°PSP°
PrPsP·P° PiCtjPµPiP°CtjP°C,Cf C” C,P°PePsCtj P¶, CtjPe PSP°PIPµPrPµPSPs PIPëC%µPµ Cf
CtjPsP·PrC-P»C- B«P”PsCtjPsCÍP»C-B».

PhCÍPeC-P»CHPePë Cf PiP°CtC-C”PSC,C-PI P·PSPëCtjPePsPIPsCtj
PSPµPrPsCÍC,P°C,PSC-CÍC,CH C,CtjP¶PePsPiPs CÍC,CfPiPµPSCtj (PeP»C-CtjPµPSCÍ
PeCtjPµP°C,PëPSC-PSCf < 30 PjP»/C...PI) PeP»C-CtjPµPSCÍ CÍPëP»PrPµPSP°C,,C-P»Cf
P·PSPëP¶PµPSPëPµ, CÍP»C-Pr CtjPsP·PiP»CtjPSCfC,Pë PjPsP¶P»PëPIC-CÍC,CH
P·P°CÍC,PsCÍCfPIP°PSPSCtj PrPsP·Pë B B B B B 25 PjPi*. P—P°P»PµP¶PSPs PIC-Pr
PµC,,PµPeC,PëPIPSPScÍC,C- C,P° PiPµCtjPµPSPScÍPëPjPsCÍC,C- PiCtjPµPiP°CtjP°C,Cf PiCtjPë
PSPµPsP±C...C-PrPSPScÍC,C- PrPsP·Cf PjPsP¶PSP° PiPsCÍC,CfPiPsPIPs P·P±C-P»CHCεPëC,Pë
PrPs 50 PjPi C,P° PrPs 100 PjPi.

PµP°CtC-C”PSC,Pë P·PiPµCfC-PSPePsPIPsCtj PSPµPrPsCÍC,P°C,PSC-CÍC,CH

PhCÍPeC-P»CHPePë Cf PiP°CtC-C”PSC,C-PI C-P·PiPµCfC-PSPePsPIPsCtj
PSPµPrPsCÍC,P°C,PSC-CÍC,CH (PSP°PiCtjPëPeP»P°Pr, C-P·CtjPëCtjPsP·PsPj) PeP»C-CtjPµPSCÍ
CÍPëP»PrPµPSP°C,,C-P»Cf P·PSPëP¶PµPSPëPµ, CÍP»C-Pr CtjPsP·PiP»CtjPSCfC,Pë
PjPsP¶P»PëPIC-CÍC,CH P·P°CÍC,PsCÍCfPIP°PSPSCtj PrPsP·Pë 25 PjPi*. P—P°P»PµP¶PSPs PIC-Pr
PµC,,PµPeC,PëPIPSPScÍC,C- C,P° PiPµCtjPµPSPScÍPëPjPsCÍC,C- PiCtjPµPiP°CtjP°C,Cf PiCtjPë
PSPµPsP±C...C-PrPSPScÍC,C- PrPsP·Cf PjPsP¶PSP° PiPsCÍC,CfPiPsPIPs P·P±C-P»CHCεPëC,Pë
PrPs 50 PjPi C,P° PrPs 100 PjPi.

*PµP°CtC-C”PSC,Pë, CtjPeC- P·P°CÍC,PsCÍPsPICéCtjC,CH C-PSCεC- P»C-PeP°CtjCÍCHPeC-
P·P°CÍPsP±Pë*

PiPeC%µPs PiP°CtC-C”PSC,Pë PsPrPSPScfP°CÍPSPs P·P°CÍC,PsCÍPsPICfCtjC,CH

C-PSiC-P±C-C,PsCḡPë CYP3A4 (PrPëPI. CḡPsP·PrC-P» B«P'P·P°C''PjPsPrC-Cḡ P· C-PSC€PëPjPë P»C-PeP°CḡCḡCḡPePëPjPë P·P°CḡPsP±P°PjPë C,P° C-PSC€C- PIPëPrPë PIP·P°C''PjPsPrC-PNëB»), CḡP»C-Pr CḡPsP·PiP»CḡPSCḡC,Pë PjPsPḡP»PëPIC-CḡC,Ch P·P°CḡC,PsCḡCḡPIP°PSPSCḡ PİPsCḡP°C,PePsPIPsC— PrPsP·Pë 25 PjPi* (P·P° PIPëPSCḡC,PePsPj CḡPëC,PsPSP°PIC-CḡCḡ, P·P°CḡC,PsCḡCḡPIP°PSPSCḡ CḡPePsPiPs PsPrPSPsCḡP°CḡPSPs P·C- CḡPëP»PrPµPSP°C,,C-P»PsPj PSPµ CḡPµPePsPjPµPSPḡCḡC,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°CḡC-C''PSC,C-PI, CḡPeC- P·P°CḡC,PsCḡPsPICḡCḡC,Ch P±P»PsPeP°C,PsCḡPë α-P°PrCḡPµPSPsCḡPµCḡPµPiC,PsCḡC-PI, PrPs PİPsCḡP°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. PÿP°PePsPḡ PİPsC,CḡC-P±PSPs CḡPsP·PiP»CḡPSCḡC,Pë PjPsPḡP»PëPIC-CḡC,Ch P·P°CḡC,PsCḡCḡPIP°PSPSCḡ PİPsCḡP°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°CḡCḡCḡPePëPjPë 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 PİCḡPsCḡPµCḡCḡ- PeP»C-PSC-CḡPSPëC... PrPsCḡP»C-PrPḡPµPSCḡ P·P° CḡCḡP°CḡC,Ch PrPsCḡPsCḡP»PëC... PrPsP±CḡPsPIPsP»ChCḡC-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-CḡPSC- CḡPµP°PeCḡC-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 PSPëPḡCḡPëC... PrPsP·P°C..., P°P»Pµ PIPëPSPëPeP°P»Pë CḡP°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 CḡPİCḡPëCḡPëPSCḡP»Ps P·CḡPsCḡC,P°PSPSCḡ PeC-P»ChPePsCḡC,C- PIPëPİP°PrPeC-PI CḡPsP·PIPëC,PeCḡ PİPsP±C-CḡPSPëC... CḡPµP°PeCḡC-PNë (PİPsP»PsPIPsPİPs P±PsP»Ch, PİCḡPëPİP»PëPIC-PI PeCḡPsPIC-, P·P°PİP°PjPsCḡPsCḡPµPSPSCḡ, 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- PIPḡP°CḡC,ChCḡCḡ PrPs P·PIPëCḡP°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 CḡPµCḡPµP· PIPëCḡPsPePëPNë CḡC,CḡPİC-PSCḡ P·PİBḡ™ 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°CḡC-C— CḡPëP»PrPµPSP°C,,C-P»Cḡ C-P· CḡPµCḡPµCḡ.

