

**Р†РќРŸРŸР RJPЛP!P†PĪ**

**РгР»CĪ PjPμPгPëC†PSPsPiPs P·P°CĪC,PsCĪCĭPIP°PSPSCĪ P»C-PeP°CĤCĪCĤPePsPiPs  
P·P°CĪPsP±Cĭ**

**РμP»P·PŸPĥP»**

**(PLETOL)**

**PŸPeP»P°Pг:**

*РгC-CĤC†P° CĤPμC†PsPIPëPSP°: cilostazol;*

1 C,P°P±P»PμC,PeP° PjC-CĪC,PëC,CĤ C†PëP»PsCĪC,P°P·PsP»Cĭ 50 PjPi P°P±Ps 100 PjPi;

C,P°P±P»PμC,PePë PĭPs 50 PjPi: PгPsPiPsPjC-PŸPSC- CĤPμC†PsPIPëPSPë:

*PeCĤPsC...PjP°P»CĤ PeCĭPeCĭCĤCĭPгP·CĪPSPëP№,*

*PiC-PгCĤPsPeCĪPëPjPμC,PëP»PĭCĤPsPiC-P»C†PμP»CĤP»PsP·P° P· 5, C†PμP»CĤP»PsP·P°*

*PjC-PeCĤPsPeCĤPëCĪC,P°P»C-C†PSP° 101, PjP°PiPSC-CĤ CĪC,PμP°CĤP°C,, C†PμP»CĤP»PsP·P°*

*PjC-PeCĤPsPeCĤPëCĪC,P°P»C-C†PSP° 12, PeP°P»CĤC†C-CĤ PeP°CĤPjPμP»PsP·P°;*

C,P°P±P»PμC,PePë PĭPs 100 PjPi: PгPsPiPsPjC-PŸPSC- CĤPμC†PsPIPëPSPë:

*PeCĤPsC...PjP°P»CĤ PeCĭPeCĭCĤCĭPгP·CĪPSPëP№,*

*PiC-PгCĤPsPeCĪPëPjPμC,PëP»PĭCĤPsPiC-P»C†PμP»CĤP»PsP·P° P· 5, C†PμP»CĤP»PsP·P°*

*PjC-PeCĤPsPeCĤPëCĪC,P°P»C-C†PSP° 101, PjP°PiPSC-CĤ CĪC,PμP°CĤP°C,, PeP°P»CĤC†C-CĤ*

*PeP°CĤPjPμP»PsP·P°.*

**P»C-PeP°CĤCĪCĤPeP° C,,PsCĤPjP°.** PŸP°P±P»PμC,PePë.

*PĥCĪPSPsPIPSC- C,,C-P·PëPePs-C...C-PjC-C†PSC- PIP»P°CĪC,PëPIPsCĪC,C-:*

C,P°P±P»PμC,PePë PĭPs 50 PjPi: PeCĤCĭPiP»C-, PĭP»P°CĪPeC- C,P°P±P»PμC,PePë PIC-Pг

*P±C-P»PsPiPs PгPs PjP°P№PŸPμ P±C-P»PsPiPs PePsP»CĤPsCĤCĭ, P·*

*PiCĤP°PIC-CĤPIP°PSPSCĪPj B«50B» P· PsPгPSPsPiPs P±PsPeCĭ;*

C,P°P±P»PμC,PePë PĭPs 100 PjPi: PeCĤCĭPiP»C-, PĭP»P°CĪPeC- C,P°P±P»PμC,PePë PIC-Pг

*P±C-P»PsPiPs PгPs PjP°P№PŸPμ P±C-P»PsPiPs PePsP»CĤPsCĤCĭ, P·*

*PiCĤP°PIC-CĤPIP°PSPSCĪPj B«100B» P· PsPгPSPsPiPs P±PsPeCĭ.*

**P»P°CĤPjP°PePsC,PμCĤP°PĭPμPIC,PëC†PSP° PiCĤCĭPĭP°.**

*PĥPSC,PëC,CĤPsPjP±PsC,PëC†PSC- P·P°CĪPsP±Pë. P†PSPiC-P±C-C,PsCĤPë*

*P°PiCĤPμPiP°C†C-C— C,CĤPsPjP±PsC†PëC,C-PI P·P° PIPëPSCĪC,PePsPj PiPμPĭP°CĤPëPSCĭ.*