PuPsP±C-C±PSC- CḡPμP°PeC†C-C—.

PḡCḡPsC,,C-P»CH P±PμP·PīPμPePē CḡPēP»PrPμPSP°C,,C-P»Cḡ P±P°P·CḡC”C,CHCḡCḡ PSP° PrP°PSPēC..., PsC,CḡPēPjP°PSPēC... C-P· 74 PīPsPrPIC-PN°PSPēC... CḡP»C-PīPēC... PīP»P°C†PμP±Ps-PePsPSC,CḡPsP»CHPsPIP°PSPēC... PeP»C-PSC-C†PSPēC... PrPsCḡP»C-PrPḡPμPSCḡ (9570 PīP°C†C-C”PSC,C-PI). PḡP°PN°C†P°CḡC,C-C€Pμ PīPsPIC-PrPsPjP»CḡP»PsCḡCḡ PīCḡPs C,P°PeC- PīPsP±C-C†PSC- CḡPμP°PeC†C-C—, CḡPe PīPsP»PsPIPSPēPN° P±C-P»CH, PīCḡPēPīP»PēPIPē PeCḡPsPIC-, PrPēCḡPīPμPīCḡC-Cḡ, P·P°PeP»P°PrPμPSC-CḡC,CH PSPsCḡP°, P±C-P»CH Cḡ CḡPīPēPSC-, P·P°PīP°PjPsCḡPsC†PμPSPSCḡ, PSCḡPrPsC,P°, PīCḡPēPīP»PēPIPē PḡP°CḡCḡ, PīPsCḡCḡC€PμPSPSCḡ P·PsCḡCḡ, C†C-P°PSPsPīCḡC-Cḡ C,P° P·P°C,CHPjP°CḡPμPSPSCḡ P·PsCḡCḡ. P†PSC,,PsCḡPjP°C†C-Cḡ C°PsPrPs PīPsP±C-C†PSPēC... CḡPμP°PeC†C-PN° Cḡ CḡP°PjPeP°C... PīPsCḡC,PjP°CḡPePμC,PēPSPiPsPIPsiPs CḡPīPsCḡC,PμCḡPμPḡPμPSPSCḡ P±CḡP»P° P·C-P±CḡP°PSP° PīCḡPsC,CḡPīPsPj PīPμCḡC-PsPrCḡ C,CḡPēPIP°P»C-CḡC,CH P±C-P»CHC€Pμ PSC-Pḡ 10 CḡPsPeC-PI. PḡCḡPeC-P»CHPePē P·P°CḡPIPSPēPePsPIC- PīPsPIC-PrPsPjP»CḡP»PsCḡCḡ PSPμ PīCḡPs PICḡC- PīPsP±C-C†PSC- CḡPμP°PeC†C-C— C,P° PSPμ PICḡC- PīPsP±C-C†PSC- CḡPμP°PeC†C-C— P±CḡP»Pē PIPeP»CḡC†PμPSC- PI P±P°P·Cḡ PrP°PSPēC... P· P±PμP·PīPμPePē, C†P°CḡC,PsC,P° C,P°PePēC... CḡPμP°PeC†C-PN° PSPμ PjPsPḡPμ P±CḡC,Pē PrPsCḡC,PsPIC-CḡPSPs PIPēP·PSP°C†PμPSP°.

PjCḡC- PeP»C-PSC-C†PSPs P·PSP°C†CḡC°C- PīPsP±C-C†PSC- CḡPμP°PeC†C-C—, C°Ps CḡPīPsCḡC,PμCḡC-PiP°P»PēCḡCḡ PīC-Pr C†P°Cḡ PeP»C-PSC-C†PSPēC... PrPsCḡP»C-PrPḡPμPSCḡ C†P°CḡC,C-C€Pμ, PSC-Pḡ PīCḡPē P·P°CḡC,PsCḡCḡPIP°PSPSC- PīP»P°C†PμP±Ps, PSP°PIPμPrPμPSPs PSPēPḡC†Pμ P·P° CḡPēCḡC,PμPjP°PjPē PsCḡPīP°PSC-PI C,P° C†P°CḡC,PsC,PsCḡ: PrCḡPḡPμ C†P°CḡC,Ps (≥1/10), C†P°CḡC,Ps (≥1/100 Bḡ” <1/10), PSPμC†P°CḡC,Ps (≥1/1000 Bḡ” <1/100), CḡC-PrPePs (≥1/10000 Bḡ” <1/1000), PrCḡPḡPμ CḡC-PrPePs (<1/10000), C†P°CḡC,PsC,P° PSPμPIC-PrPsPjP° (PSPμ PjPsPḡPμ P±CḡC,Pē PICḡC,P°PSPsPIP»PμPSP° PIPēC...PsPrCḡC†Pē P· PSP°CḡPIPSPēC... PrP°PSPēC...). PḡP°CḡC,PsC,P° PeP»C-PSC-C†PSPs P·PSP°C†CḡC°PēC... PīPsP±C-C†PSPēC... CḡPμP°PeC†C-PN°, PīCḡPs CḡPeC- PīPsPIC-PrPsPjP»CḡP»PsCḡCḡ Cḡ CḡP°PjPeP°C... PīPsCḡC,PjP°CḡPePμC,PēPSPiPsPIPēC... CḡPīPsCḡC,PμCḡPμPḡPμPSCḡ, PIPēP·PSP°C†PμPSP° CḡPe PSPμPIC-PrPsPjP°. Pj PjPμPḡP°C... PePsPḡPSPsC— PīCḡCḡPīPē P·P° C†P°CḡC,PsC,PsCḡ PīPsP±C-C†PSC- CḡPμP°PeC†C-C— PSP°PIPμPrPμPSC- Cḡ PīPsCḡCḡPrPeCḡ P·PjPμPSC€PμPSPSCḡ C—C... CḡPμCḡPN°PsP·PSPsCḡC,C-.

P†PSC,,PμPeC†C-PN°PSC- C,P° C-PSPIP°P·PēPIPSC- P·P°C...PIPSCḡCḡPIP°PSPSCḡ

PḡPμC†P°CḡC,Ps: CḡPēPSC-C,.

P— P±PsPeCḡ C-PjCḡPSPSPsC— CḡPēCḡC,PμPjPē

PḡPμC†P°CḡC,Ps: CḡPμP°PeC†C-C— PīC-PrPIPēC°PμPSPsC— C†CḡC,P»PēPIPSCḡC,C-.

P— P±PsPeCḡ PSPμCḡPIPsiPsC— CḡPēCḡC,PμPjPē

P”CḡPḡPμ C†P°CḡC,Ps: PīPsP»PsPIPSPēPN° P±C-P»CH.