*РлPsPг PĥPŸPГ B01A C23.*

**P»P°CĤPjP°PePsP»PsPiC-C†PSC- PIP»P°CĪC,PëPIPsCĪC,C-.**

*PαP°CḡPjP°PePsPrPëPSP°PjC-PeP°.*

PjPëP»PsCfC,P°P·PsP» C” C-PSPiC-P±C-C,PsCḡPsPj P°PiCḡPμPiP°C†C-C—  
C,CḡPsPjP±PsC†PëC,C-PI. PμCḡPμPiP°CḡP°C, PiPsPeCḡP°C%oCfC” P·PrP°C,PSC-CfC,CH  
PiPμCḡPμPSPsCfPëC,Pë C,,C-P·PëC±PSC- PSP°PIP°PSC,P°P¶PμPSPSCf, C¶PeCf  
PsC†C-PSCḡCḡC,CH P·P° P°P±CfPsP»CḡC,PSPsCḡ PrPëCfC,P°PSC†C-C”Cḡ PiCḡPë  
PiPμCḡPμPjC-P¶PSC-PMë PeCfP»CHPiP°PiPsCfC,C- (P°P±Ps P·P° PjP°PeCfPëPjP°P»CHPSPsCḡ  
PrPëCfC,P°PSC†C-C”Cḡ C...PsPrCHP±Pë (PḡP”PΓ)) C,P° PiPsC±P°C,PePsPiPsCḡ  
PrPëCfC,P°PSC†C-C”Cḡ PiPμCḡPμPjC-P¶PSPsC— PeCfP»CHPiP°PiPsCfC,C- (P°P±Ps P·P°  
P±PμP·P±PsP»C-CfPSPsCḡ PrPëCfC,P°PSC†C-C”Cḡ C...PsPrCHP±Pë (P’P”PΓ)) Cf  
C,PμCfC,CfPIP°PSPSC- PSP° P±C-PiPsPIC-PMë PrPsCḡC-P¶C†C-. P—P°  
CḡPμP·CfP»CHC,P°C,P°PjPë PrPsCfP»C-PrP¶PμPSCḡ PiCḡPë CḡC-P·PSPëC...  
PSP°PIP°PSC,P°P¶PμPSPSCfC... P±CfP»Ps PICfC,P°PSPsPiP»PμPSPs P·PSP°C±PSPμ  
P°P±CfPsP»CḡC,PSPμ PiPsPeCḡP°C%oPμPSPSCf PSP° 42 PjPμC,CḡPë PḡP”PΓ PiCḡPë  
P·P°CfC,PsCfCfPIP°PSPSC- PiCḡPμPiP°CḡP°C,Cf PiPsCḡC-PIPSCfPSPs P· C,P°PePsCḡ  
PiCḡPëB PiCḡPëPMëPsPjC- PiP»P°C†PμP±Ps. PjPμ PIC-PrPiPsPIC-PrP°C”  
PIC-PrPSPsCfPSPsPjCf PiPsPeCḡP°C%oPμPSPSCḡ PSP° 100 % PiPsCḡC-PIPSCfPSPs P·  
PiP»P°C†PμP±Ps. PjPμPMë PμC,,PμPeC, P±CfPI PrPμC%oPs PSPëP¶C†PëPj Cf  
C...PiPsCḡPëC... PSP° C†CfPeCḡPsPiPëPMë PrC-P°P±PμC,,.

PjPëP»PsCfC,P°P·PsP» PiPëC¶PIP»C¶C” PiP°P·PsPrPëP»P°C,P°C,PsCḡPSPëPMë PμC,,PμPeC,,  
C%oPs P±CfP»Ps PiC-PrC,PIPμCḡPrP¶PμPSPs PiPëPjC-CḡCḡPiP°PSPSCfPj PeCḡPsPiPsC,PsPeCf  
PSPëP¶PSC-C... PeC-PSC†C-PIPsPe P·P° PrPsPiPsPjPsPiPsCḡ C,PμPSP·PsPjPμC,CḡPëC±PSPsC—  
PiP»PμC,PëP·PjPsPiCḡP°C,,C-C—. PiPëP»PsCfC,P°P·PsP» C,P°PePsP¶ C-PSPiC-P±CfC”  
PiCḡPsP»C-C,,PμCḡP°C†C-Cḡ PeP»C-C,PëPS PiP»P°PrPePëC... PjBḡ™ C¶P·C-PI C,P°  
C-PSPiC-P±CfC” CḡPμP°PeC†C-Cḡ PiPëPIC-P»CHPSPμPSPSCf C,CḡPsPjP±PsC†PëC,C-PI  
C,CḡPsPjP±PsC†PëC,P°CḡPSPsPiPs C,,P°PeC,PsCḡP° CḡPsCfC,Cf C,P° PF-4 Cf  
C,CḡPsPjP±PsC†PëC,P°C... P»CḡPrPëPSPë.

P”PsCfP»C-PrP¶PμPSPSCf PiCḡPsPrPμPjPsPSCfC,CḡCfPIP°P»Pë, C%oPs C†PëP»PsCfC,P°P·PsP»  
CfPiCḡPëC±PëPSCfC” PsP±PsCḡPsC,PSPμ PiCḡPëPiPSC-C±PμPSPSCf P°PiCḡPμPiP°C†C-C—  
C,CḡPsPjP±PsC†PëC,C-PI. P†PSPiC-P±CfPIP°PSPSCf PμC,,PμPeC,PëPiPSPμ PiCḡPsC,Pë  
PSPëP·PePë P°PiCḡPμPiP°PSC,C-PI (PiPeP»CḡC±P°CḡC±Pë P°CḡP°C...C-PrPsPSPsPICf  
PePëCfP»PsC,Cf, PePsP»P°PiPμPS, P°PrPμPSPsP·PëPSPrPëC,,PsCfC,,P°C, (PḡP”Pα) C,P°  
P°PrCḡPμPSP°P»C-PS); Cf PiP°C†C-C”PSC,C-PI C-PSPiC-P±CfPIP°PSPSCf C,CḡPëPiP°C” PrPs  
12 PiPsPrPëPS, P° PiC-CfP»C¶ P·P°PIPμCḡC€PμPSPSCf PiCḡPëPMëPsPjCf  
C†PëP»PsCfC,P°P·PsP»Cf PIC-PrPSPsPiP»PμPSPSCf P°PiCḡPμPiP°C†C-C—  
PiCḡPsC...PsPrPëP»Ps PI PjPμP¶P°C... 48-96 PiPsPrPëPS P±PμP· PμC,,PμPeC,Cf  
CḡPëPePsC€PμC,Cf (PiC-PiPμCḡP°PiCḡPμPiP°C†C-C—). PÿP°PePsP¶ PICfC,P°PSPsPiP»PμPSPs  
PiP»PëPI C†PëP»PsCfC,P°P·PsP»Cf PSP° P»C-PiC-PrPë, C¶PeC- C†PëCḡPeCfP»CḡCḡC,CH Cf  
PiP»P°P·PjC- PeCḡPsPIC-. PμCḡPëPMëPsPj PiCḡPμPiP°CḡP°C,Cf P·PSPëP¶CfC” CḡC-PIPμPSCḡ  
C,CḡPëPiP»C-C†PμCḡPëPrC-PI C,P° PiC-PrPiPëC%oCfC” CḡC-PIPμPSCḡ HDL-  
C...PsP»PμCfC,PμCḡPsP»Cf. PÿCḡPëPiP°P»Pμ P·P°CfC,PsCfCfPIP°PSPSCf PiCḡPμPiP°CḡP°C,Cf  
PSPμ PiPëPeP»PëPeP°P»Ps P·P±C-P»CHC€PμPSPSCf CḡC-PIPSCf P»PμC,P°P»CHPSPsCfC,C-  
CfPμCḡPμPr PiP°C†C-C”PSC,C-PI PiPsCḡC-PIPSCfPSPs P· PiP»P°C†PμP±Ps.

*PαP°CḡPjP°PePsPeC-PSPμC,PëPeP°.*

PμCḡPë CḡPμPiCfP»C¶CḡPSPsPjCf PiCḡPëPMëPsPjC- C†PëP»PsCfC,P°P·PsP»Cf PI PrPsP·C- 100  
PjPi 2 CḡP°P·Pë PSP° PrPsP±Cf Cf PiP°C†C-C”PSC,C-PI C-P· P·P°C...PiPsCḡCḡPiP°PSPSCfPjPë  
PiPμCḡPëC,,PμCḡPëC±PSPëC... CfCfPrPëPS CfC,P°P±C-P»CHPSPëPMë CfC,P°PS

PrPsCfCfCfPiP°C"C,CHbCfCfCf PiCfPSc,CfPiPsPj 4 PrPSC-PI. PыP°PeCfPөPjP°P»CHbPSP°  
PePsPSCfPµPSC,CfP°CfC-CfCf (PŸ<sub>max</sub>) C†PөP»PsCfC,P°P·PsP»Cf C,P° P№PsPiPs  
PiPµCfPiPөPSPSPөC... PjPµC,P°P±PsP»C-C,C-PI PiC-PrPiPөC%oCfC"C,CHbCfCfCf PjPµPSCө  
PiCfPScPiPsCfCfCfC-PMөPSPs P· PiC-PrPiPөC%oPµPSPSCfPj PrPsP·Pө, PiCfPSc,Pµ PiP»PsC%oP°  
PiC-Pr PeCfPөPiPsCf B«PePsPSCfPµPSC,CfP°CfC-CfCfCf» (AUC)  
C†PөP»PsCfC,P°P·PsP»Cf C,P° P№PsPiPs PjPµC,P°P±PsP»C-C,C-PI  
P·P±C-P»CHbCөCfC"C,CHbCfCfCf PiCfPөP±P»PөP·PSPs PiCfPScPiPsCfCfCfC-PMөPSPs PrPs  
PrPsP·CfPiP°PSPSCfCf. PiPiPSPөP№ PiPµCfCf-PsPr PSP°PiC-PIPiPөPiPµPrPµPSPSCfCf  
C†PөP»PsCfC,P°P·PsP»Cf CfPөP»P°PrP°C" 10,5 PiPsPrPөPSPө. P†CfPSCfC" PrPiP°  
PiPsP»PsPiPSPөC... PjPµC,P°P±PsP»C-C,Pө бf" PrPµPiC-PrCfPScC†PөP»PsCfC,P°P·PsP» C,P° 4'-  
C,CfP°PSCf-PiC-PrCfPScPeCfPөC†PөP»PsCfC,P°P·PsP», C%oPs PjP°CfC,CHb P±P»PөP·CHbPeC-  
PiPsPeP°P·PSPөPePө PSP°PiC-PIPiPөPiPµPrPµPSPSCfCf. P" PµPiC-PrCfPScPjPµC,P°P±PsP»C-C,  
PjP°C" Cf 4-7 CfP°P·C-PI PiPөC%oCf P°PSC,PөC,CfPScPjP±PsC,PөC†PSCf  
P°PeC,PөPiPSC-CfC,CHb, PSC-P¶ PiPөC...C-PrPSP° CfPµC†PsPiPөPSP°, P° 4'-C,CfP°PSCf-  
PiC-PrCfPScPeCfPөPjPµC,P°P±PsP»C-C, бf" 1/5 PiC-Pr P°PeC,PөPiPSPsCfC,C-  
C†PөP»PsCfC,P°P·PsP»Cf. PыPsPSCfPµPSC,CfP°CfC-C— Pi PiP»P°P·PjC- PeCfPScPIC-  
(PsC,CfPөPjP°PSC- P·P° PrPsPiPsPjPsPiPsCf AUC) PrPµPiC-PrCfPSc- C,P° 4'-C,CfP°PSCf-  
PiC-PrCfPScPeCfPөPjPµC,P°P±PsP»C-C,C-PI PiCfPөP±P»PөP·PSPs CfPөP»P°PrP°CfC,CHb 41 %  
C,P° 12 % PiC-Pr PePsPSCfPµPSC,CfP°CfC-C— C†PөP»PsCfC,P°P·PsP»Cf PiC-PrPiPsPIC-PrPSPs.