PḡP°CḡC,Ps: P·P°PīP°PjPsCḡPsC†PμPSPSCḡ.

PḡPμC†P°CḡC,Ps: CḡPsPSP»PēPIC-CḡC,CH, PīC-PīPμCḡC,PμP·C-Cḡ.

P C-PrPePs: C-PSCÍCíP»CHC,, CÍPëPSPePsPíPµ, C,CHP°PSP·PëC,PsCḡPSP° C-CëPµPjC-C±PSP° P°C,P°PeP°, CÍCíPrPsPjPë, CḡPµC†PëPrPëPIPë CÍCíPrPsPj.

P— P±PsPeCá PsCḡPiP°PSC-PI P·PsCḡCá

PŞP°CÍC,Ps: PíPsCḡCíCëPµPSPSCÍ CÍPíCḡPëPñPSCÍC,C,CÍ PëPsP»CHPsCḡCí*, CḡPsP·P»P°PrPë P·PsCḡCí, P·P°C,CHPjP°CḡPµPSPSCÍ P·PsCḡCí.

PќPµC±P°CÍC,Ps: CḡPsP·P»P°PrPë CÍP»CHPsP·PsPíPëPrC-P»PµPSPSCÍ**, P±C-P»CH PI PsC±P°C..., C,,PsC,PsC,,PsP±C-CÍ, C,,PsC,PsPíCÍC-CÍ, PiC-PíPµCḡPµPjC-CÍ PsC±PµPñ, CÍCÍPëCḡP°PIC-CÍC,CH P·PsCḡCí, PePsPSḡḡ™ CḡPSPeC,PëPIC-C,.

P C-PrPePs: PSPµP°CḡC,PµCḡC-P°P»CHPSP° PíPµCḡPµPrPSCÍ C-CëPµPjC-C±PSP° PSPµPñCḡPsPíP°C,C-CÍ P·PsCḡPsPíPíPSPµCḡPIP°, PsPeP»CHP·C-CÍ CÍCíPrPëPS CÍC-C,PeC-PIPePë, CḡPµC,PëPSP°P»CHPSPëPñ PeCḡPsPíPíPëP»PëPI, P°CḡC,PµCḡC-PsCÍPëP»PµCḡPsC,PëC±PSP° CḡPµC,PëPSPsPíP°C,C-CÍ, PíPsCḡCíCëPµPSPSCÍ P· P±PsPeCí CÍC-C,PeC-PIPePë, PiP»P°CíPePsPjP°, PrPµC,,PµPeC,Pë PíPsP»CÍ P·PsCḡCí, PrPëPíP»PsPíC-CÍ, P·PSPëP¶PµPSPSCÍ PiPsCÍC,CḡPsC,Pë P·PsCḡCí, PjC-PsPíC-CÍ, P°CÍC,PµPSPsPíC-CÍ, PíP»P°PIP°CḡC±C- PíPsPjCíC,PSC-PSPSCÍ CÍPëP»PëCÍC,PsPíPs C,C-P»P°, PíPsCḡCíCëPµPSPSCÍ P· P±PsPeCí CḡP°PñPrCíP¶PSPsC— PsP±PsP»PsPSPePë, PjC-PrCḡC-P°P, PíPsCÍPIP° CÍCÍCḡC±PëC... PeCḡCíPiC-PI PSP°PIPePsP»Ps PrP¶PµCḡPµP»P° CÍPIC-C,P»P° (PiP°P»Ps) Cí PíPsP»C- P·PsCḡCí, PSP°P±CḡCÍPe PsC±PµPñ, PíCḡPëPíCíC...P»C-CÍC,CH PsC±PµPñ, PíPsCḡCíCëPµPSPSCÍ P· P±PsPeCí PsC±PµPñ, PiC-PíPµCḡPµPjC-CÍ PePsPSḡḡ™ CḡPSPeC,PëPIPë, PíPsPrCḡP°P·PSPµPSPSCÍ PsC±PµPñ, P°PSPsPjP°P»CHPSC- PIC-PrC±CíC,C,CÍ PI PsC±P°C..., PSP°P±CḡCÍPe PíPsPIC-Pe, P·PSPµP±P°CḡPIP»PµPSPSCÍ CÍPëP»PµCḡPë.

P— P±PsPeCá PsCḡPiP°PSC-PI CÍP»CáC...Cá C,P° PIPµCÍC,PëP±CáP»CÍCḡPSPsPíPs P°PíP°CḡP°C,Cá

PќPµC±P°CÍC,Ps: P·P°PíP°PjPsCḡPsC±PµPSPSCÍ, PIPµCḡC,PëPíPs, PrP·PIC-PS Cí PICíC...P°C....

P C-PrPePs: PiP»CíC...PsC,P°.

P— P±PsPeCá CÍPµCḡC†PµPIPí-CÍCáPrPëPSPSPsC— CÍPëCÍC,PµPjPë

PŞP°CÍC,Ps: PíCḡPëPíP»PëPIPë PeCḡPsPIC- PrPs PsP±P»PëC±C±CÍ, PíCḡPëPíP»PëPIPë P¶P°CḡCí.

PќPµC±P°CÍC,Ps: C,P°C...C-PeP°CḡPrC-CÍ, PíPsCÍPëP»PµPSPµ CÍPµCḡC†PµP±PëC,C,CÍ, P°CḡC,PµCḡC-P°P»CHPSP° PiC-PíPµCḡC,PµPSP·C-CÍ, P°CḡC,PµCḡC-P°P»CHPSP° PiC-PíPsC,PµPSP·C-CÍ.

P C-PrPePs: CḡP°PíC,PsPIP° CÍPµCḡC†PµPIP° CÍPjPµCḡC,CH, C-PSC,,P°CḡPeC, PjC-PsPeP°CḡPrP°, CëP»CíPSPsC±PePsPIP° P°CḡPëC,PjC-CÍ, C,,C-P±CḡPëP»CÍC†C-CÍ PíPµCḡPµPrCÍPµCḡPrCH, PSPµCÍC,P°P±C-P»CHPSP° CÍC,PµPSPsPeP°CḡPrC-CÍ.

P— P±PsPeCá PrPëC...P°P»CHPSPsC— CÍPëCÍC,PµPjPë, PiCḡCáPrPSPsC— PeP»C-C,PePë C,P° CÍPµCḡPµPrPsCÍC,C-PSPSCÍ

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

PќPµC±P°CÍC,Ps: PSPsCÍPsPIP° PeCḡPsPIPíC,PµC±P°, P·P°PeP»P°PrPµPSC-CÍC,CH

PiC7P8PrP°C,PePsPIP8C... PiP°P·C7C... PSPsC7P°.