*PµPµC,P°P±PsP»C-P·Pj*

PiPөP»PsCfC,P°P·PsP» PiPөPiPsPrPөC,CHbCfCfCf PiPµCfPµPiP°P¶PSPs P·P° PrPsPiPsPjPsPiPsCf  
PjPµC,P°P±PsP»C-P·PjCf C,P° PiPsPrP°P»CHbCөPsPiPs PiPөPiPµPrPµPSPSCfCf P№PsPiPs  
PjPµC,P°P±PsP»C-C,C-PI P·C- CfPµC†PµCf. PµPµCfPiPөPSPSC- C-P·PsC,,PµCfPjPµPSC,Pө  
C†PөC,PsC...CfPScPjCf P 450, CfPeC- P±PµCfCfC,CHb CfC†P°CfC,CHb Cf P№PsPiPs  
PjPµC,P°P±PsP»C-P·PjC-, бf" CYP3A4, PjPµPSCөPsCf PjC-CfPScCf бf" CYP2C19 C,P° C%oPµ  
PjPµPSCөPsCf PjC-CfPScCf бf" CYP1A2.

*P'PөPiPµPrPµPSPSCfCf*

P"PsP»PsPiPSPөP№ CөP»CfC... PiPөPiPµPrPµPSPSCfCf бf" P·C- CfPµC†PµCf (74 %),  
P·P°P»PөCөPePsPIC- PeC-P»CHbPePsCfC,C- PiPөPrC-P»CfCfC,CHbCfCfCf P· PeP°P»PsPj.  
PкPµP·PSP°C†PSC- PeC-P»CHbPePsCfC,C- PSPµP·PjC-PSPµPSPsPiPs C†PөP»PsCfC,P°P·PsP»Cf  
PiPөPrC-P»CfCfC,CHbCfCfCf P·C- CfPµC†PµCf, P° PjPµPSCө PSC-P¶ 2 % P№PsPiPs PrPsP·Pө  
PiPөPrC-P»CfC"C,CHbCfCfCf Cf PiPөPiP»CfPrC- PrPµPiC-PrCfPScC†PөP»PsCfC,P°P·PsP»Cf.  
PµCfPөP±P»PөP·PSPs 30 % PiC-Pr PiPsC†P°C,PePsPiPsC— PrPsP·Pө PiPөPrC-P»CfC"C,CHbCfCfCf  
P·C- CfPµC†PµCf CfPe 4'-C,CfP°PSCf-PiC-PrCfPScPeCfPөPjPµC,P°P±PsP»C-C,.  
P—P°P»PөCөPePsPiP° PeC-P»CHbPeC-CfC,CHb PiPөPrC-P»CfC"C,CHbCfCfCf CfPe CfCfPjP°  
PjPµC,P°P±PsP»C-C,C-PI, P¶PsPrPµPS P· CfPePөC... PSPµ PiPµCfPµPiPөC%oCfC" 5 % PiC-Pr  
P·P°PiP°P»CHbPSPsC— PeC-P»CHbPePsCfC,C-.

*P PsP·PiPsPrC-P»*

PiPөP»PsCfC,P°P·PsP» P·Piбf™ CfP·CfC"C,CHbCfCfCf P· P±C-P»PeP°PjPө PSP° 95-98 %,  
PiPµCfPµPiP°P¶PSPs P· P°P»CHbP±CfPjC-PSPsPj. P" PµPiC-PrCfPScPjPµC,P°P±PsP»C-C, C,P° 4'-  
C,CfP°PSCf-PiC-PrCfPScPeCfPөPjPµC,P°P±PsP»C-C, P·Piбf™ CfP·CfCfC,CHbCfCfCf P·  
P±C-P»PeP°PjPө PSP° 97,4 % C,P° 66 % PiC-PrPiPsPIC-PrPSPs.

PкPµPjP°C" PiC-PrC,PiPµCfPµPrP¶PµPSPSCfCf P·PrP°C,PSPsCfC,C- C†PөP»PsCfC,P°P·PsP»Cf  
C-PSPrCfPeCfPiP°C,Pө PjC-PeCfPScCfPsPjP°P»CHbPSC- C,,PµCfPjPµPSC,Pө PiPµC†C-PSPөPө.  
PαP°CfPjP°PePsPeC-PSPµC,PөPeP° C†PөP»PsCfC,P°P·PsP»Cf C,P° P№PsPiPs

PjPμC,P°P±PsP»C-C,C-PI PSPμ P·P°P»PμP¶P°P»P° P·PSP°C±PSPsC¶ PjC-C¶PpsC¶ PIC-Pr PIC-PeCf P°P±Ps C¶C,P°C,C- PIP°C†C-C”PSC,C-PI PIC-PePsPj 50-80 C¶PpsPeC-PI.

PJ PIP°C†C-C”PSC,C-PI P· C,C¶P¶PePsC¶ PSPeC¶PePsIPsC¶ PSPμPrPsC¶C,P°C,PSC-C¶C,C¶ PIC-P»CHPSP° C,,C¶P°PeC†C-C¶ C†PëP»PsC¶C,P°P·PsP»Cf P±CfP»P° PSP° 27 % PIPeC%PsC¶, P° PŸ<sub>max</sub> C,P° AUC P±CfP»Pë PIC-PrPipPsPIC-PrPSPs PSP° 29 % C,P° 39 % PSPeP¶C±PëPjPë, PSC-P¶ Cf PsC¶C-P± P· PSPsC¶PjP°P»CHPSPsC¶ C,,CfPSPeC†C-C”C¶ PSPeC¶PsPe. PŸ<sub>max</sub> C,P° AUC PrPμPic-PrC¶PpsPjPμC,P°P±PsP»C-C,Cf P±CfP»Pë PIC-PrPipPsPIC-PrPSPs PSP° 41 % C,P° 47 % PSPeP¶C±PëPjPë Cf PIP°C†C-C”PSC,C-PI P· C,C¶P¶PePëPjPë PipsC¶CfCεPμPSPSC¶PjPë PSPeC¶PsPe PipsC¶C-PIPSC¶PSPs P· PIP°C†C-C”PSC,P°PjPë P· PSPsC¶PjP°P»CHPSPsC¶ C,,CfPSPeC†C-C”C¶ PSPeC¶PsPe. PŸ<sub>max</sub> C,P° AUC 4'-C,C¶P°PSC¶- Pic-PrC¶PpsPeC¶PëC†PëP»PsC¶C,P°P·PsP»Cf P±CfP»Pë PSP° 173 % C,P° 209 % PIC-PrPipPsPIC-PrPSPs PIPeC%PëPjPë Cf PIP°C†C-C”PSC,C-PI P· C,C¶P¶PePsC¶ PSPeC¶PePsIPsC¶ PSPμPrPsC¶C,P°C,PSC-C¶C,C¶. PkPμPjP°C” PrP°PSPeC... C%PsPrPs PIP°C†C-C”PSC,C-PI P· PipsPjC-C¶PSPsC¶ C,P° C,C¶P¶PePsC¶ PIPμC±C-PSPePsIPsC¶ PSPμPrPsC¶C,P°C,PSC-C¶C,C¶.

**РљР»С-РSC-С±РSC- С...Р°С¶Р°РeС,РμС¶РeС¶С,РeРeРe.**

**РμРsРeР°Р·Р°РSPSC¶.**

P”P»C¶ P·P±C-P»CHCεPμPSPSC¶ PjP°PeC¶PëPjP°P»CHPSPsC— P±PμP·P±PsP»C-C¶PSPsC— PrPëC¶C,P°PSC†C-C— C...PsPrCHP±Pë Cf PIP°C†C-C”PSC,C-PI P· PIPμC¶PμPjC-P¶PSPsC¶ PeCfP»CHPip°PIC-C¶C,C¶, C¶PeC- PSPμ PjP°C¶C,C¶ P±PsP»C¶ Cf C¶C,P°PSC- C¶PipsPePsC¶ C,P° PsP·PSP°Pe PSPμPeC¶PsP·Cf PIPμC¶PëC,,PμC¶PëC±PSPeC... C,PeP°PSPëPs (P·P°C...PIPpsC¶PIP°PSPSC¶ PIPμC¶PëC,,PμC¶PëC±PSPeC... P°C¶C,PμC¶C-PNε, C¶C,P°PrC-C¶ II P·P° PαPsPSC,PμPNεPSPsPj).

P—P°C¶C,PsC¶PsPICfPIP°C,Pë C¶Pe C,PμC¶P°Pic-C¶ PrC¶CfPipsC— P»C-PSC-C— PrP»C¶ PIP°C†C-C”PSC,C-PI, Cf C¶PePëC... P·PjC-PSP° C¶C,PëP»C¶ P¶PëC,C,C¶ (PIPeP»C¶C±P°C¶C±Pë PIC-PrPjPsPICf PIC-Pr PeCfC¶C-PSPSC¶ C,P° PePsPSC,C¶PsP»CHPpsPIP°PSC- PïC¶PsPïC¶P°PjPë P·P°PSC¶C,C¶ C¶PipsC¶C,PsPj) C,P° C-PSCεC- PIC-PrPipPsPIC-PrPSC- P·P°C...PsPrPë PSPμ PïC¶PëP·PIPμP»Pë PrPs P·PSP°C±PSPsPips PipsC¶P»P°P±P»PμPSPSC¶ C¶PëPjPïC,PsPjC-PI PIPμC¶PμPjC-P¶PSPsC— PeCfP»CHPip°PIPpsC¶C,-.