P C-PrPePs: PIC-PrC7C7C,C,C7 C7C,P8C7PSPµPSPSC7 C7 PiPsC7P»C-, PSP°P±C7C7Pe C7P»P8P·PsPIPSC— PsP±PsP»PsPSPeP8 PSPsC7P°, C7C7C...C-C7C,C7 C7 C7 PSPsC7C-.

P— P±PsPeC7 C7P»C7PSPePsPIP8-PeP8C7PePsPIP8PiPs C,C7P°PeC,C7

P8P°C7C,Ps: PSC7PrPsC,P°, PrP8C7PiPµPiC7C-C7.

PkPµC7P°C7C,Ps: PiP°C7C,C7PsPµP·PsC,,P°PiPµP°P»C7PSP° C7PµC,,P»C7PeC7PSP° C...PIP8C7PsP±P°, P±P»C7PIP°PSPSC7, P±C-P»C7 C7 PiPµC7C...PSC-PM8 C7P°C7C,P8PSC-P7P8PIP8C,P°, C7C7C...C-C7C,C7 C7 C7PsC,C-.

P C-PrPePs: PiC-PiPµC7C,PµP·C-C7 C7PsC,PsPIP8C— PiPsC7PsP7PSP8PSP8.

P— P±PsPeC7 C7PeC-C7P8 C,P° PiC-PrC7PeC-C7PSP8C— PeP»C-C,PePsPIP8PSP8

PkPµC7P°C7C,Ps: PIP8C7P8PiP°PSPSC7.

P C-PrPePs: C7P8PSP7C7PsPj P7C,C-PIPµPSC7P°P7P”P7P7PsPSC7PsPSP°, C,PsPeC7P8C7PSP8PM8 PµPiC-PrPµC7PjP°P»C7PSP8PM8 PSPµPeC7PsP»C-P·.

P— P±PsPeC7 PsPiPsC7PSP8-C7C7C...PsPIP8PiPs P°PiP°C7P°C,C7 C,P° C7PiPsP»C7C7PSP8C— C,PeP°PSP8PSP8

PkPµC7P°C7C,Ps: PjC-P°P»PiC-C7, P±C-P»C7 C7 PeC-PSC7C-PIP8P°C....

P— P±PsPeC7 C7PµC7PsPIP8PrC-P»C7PSP8C— C7P8C7C,PµPjP8

PkPµC7P°C7C,Ps: PiPµPjP°C,C7C7C-C7.

P— P±PsPeC7 C7PµPiC7PsPrC7PeC,P8PIP8PsC— C7P8C7C,PµPjP8 C,P° PjPsP»PsC7PSP8C... P·P°P»PsP·

P C-PrPePs: PeC7PsPIP8C,PµC7P° P·C- C7C,P°C,PµPIP8PiPs C7P»PµPSP°, PiC7C-P°PiC-P·Pj, PiPµPjP°C,PsC7PiPµC7PjC-C7, PiPsPrPsPIP7PµPSP° PµC7PµPeC7C-C7.

P— P°PiP°P»C7PSC- C7PsP·P»P°PrP8 C,P° C7PµP°PeC7C-C— C7 PjC-C7C7C- PIPiPµPrPµPSPSC7

PkPµC7P°C7C,Ps: P±C-P»C7 C7 PiC7C7PrC7C..., PiC-PrPIP8C%PµPSP° PIC,PsPjP»C7PIP°PSC-C7C,C7, PIC-PrC7C7C,C,C7 P7P°C7C7.

P C-PrPePs: PiPsPrC7P°P·PSPµPSPSC7.

P»P°P±PsC7P°C,PsC7PSC- C,PµC7C,P8

PkPµC7P°C7C,Ps: PiC-PrPIP8C%PµPSP° C7P°C7C,PsC,P° C7PµC7C7PµPIP8C... C7PePsC7PsC7PµPSC7.

* B PµPsC7C7C7PµPSPSC7 C7PiC7P8PM8PSC7C,C,C7 PePsP»C7PsC7C7: C...P»PsC7PsPiC7C-C7, C...C7PsPjP°C,PsPiC7C-C7, C7C-P°PSP8PiC7C-C7, PµC7P8C,C7PsPiC7C-C7, PeC7P°PSC,PsPiC7C-C7.

** PµPsC7C7C7PµPSPSC7 C7P»C7PsP·PsPIP8PrC-P»PµPSPSC7: C7C7C...C-C7C,C7 PI

PsC±P°C..., PīPsCḡCfC€PμPSPSCḡ CḡP»CHPsP·PsPIPëPrC-P»PμPSPSCḡ C,P°
PīC-PrPIPëC%PμPSPSCḡ CḡP»CHPsP·PsPIPëPrC-P»PμPSPSCḡ.

PkPëPḡC±PμP·P°P·PSP°C±PμPSC- CḡPIPëC%P° CḡPīPsCḡC,PμCḡC-PiP°P»PëCḡCḡ Cḡ < 2 %
PīP°CḡC-C”PSC,C-PI Cḡ PīCḡPsC±PμCḡC- PePsPSC,CḡPsP»CHPsPIP°PSPëC...
PeP»C-PSC-C±PSPëC... PrPsCḡP»C-PrPḡPμPSCḡ; PīCḡPëC±PëPSPSPëPNḡ
PIP·P°C”PjPsP·PIBḡ™ CḡP·PsPe PSPμ PIPëP·PSP°C±PμPSPëPNḡ. PμPsPIC-PrPsPjP»PμPSPSCḡ
PIPëP»CHC±P°P»Pë CḡPIPëC%P°, C%Ps PjP°P»Pë PIC-CḡPsPiC-PrPSPëPNḡ P·PIBḡ™ CḡP·PsPe
C-P·P°CḡC,PSCḡCḡPIP°PSPSCḡPj PīCḡPμPīP°CḡP°C,Cḡ. PīPIPëC%P°, CḡPeC- PSPμ P±CḡP»Pë
P·P°P·PSP°C±PμPSC-, P±CḡP»Pë P»PμPiPePëPjPë C,P° PīPsPIC-PrPsPjP»PμPSPSCḡ P±CḡP»Pë
PrCḡPḡPμ PSPμC,PSC±PSPëPjPë, C%PsP± PjP°C,Pë P·PSP°C±PμPSPSCḡ.

P—P°PiP°P»CHPSC-. PkP°P±CḡCḡPe PsP±P»PëC±CḡCḡ, CḡPμP°PeCḡC-C—
C,,PsC,PSC±CḡC,P»PëPIPSCḡC-, C€PsPe, P°CḡC,PμPSC-Cḡ, P±C-P»CH, CḡP°PiC,PsPIPμ
PīP°PrC-PSPSCḡ, P±C-P»CH Cḡ PḡPëPīPsC,C-, CḡP°PīC,PsPIPμ PīPsC€PePsPrPḡPμPSPSCḡ.

P— P±PsPeCḡ CḡPμCḡCḡPμPIPSCḡCḡPrPëPSPSPSC— CḡPëCḡC,PμPjPë:
CḡC,PμPSPPsPeP°CḡPrC-Cḡ, PḡV-P±P»PsPeP°PrP°, PjC-PiCḡPμPSCḡ, PīPsCḡC,CḡCḡP°P»CHPSP°
PiC-PīPsC,PμPSP·C-Cḡ, C-C€PμPjC-Cḡ PjC-PsPeP°CḡPrP°, C,CḡPsPjP±PsP· CḡCḡPrPëPS
PīPsP»PsPIPSPsPiPs PjPsP·PeCḡ, CḡP°PīC,PsPIP° P·CḡPīPëPSPeP° CḡPμCḡCḡCḡ,
PIC-PrC...PëP»PμPSPSCḡ PīPsPeP°P·PSPëPeC-PI P·PḡP”, PeP°CḡPrC-PsPjC-PsPīP°C,C-Cḡ.

P— P±PsPeCḡ C€P»CḡPSPePsPIPs-PePëC€PePsPIPsPiPs C,CḡP°PeC,Cḡ: PiP»PsCḡPëC,, PePsP»C-C,,
PrPëCḡC,,P°PiC-Cḡ, PiP°CḡC,CḡPëC,, PiP°CḡC,CḡPsPμPSC,PμCḡPëC,, PμP·PsC,,P°PiC-C,,
CḡC,PSPjP°C,PëC,, PīPsCḡCḡC€PμPSPSCḡ CḡPμP·CḡP»CHC,P°C,C-PI PīPμCḡC-PSPePsPIPëC...
PīCḡPsP±, CḡPμPeC,P°P»CHPSP° PeCḡPsPIPsC,PμCḡP°, PiC-PSPiC-PIC-C,.

P— P±PsPeCḡ CḡPëCḡC,PμPjPë PeCḡPsPIC- C,P° P»C-PjC,,P°C,PëC±PSPSC— CḡPëCḡC,PμPjPë:
P°PSPμPjC-Cḡ, P»PμPNḡPePsPīPμPSC-Cḡ.

P PsP·P»P°PrPë PjPμC,P°P±PsP»C-P·PjCḡ C,P° C...P°CḡCḡCḡPIP°PSPSCḡ: CḡPīCḡP°PiP°,
PSP°P±CḡCḡPe, PīPsPrP°PiCḡP°, PSPμCḡC,P°P±C-P»CHPSPëPNḡ PrC-P°P±PμC,,
PiC-PīPμCḡPīP»C-PePμPjC-Cḡ, PīPμCḡPëC,,PμCḡPëC±PSC- PSP°P±CḡCḡPePë,
PiC-PīPμCḡCḡCḡPëPePμPjC-Cḡ, PiC-PīPsPiP»C-PePμPjC-Cḡ, PiC-PīPμCḡPSP°C,CḡC-C”PjC-Cḡ.

P— P±PsPeCḡ CḡPePμP»PμC,PSPs-Pjḡ™ CḡP·PsPIPsC— CḡPëCḡC,PμPjPë: P°CḡC,CḡPëC,,
P°CḡC,CḡPsP·, CḡPsP·CḡPëPI CḡCḡC...PsPḡPëP»P»Cḡ, C,PμPSPsCḡPëPSPsPIC-C,, P±C-P»CH Cḡ
PeC-CḡC,PeP°C..., PjC-P°CḡC,PμPSC-Cḡ, CḡPëPSPsPIC-C,.

P— P±PsPeCḡ PSPμCḡPIPsPIPsC— CḡPëCḡC,PμPjPë: P°C,P°PeCḡC-Cḡ, PSPμPICḡP°P»PiC-Cḡ,
PSPμPNḡCḡPsPīP°C,C-Cḡ, PīP°CḡPμCḡC,PμP·C-Cḡ, C,CḡPμPjPsCḡ, PIPμCḡC,PëPīPs,
PrPμPiCḡPμCḡC-Cḡ, P±PμP·CḡPsPSPSCḡ, P°PSPsPjP°P»CHPSC- CḡPSPsPIPëPrC-PSPSCḡ,
P·PSPëPḡPμPSPSCḡ CḡPμC,,P»PμPeCḡC-PI.

P— P±PsPeCḡ PrPëC...P°P»CHPSPsC— CḡPëCḡC,PμPjPë: P°CḡC,PjP°, PrPëCḡPīPSPsPμ,
P»P°CḡPëPSPiC-C,, C,,P°CḡPëPSPiC-C,, CḡPëPSCḡCḡPëC,, P±CḡPsPSC...C-C,, PīPsCḡPëP»PμPSPμ
CḡP»PëPSPsPIPëPrC-P»PμPSPSCḡ, PīPsCḡPëP»PμPSPSCḡ PeP°C€P»Cḡ.

P PsP·P»P°PrPë P· P±PsPeCḡ C€PeC-CḡPë: PeCḡPsPīPëPIBḡ™ CḡPSPeP°, PiPμCḡPīPμCḡ,
CḡPIPμCḡP±C-Pḡ, PīC-PrPIPëC%PμPSP° PīC-C,P»PëPIC-CḡC,CH, PIPëCḡP°P·PePë C€PeC-CḡPë,
PePsPSC,P°PeC,PSPëPNḡ PrPμCḡPjP°C,PëC,, PμPeCḡC,,PsP»C-P°C,PëPSPëPNḡ PrPμCḡPjP°C,PëC,.

PŸPīPμCḡPëC,,C-C±PSC- PIC-PrCḡCḡC,C,Cḡ: CḡP°PīC,PsPIPμ P·PSPëPḡPμPSPSCḡ C±Pë

PIC, C₇H₆P^oC, P^o C₁F₄»C₁F₃...C₁F₂, P±C-P»CH C₁ PIC₁F₃...P^oC..., PeC₇H₆PsPIP₆PIP₆P»P₆PI C₁ PsPePs, PeP^oC, P^oC₇H₆P^oPeC, P^o, C₁F₃C₁...C-C₁F₃, CH PI PsC₁P^oC...