**РμС¶РsС,РeРPipsРeР°Р·Р°РSPSC¶.**

- P’C-PrPsPjP° Pic-PipμC¶C±CfC,P»PëPIC-C¶C,C¶ PrPs C†PëP»PsC¶C,P°P·PsP»Cf P°P±Ps PrPs P±CfPrCH-C¶PePsPips PePsPjPipsPSPμPSC,P° PïC¶PμPip°C¶P°C,Cf;
- C,C¶P¶PeP° PSPeC¶PePsPIP° PSPμPrPsC¶C,P°C,PSC-C¶C,C¶ (PeP»C-C¶PμPSC¶ PeC¶PμP°C,PëPSC-PSCf £ 25 PjP»/C...PI);
- PipsPjC-C¶PSP° P°P±Ps C,C¶P¶PeP° PIPμC±C-PSPePsPIP° PSPμPrPsC¶C,P°C,PSC-C¶C,C¶;
- P·P°C¶C,C-PNεPSP° C¶PμC¶C†PμPIP° PSPμPrPsC¶C,P°C,PSC-C¶C,C¶;
- PIP°Pic-C,PSC-C¶C,C¶;

- P±CfPrCH-ClPeP° PIC-PrPsPjP° CÍC...PëP»CHPSC-CÍC,CH PrPs PeCĤPsPIPsC,PµC±C- (PSP°PíCĤPëPeP»P°Pr, PIPëCĤP°P·PeP° CëP»CfPSPeP° P°P±Ps PrPIP°PSP°PrC†CĤC,PëPíP°P»PsC— PePëCëPePë Cf CÍC,P°PrC-C— P·P°PiPsCÍC,CĤPµPSPSCĤ, PSPµC%PsPrP°PIPsC-PNë PiPµPjPsCĤP°PiC-C†PSPëPNë C-PSCÍCfP»CHC, (PíCĤPsC,ClPiPsPj 6 PjC-CÍCĤC†C-PI), PíCĤPsP»C-C,,PµCĤP°C,PëPIPsP° C,,PsCĤPjP° PrC-P°P±PµC,PëC†PSPSC— CĤPµC,PëPSPsPíP°C,C-C—, CÍP»P°P±PsPePsPSC,CĤPsP»CHPsPIP°PSP° P°CĤC,PµCĤC-P°P»CHPSP° PiC-PíPµCĤC,PµPSP·C-Cl);

- CëP»CfPSPsC†PePsPIP° C,P°C...C-PeP°CĤPrC-Cl, C,,C-P±CĤPëP»ClC†C-Cl CëP»CfPSPsC†PeC-PI P°P±Ps PjCfP»CHC,PëP»PsPeCfP»ClCĤPSP° CëP»CfPSPsC†PePsPIP° PµPeC,PsPíC-Cl PI P°PSP°PjPSPµP·C- Cf PíP°C†C-C”PSC,C-PI, ClPeC- PíCĤPsC...PsPrPëP»Pë P°P±Ps PSPµ PíCĤPsC...PsPrPëP»Pë PIC-PrPíPsPIC-PrPSCr C,PµCĤP°PíC-CĤ, P° C,P°PePsP¶ PíPsPrPsPIP¶PµPSPSCĤ C-PSC,PµCĤPIP°P»Cf QT;

- C,ClP¶PeP° C,P°C...C-P°CĤPëC,PjC-Cl PI P°PSP°PjPSPµP·C-;

- PsPrPSPsC†P°CÍPSPµ P»C-PeCfPIP°PSPSCĤ PrPIPpsPjP° P°P±Ps P±C-P»CHCëPµ PrPsPrP°C,PePsPIPëPjPë P°PSC,PëC,CĤPsPjP±PsC†PëC,P°CĤPSPëPjPë P·P°CÍPsP±P°PjPë P°P±Ps P°PSC,PëPePsP°PiCfP»ClPSC,P°PjPë (PSP°PíCĤPëPeP»P°Pr, P°C†PµC,PëP»CÍP°P»C-C†PëP»PsPIP° PePëCÍP»PsC,P°, PeP»PsPíC-PrPsPíCĤPµP»CH, PiPµPíP°CĤPëPS, PIP°CĤC,,P°CĤPëPS, P°C†PµPSPsPeCfPjP°CĤPsP», PrP°P±C-PiP°C,CĤP°PS, CĤPëPIP°CĤPsPeCÍP°P±P°PS P°P±Ps P°PiC-PeCÍP°P±P°PS);

- PSPµCÍC,P°P±C-P»CHPSP° CÍC,PµPSPsPeP°CĤPrC-Cl, C-PSC,,P°CĤPeC, PjC-PsPeP°CĤPrP° PíCĤPsC,ClPiPsPj PsCÍC,P°PSPSC-C... 6 PjC-CÍCĤC†C-PI P°P±Ps PePsCĤPsPSP°CĤPSPµ PIC,CĤCfC†P°PSPSCĤ PíCĤPsC,ClPiPsPj PsCÍC,P°PSPSC-C... 6 PjC-CÍCĤC†C-PI.