P— P±PsPeC₁ C₁C₇H₆PsPiP₆PSC-C, P^oP»CH₆PSPsC— C₁F₃P₆C₁F₃, P₆PjP₆: C₁P₆C₁F₃, P₆C,, PSC-PeC, C₁C₇H₆-C₁F₃, PiC-PrPIP₆C%₀P₆PSP^o C₁P^oC₁F₃, PsC, P^o C₁P₆C₁F₃PsPIP₆PiC₁F₃Pe^oPSC₆H, P·P±C-P»CH₆C₆P₆PSPSC₁F₃PjPsP»PsC₁PSP₆Pe... P·P^oP»PsP·, PSP₆C, C₇H₆P₆PjP^oPSPSC₁F₃ C₁P₆C₁F₃-, PiPsC₇H₆C₆P₆PSPSC₁F₃ P₆C₁F₃PeC₁F₃»C₁F₃C₁F₃-C-, PSP^oP±C₇H₆C₁F₃Pe C₁F₃, P^oC, P₆PIP₆C... PsC₇H₆PIP^oPSC-PI, P^oPSPsC₇H₆PIP^oP·PjC-C₁F₃.

P^oPsC₁F₃PIC-Pr P·P^oC₁F₃, PsC₁F₃C₁F₃PIP^oPSPSC₁F₃ PiC-C₁F₃»C₁F₃ PIP₆C...PsPrC₁ PSP^o C₇H₆P₆PsPe.
P₆C-C₁F₃»C₁F₃ PIP₆C...PsPrC₁ PiC₇H₆P₆PiP^oC₇H₆P^oC, C₁ PSP^o C₇H₆P₆PsPe P±C₁F₃»P₆
C-PrP₆PSC, P₆C,, C-PePsPIP^oPSC- PSP₆P₆C₁F₃P₆P·P^oP·PSP^oC₁F₃PSC- PiPsP±C-C₁F₃PSC-
C₇H₆P₆P^oPeC₁F₃-C-. P₆C₁F₃PeC-P»CH₆PeP₆ PiC₇H₆Ps C, P^oPeC- C₇H₆P₆P^oPeC₁F₃-C-
PiPsPIC-PrPsPjP»C₁F₃C₁F₃, CH PrPsP±C₇H₆PsPIC-P»CH₆PSPs C, P^o PiPsPIC-PrPsPjP»P₆PSPSC₁F₃
PSP^oPrC...PsPrC₁F₃, CH PIC-Pr PiPsPiC₁F₃»C₁F₃C₁F₃-C- PSP₆PIC-PrPsPjPsC-
C₁F₃P₆C₁F₃»CH₆PSPsC₁F₃, C-, PSP₆ P·P^oPIP₆PrP₆ PjPsP₆PSP^o PrPsC₁F₃, PsPIC-C₇H₆PSPs
PsC₁F₃-PSP₆C, P₆C-C... C₁F₃P^oC₁F₃, PsC, C₁ C, P^o PIC₁F₃, P^oPSPsPIP₆C, P₆ PiC₇H₆P₆C₁F₃PSP₆P₆Pe
P·PiB₆F₆TM C₁F₃P·PsPe C-P· P₆PeC₁F₃PiPsP·P₆C₁F₃-C₁F₃ P»C-Pe^oC₇H₆C₁F₃PePsPiPs P·P^oC₁F₃PsP±C₁F₃.
P₁C- C₁F₃PIP₆C%₀P^o P±C₁F₃»P₆ P·P^oP·PSP^oC₁F₃PSC- C₁F₃Pe C₁F₃P₆P₆P· C-C...
C₁F₃P₆C₇H₆PsP·PSC-C₁F₃, CH, C₁F₃P^oC₁F₃, PsC, C₁ PiPsPIC-PrPsPjP»P₆PSPSC₁F₃,
PIC-PrC₁F₃C₁F₃, PSC-C₁F₃, CH C₁F₃-C, PePsPiPs P^oP»CH₆C, P₆C₇H₆PSP^oC, P₆PIPSPsPiPs
P·PiB₆F₆TM C₁F₃P·PeC₁, C, P^oPe C- C₁F₃P₆P₆P· PePsPjP±C-PSP^oC₁F₃-C₁F₃ C₁F₃»C... C,, P^oPeC, PsC₇H₆-PI.

P₆P₆C₇H₆C₁F₃PIP₆Ps-C₁F₃PrP₆PSPSC- C, P^o C₁F₃P₆C₇H₆P₆P±C₇H₆PsPIP^oC₁F₃PrP₆P»C₁F₃C₇H₆PSC-
C₁F₃PIP₆C%₀P^o. P₆PsPIC-PrPsPjP»C₁F₃»PsC₁F₃ PiC₇H₆Ps C₁F₃P₆C₇H₆PsP·PSC- C₁F₃P₆C₇H₆C₁F₃PIP₆Ps-
C₁F₃PrP₆PSPSC-, C₁F₃P₆C₇H₆P₆P±C₇H₆PsPIP^oC₁F₃PrP₆P»C₁F₃C₇H₆PSC- C, P^o C₁F₃PrP₆PSPSC-
C₁F₃PIP₆C%₀P^o, PIP₆Pe»C₇H₆C₁F₃P₆C₁F₃ C₁F₃P₆C₇H₆P₆P±C₇H₆PsPIP^oC₁F₃PrP₆P»C₁F₃C₇H₆PSC₁
PeC₇H₆PsPIP₆Ps, P₆C₁F₃, C₁F₃PrP±P^oC₇H₆P^oC...PSPsC-PrP^oP»CH₆PSC₁ C, P^o
PIPSC₁C₇H₆-C₆PsCH₆PsC₁F₃P₆P±C₇H₆P^oP»CH₆PSC₁ PeC₇H₆PsPIP₆Ps, P₆C₁F₃ C, P^o
P»P₆PiP₆PSP₆PI C₁ PeC₇H₆PsPIP₆Ps, P₆C₁F₃, C%₀Ps P±C₁F₃»P₆ PiPsPiB₆F₆TM C₁F₃P·P^oPSC- C₁ C₁F₃P^oC₁F₃-
C-P· P·P^oC₁F₃, PsC₁F₃PrP^oPSPSC₁F₃Pj C₁F₃P₆»PrP₆PSP^oC,, C-P»C₁F₃. Pj P±C-P»CH₆C₆PsC₁F₃, C-
PiP^oC₁F₃-C^oPSC, C-PI P±C₁F₃»P₆ PSP^oC₁F₃PIPSC- C,, P^oPeC, PsC₇H₆P₆ C₁F₃P₆C₇H₆C₁F₃PIP₆Ps-
C₁F₃PrP₆PSPSPsPiPs C₇H₆P₆P·P₆PeC₁. P₆PsPIC-PrPsPjP»C₁F₃»PsC₁F₃, C%₀Ps P±P^oPiP^oC, Ps P·
C₁F₃»C... C₁F₃PIP₆C%₀ PIP₆PSP₆PeP»P₆ PiC-Pr C₁F₃P^oC₁F₃ P±Ps PsPrC₇H₆P^oC₁ PiC-C₁F₃»C₁F₃
C₁F₃P₆PeC₁F₃PrP^oP»CH₆PSPsC— P^oPeC, P₆PIPSPsC₁F₃, C- C, P^o PrP₆PeC-P»CH₆Pe^o C₁F₃PIP₆C%₀
PIP₆PSP₆PeP»P₆ PsPrC₇H₆P^oC₁ PiC-C₁F₃»C₁F₃ P·P^oC₁F₃, PsC₁F₃PrP^oPSPSC₁F₃
C₁F₃P₆»PrP₆PSP^oC,, C-P»C₁F₃ P±P₆P· C₁F₃P₆PeC₁F₃PrP^oP»CH₆PSPsC— P^oPeC, P₆PIPSPsC₁F₃, C-.
P₁PSC₆C- C₁F₃PIP₆C%₀P^o PIP₆PSP₆PeP»P₆ PiC₇H₆PsC, C₁F₃PiPsPj PSP^oC₁F₃, C₁PiPSP₆C... PiPsPrP₆PS
C₁F₃PrPSC-PI PiC-C₁F₃»C₁F₃ P·P^oC₁F₃, PsC₁F₃PrP^oPSPSC₁F₃ C₁F₃P₆»PrP₆PSP^oC,, C-P»C₁F₃ C, P^o
C₁F₃P₆PeC₁F₃PrP^oP»CH₆PSPsC— P^oPeC, P₆PIPSPsC₁F₃, C-. P₆P₆PjPsP₆P»P₆PIPs
PIC₁F₃, P^oPSPsPIP₆C, P₆, C₁F₃ C₁F₃- C₁F₃PIP₆C%₀P^o PiPsPiB₆F₆TM C₁F₃P·P^oPSC- C-P·
P·P^oC₁F₃, PsC₁F₃PrP^oPSPSC₁F₃Pj PiC₇H₆P₆PiP^oC₇H₆P^oC, C₁, C-P· C₁F₃P₆PeC₁F₃PrP^oP»CH₆PSPsC₇H
P^oPeC, P₆PIPSC-C₁F₃, C₇H, C-P· PSP^oC₁F₃PIPSP₆PjP₆ C,, P^oPeC, PsC₇H₆P^oPjP₆ C₇H₆P₆P·P₆PeC₁ C₁F₃ C-P·
PePsPjP±C-PSP^oC₁F₃-C^oC₇H C₁F₃»C... C,, P^oPeC, PsC₇H₆-PI, C₁F₃ P· C-PSC₆PePjP₆
C,, P^oPeC, PsC₇H₆PjP₆.

P₆C₇H₆PsPIP₆PSPsC₁F₃PSP^o C, P^o P»C-PjC,, P^oC, P₆C₁F₃PSP^o C₁F₃P₆C₁F₃, P₆PjP₆:
PIP^oP·PsPsPeP»CH₆P·P₆PIPSP₆Pe₆ PeC₇H₆P₆P· Pj PSP₆PIP₆P»P₆PePsPjC₁ P·P^oPI C₁F₃P^oC₁F₃PSPs
PiC₇H₆P₆PiP₆PSP₆PSPsPjC₁ PrPsC₁F₃»C-PrP₆PIP₆PSPSC- P·P^oC₁F₃, PsC₁F₃PrP^oPSPSC₁F₃
PiC₇H₆P₆PiP^oC₇H₆P^oC, C₁ C₁F₃P₆»PrP₆PSP^oC,, C-P» P·P^o C₁F₃P^oC₁F₃, C₇H PiP^oC₁F₃-C^oPSC, C-PI P·
P»P₆PiP₆PSP₆PIP₆PsC₇H P^oC₇H₆, P₆C₇H₆-P^oP»CH₆PSPsC₇H PiC-PiP₆C₇H₆, P₆PSP·C-C^oC₇H,
PIC, PsC₇H₆P₆PSPSPsC₇H C%₀PsPrPs C₁F₃P₆C₇H₆PiPsPiPsPrC-P±PSPsPeP»C-C, P₆PIPSPsC—

P°PSPµPjC-C—, PiCḡPë P·P°CÍC,PsCÍCfPIP°PSPSC- CÍPëP»PrPµPSP°C,,C-P»Cf PiCḡPs
CḡPsP·PIPëC,PsPe PIP°P·PsPsPeP»CḡP·PëPIPSPëC... PeCḡPëP·C-PI, C%Ps
PiPsC,CḡPµP±CfPIP°P»Pë PiPsCÍPiC-C,P°P»C·P·P°C†C-C—, PiPsPIC-PrPsPjP»CḡP»PsCÍCḡ
C±P°CÍC,C-CëPµ, PSC-Pḡ PiCḡPë P·P°CÍC,PsCÍCfPIP°PSPSC- PiP»P°C†PµP±Ps.
PḡP»C-PSC-C±PSPµ P·PSP°C±PµPSPSCḡ C†C-C— C-PSC,,PsCḡPjP°C†C-C— PrP»Cḡ
PiP°C†C-C”PSC,C-PI, CḡPeC- P·P°CÍC,PsCÍCfPICfCḡC,Cḡ CÍPëP»PrPµPSP°C,,C-P» P·
PjPµC,PsCḡ P»C-PeCfPIP°PSPSCḡ PµCḡPµPeC,PëP»CḡPSPsC— PrPëCÍC,,CfPSPeC†C-C—,
PSPµPIC-PrPsPjPs.

P— P±PsPeCz PSPµCḡPIPSPiPsC— CÍPëCÍC,PµPjPë: C,CḡPëPIPSPiP°, C,CḡP°PSP·PëC,PsCḡPSP°
PiP»PsP±P°P»CḡPSP° P°PjPSPµP·C-Cḡ.