**P·P·P°C”PjPsPrC-Cl P· C-PSCëPëPjPë P»C-PeP°CĤCÍCHPePëPjPë P·P°CÍPsP±P°PjPë C,P° C-PSCëC- PIPëPrPë PIP·P°C”PjPsPrC-PNë.**

PĥPSC,PëC,CĤPsPjP±PsC,PëC†PSC- P·P°CÍPsP±Pë. PíPëP»PsCÍC,P°P·PsP» C” C-PSPiC-P±C-C,PsCĤPsPj C,,PsCÍC,,PsPrC-PµCÍC,PµCĤP°P·Pë P†P†P† P· P°PSC,PëC,CĤPsPjP±PsC†PëC,P°CĤPSPsCĤ P°PeC,PëPIPsC-CÍC,CĤ. P™ PsPiPs P·P°CÍC,PsCÍCfPIP°PSPSCĤ P·PrPsCĤPsPIPëPj PrPsP±CĤPsPIPsP»CHC†ClPj Cf PrPsP·C- 150 PjPi PíCĤPsC,ClPiPsPj 5 PrPSC-PI PSPµ PíCĤPëP·PIPsPrPëP»Ps PrPs PíPsPrPsPIP¶PµPSPSCĤ C†P°CÍCf PeCĤPsPIPsC,PµC±C-.

PĥC†PµC,PëP»CÍP°P»C-C†PëP»PsPIP° PePëCÍP»PsC,P° (PĥPŸPľ). PŸCfPjC-CÍPSPµ P·P°CÍC,PsCÍCfPIP°PSPSCĤ P· PĥPŸPľ PíCĤPsC,ClPiPsPj PePsCĤPsC,PePsPiPs C†P°CÍCf (PrPs 4 PrPSC-PI) P±CfP»Ps PíPsPíPŸ™ ClP·P°PSPµ P· PíC-PrPIPëC%PµPSPSCĤPj PSP° 23-25 % PíCĤPëPiPSC-C†PµPSPSCĤ PĥP”Pα-C-PSPrCfPePsPIP°PSPSC— P°PiCĤPµPiP°C†C-C— C,CĤPsPjP±PsC†PëC,C-PI PíPsCĤC-PIPsCĤPSPs P· PíCĤPëPNëPsPjPsPj C,C-P»CHPePë PĥPŸPľ. PkPµ CÍPíPsCÍC,PµCĤC-PiP°P»PsCÍCĤ PsC†PµPIPëPrPSPëC... C,PµPSPPrPµPSC†C-PNë PrPs P·P±C-P»CHCëPµPSPSCĤ CĤC-PIPsCĤ PiPµPjPsCĤP°PiC-C†PSPëC... PíPsP±C-C†PSPëC... PµC,,PµPeC,C-PI Cf PíP°C†C-C”PSC,C-PI, ClPeC- PíCĤPëPNëPjP°P»Pë PĥPŸPľ C,P° C†PëP»PsCÍC,P°P·PsP», PíPsCĤC-PIPsCĤPSPs P· PíP°C†C-C”PSC,P°PjPë, ClPeC- PíCĤPëPNëPjP°P»Pë PíP»P°C†PµP±Ps C,P° PµPePIC-PIP°P»PµPSC,PSC- PrPsP·Pë PĥPŸPľ.