PŸPiPµC†PëC,,C-C±PSC- PIC-PrC±CzC,C,Cḡ

PŸP»CzC.... PµC-CÍP»Cḡ PIPëC...PsPrCf PiCḡPµPiP°CḡP°C,Cf PSP° CḡPëPSPsPe
PiPsPIC-PrPsPjP»CḡP»PsCÍCḡ PiCḡPs PIPëPiP°PrPePë CḡP°PiC,PsPIPSPiPs P·PSPëPḡPµPSPSCḡ
C±Pë PIC,CḡP°C,Pë CÍP»CfC...Cf, PiPsPIBḡ™ CḡP·P°PSC- Cf C±P°CÍC- C-P·
P·P°CÍC,PsCÍCfPIP°PSPSCḡPj CÍPëP»PrPµPSP°C,,C-P»Cf. Pj PrPµCḡPePëC... PIPëPiP°PrPeP°C...
PiPsPIC-PrPsPjP»CḡP»PsCÍCḡ PiCḡPs PSP°CḡPIPSC-CÍC,Cḡ PjPµPrPëC±PSPëC... CÍC,P°PSC-PI
C,P° C-PSCëPëC... C,,P°PeC,PsCḡC-PI, C%Ps PjPsPiP»Pë PIC-PrC-PiCḡP°C,Pë CḡPsP»Cḡ Cf
CḡPsP·PIPëC,PeCf PiPsP±C-C±PSPëC... CḡPµP°PeC†C-PNë P· P±PsPeCf CÍP»CfC...Cf. Pj
P±P°PiP°C,CḡPsC... PIPëPiP°PrPeP°C... C-PSC,,PsCḡPjP°C†C-Cḡ C%PsPrPs
PiPsPrP°P»CḡCëPsPiPs PjPµPrPëC±PSPSPiPs CÍPiPsCÍC,PµCḡPµPḡPµPSPSCḡ
PIC-PrCÍCfC,PSCḡ. P·PëP·PSP°C±PëC,Pë, C±Pë C†C- CḡPIPëC%P° PiCḡCḡPjPs
PiPsPIBḡ™ CḡP·P°PSC- C-P· P·P°CÍC,PsCÍCfPIP°PSPSCḡPj CÍPëP»PrPµPSP°C,,C-P»Cf, P·
PSP°CḡPIPSPëPjPë C,,P°PeC,PsCḡP°PjPë CḡPëP·PëPeCf PIC,CḡP°C,Pë CÍP»CfC...Cf, P·
PePsPjP±C-PSP°C†C-C”Cḡ C†PëC... C,,P°PeC,PsCḡC-PI C±Pë P· C-PSCëPëPjPë
C,,P°PeC,PsCḡP°PjPë, PSPµPjPsPḡP»PëPIPs.

P—C-Cḡ. PŸPëPjC±P°CÍPsPIP° PIC,CḡP°C,P° P·PsCḡCf, PiPsC±PµCḡPIPSPSC-PSPSCḡ PsC±PµPNë,
PiPµC±C-PSPSCḡ PI PsC±P°C..., PiC-PrPIPëC%PµPSPSCḡ PIPSCfC,CḡC-CëPSCḡPsPsC±PSPSPiPs
C,PëCÍPeCf, PSP°P±CḡCḡPe CÍC-C,PeC-PIPePë, CÍCfPrPëPSPSC-
P·P°C...PIPSPḡCḡPIP°PSPSCḡ CÍC-C,PeC-PIPePë C±Pë PeCḡPsPIPsC,PµC±P°,
PIC-PrCëP°CḡCfPIP°PSPSCḡ CÍPeP»PëCÍC,PsPiPs C,C-P»P°.

PµC-CÍP»Cḡ PIPëC...PsPrCf PiCḡPµPiP°CḡP°C,Cf PSP° CḡPëPSPsPe CḡC-PrPePs
PiPsPIC-PrPsPjP»CḡP»PsCÍCḡ PiCḡPs PIPëPiP°PrPePë PSPµP°CḡC,PµCḡC-P°P»CḡPSPsC—
PiPµCḡPµPrPSCḡPsC— C-CëPµPjC-C±PSPsC— PSPµPICḡPsPiP°C,C-C— P·PsCḡPsPIPsPiPs
PSPµCḡPIP°, C%Ps C” PiCḡPëC±PëPSPsCḡ P·PSPëPḡPµPSPSCḡ P·PsCḡCf,
PIPeP»CḡC±P°CḡC±Pë PiPsCÍC,C-PNëPSCf PIC,CḡP°C,Cf P·PsCḡCf, CḡPeC- P±CfP»Pë
PiPsPIBḡ™ CḡP·P°PSC- Cf C±P°CÍC- C-P· P·P°CÍC,PsCÍCfPIP°PSPSCḡPj
C-PSPiC-P±C-C,PsCḡC-PI P±P”P·5, PIPeP»CḡC±P°CḡC±Pë CÍPëP»PrPµPSP°C,,C-P». Pj
P±P°PiP°C,CḡPsC...B PiP°C†C-C”PSC,C-PI, P°P»Pµ PSPµ Cf PICÍC-C..., P±CfP»Pë
PSP°CḡPIPSC- P°PSP°C,PsPjC-C±PSC- P°P±Ps CÍCfPrPëPSPSC- C,,P°PeC,PsCḡPë CḡPëP·PëPeCf
CḡPsP·PIPëC,PeCf PSPµP°CḡC,PµCḡC-P°P»CḡPSPsC— PiPµCḡPµPrPSCḡPsC—
C-CëPµPjC-C±PSPsC— PSPµPICḡPsPiP°C,C-C— P·PsCḡPsPIPsPiPs PSPµCḡPIP°,
PIPeP»CḡC±P°CḡC±Pë (P°P»Pµ PSPµ PsP±PjPµPḡCfCḡC±PëCÍCḡ) C,P°PeC-: PSPëP·CḡPePµ
CÍPiC-PIPIC-PrPSPsCëPµPSPSCḡ PrC-P°PjPµC,CḡP° PµPeCÍPeP°PIP°C†C-C— C,P° PrPëCÍPeP°
P·PsCḡPsPIPsPiPs PSPµCḡPIP° (P·P°CÍC,C-PNëPSPëPNë PrPëCÍPe P·PsCḡPsPIPsPiPs PSPµCḡPIP°),
PIC-Pe PiPsPSP°Pr 50 CḡPsPeC-PI, P°CḡC,PµCḡC-P°P»CḡPSP° PiC-PiPµCḡC,PµPSP·C-Cḡ,
P·P°C...PIPSPḡCḡPIP°PSPSCḡ PePsCḡPsPSP°CḡPSPëC... P°CḡC,PµCḡC-PNë,
PiC-PiPµCḡP»C-PiC-PrPµPjC-Cḡ C,P° PiP°P»C-PSPSCḡ. PkPµPjPsPḡP»PëPIPs

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. PjPµ PrP°C” P·PjPsPiCf PïC½PsPIPsPrPëC,Pë
P±PµP·PïPµC½PµC½PIPSPëPñ 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°.

PÿPµC½PjC-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°Pÿ.

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

PjPïP°PePsPIPëP°.

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-PMñ PïP°C±C†C-.

P½P°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.

**P½C-CÍC†PµP·PSP°C...PsPrP¶PµPSPSC¼ PIPëC½PsP±PSPëPeP° C,P° P°PrC½PµCÍP°
PjC-CÍC†C¼ PïC½PsPIP°PrP¶PµPSPSC¼ PñPsPiPs PrC-C¼P»C½PSPsCÍC,C-.**

P’CfP». P”P°PSC-C†P° 5, 48000 P½PsPïC½PëPIPSPëC†C¼, P½PsC½PIP°C,C-C¼.