PľP»PsPíC-PrPsPíCĤPµP»CH C,P° C-PSCëC- P°PSC,PëC,CĤPsPjP±PsC†PëC,P°CĤPSC-

P·P°CÍPsP±Pë. PŸCfPjC-CÍPSPëPNë PíCĤPëPNëPsPj C†PëP»PsCÍC,P°P·PsP»Cf C,P°  
PeP»PsPíC-PrPsPíCĤPµP»CĤ PSPµ PIPiP»PëPIP°PI PSP° PeC-P»CHPeC-CÍC,CH  
C,CĤPsPjP±PsC†PëC,C-PI, PíCĤPsC,CĤPsPjP±C-PSPsPIPëPNë C†P°CÍ (PµPŞ) P°P±Ps  
P°PeC,PëPIPsPIP°PSPëPNë C†P°CÍC,PePsPIPëPNë C,CĤPsPjP±PsPíP»P°CÍC,PëPSPsPIPëPNë  
C†P°CÍ (PĥPŞPŸPŞ). PJCÍC- P·PrPsCĤPsPIC- PrPsP±CĤPsPIPsP»CHC†C- PjP°P»Pë  
PíPsPrPsPIPµPSPëPNë C†P°CÍ PeCĤPsPIPsC,PµC†C- PíCĤPë PíCĤPëPNëPsPjC-  
PeP»PsPíC-PrPsPíCĤPµP»CĤ CµPe PjPsPSPsC,PµCĤP°PíC-C— C,P° PíCĤPë  
CÍCfPjC-CÍPSPsPjCf P·P°CÍC,PsCÍCfPIP°PSPSC- P· C†PëP»PsCÍC,P°P·PsP»PsPj, C%Ps  
PSPµ PíCĤPëP·PIPsPrPëP»Ps PrPs P·PSP°C†PSPsPiPs CÍCfPjP°CĤPSPsPiPs PIPiP»PëPICf  
PSP° C†P°CÍ PeCĤPsPIPsC,PµC†C-. PµCĤPsC,Pµ PíPsC,CĤC-P±PSPs  
PrPsC,CĤPëPjCfPIP°C,PëCÍCµ PsP±PµCĤPµPµPSPsCÍC,C- PíCĤPë  
PePsPjP±C-PSPsPIP°PSPsPjCf P·P°CÍC,PsCÍCfPIP°PSPSC- C†PëP»PsCÍC,P°P·PsP»Cf P·  
P±CfPrCH-CµPePëPjPë P°PSC,PëC,CĤPsPjP±PsC,PëC†PSPëPjPë P·P°CÍPsP±P°PjPë.  
PŸP»C-Pr CĤPsP·PiP»CµPSCfC,Pë PjPsPµP»PëPIC-CÍC,CH PíPµCĤC-PsPrPëC†PSPsPiPs  
PjPsPSC-C,PsCĤPëPSPiCf C†P°CÍCf PeCĤPsPIPsC,PµC†C-. P>C-PeCfPIP°PSPSCµ  
C†PëP»PsCÍC,P°P·PsP»PsPj PíCĤPsC,PëPíPsPeP°P·P°PSPs PíP°C†C-C”PSC,P°Pj, CµPeC-  
PíCĤPëPNëPjP°CĤC,CH PrPIP° P°P±Ps P±C-P»CHC€Pµ PrPsPrP°C,PePsPIPëC...  
P°PSC,PëC,CĤPsPjP±PsC†PëC,P°CĤPSPëC.../P°PSC,PëPePsP°PíCfP»CµPSC,PSPëC...  
P·P°CÍPsP±C-PI. P’C-P»CHC€ PIPëCÍPsPeP° C†P°CÍC,PsC,P° PIPëPíP°PrPeC-PI  
PeCĤPsPIPsPIPëP»PëPIC-PI CÍPíPsCÍC,PµCĤC-PiP°P»P°CÍCµ PíCĤPë  
PsPrPSPsC†P°CÍPSPsPjCf P·P°CÍC,PsCÍCfPIP°PSPSC- PeP»PsPíC-PrPsPíCĤPµP»CĤ,  
PĥPŸPŞ C- C†PëP»PsCÍC,P°P·PsP»Cf PI PíCĤPsC†PµCÍC- PrPsCÍP»C-PrPµPSPSCµ  
CASTLE.

PµPµCĤPsCĤP°P»CHPSC- P°PSC,PëPePsP°PíCfP»CµPSC,Pë (PSP°PíCĤPëPeP»P°Pr,  
PIP°CĤC,,P°CĤPëPS). PµCĤPë PsPrPSPsCĤP°P·PsPIPsPjCf PíCĤPëPNëPsPjC- PSPµ P±CfP»Ps  
PIPëCµPIP»PµPSPs PíCĤPëPiPSC-C†PµPSPSCµ PjPµC,P°P±PsP»C-P·PjCf PIP°CĤC,,P°CĤPëPSCf  
P°P±Ps PIPiP»PëPICf PSP° PíP°CĤP°PjPµC,CĤPë PePsP°PíCfP»CµC†C-C— (PµPŞ, PĥPŞPŸPŞ,  
C†P°CÍ PeCĤPsPIPsC,PµC†C-). PµCĤPsC,Pµ CĤPµPePsPjPµPSPPrPsPIP°PSPs  
PrPsC,CĤPëPjCfPIP°C,PëCÍCµ PsP±PµCĤPµPµPSPsCÍC,C- PíP°C†C-C”PSC,P°Pj, CµPeC-  
PíCĤPëPNëPjP°CĤC,CH C†PëP»PsCÍC,P°P·PsP» P· P±CfPrCH-CµPePëPj  
P°PSC,PëPePsP°PíCfP»CµC†C-PNëPSPëPj P·P°CÍPsP±PsPj, C,P° PíCĤPsPIPsPrPëC,Pë  
PíPµCĤC-PsPrPëC†PSPëPNë PjPsPSC-C,PsCĤPëPSPi PrP»Cµ PjC-PSC-PjC-P·P°C†C-C—  
PjPsPµP»PëPIPsCÍC,C- PeCĤPsPIPsC,PµC†C-. P>C-PeCfPIP°PSPSCµ C†PëP»PsCÍC,P°P·PsP»PsPj  
PíCĤPsC,PëPíPsPeP°P·P°PSPs PíP°C†C-C”PSC,P°Pj, CµPeC- PíCĤPëPNëPjP°CĤC,CH PrPIP° P°P±Ps  
P±C-P»CHC€Pµ PrPsPrP°C,PePsPIPëC...  
P°PSC,PëC,CĤPsPjP±PsC†PëC,P°CĤPSPëC.../P°PSC,PëPePsP°PíCfP»CµPSC,PSPëC...  
P·P°CÍPsP±C-PI.

P†PSPiC-P±C-C,PsCĤPë C†PëC,PsC...CĤPsPjCf P 450 (CYP). PíPëP»PsCÍC,P°P·PsP»  
P·PSP°C†PSPsCĤ PjC-CĤPsCĤ PjPµC,P°P±PsP»C-P·CfC”C,CHCÍCµ C,,PµCĤPjPµPSC,P°PjPë CYP,  
PsCÍPsP±P»PëPIPs CYP3A4 C,P° CYP2C19, C- PjPµPSC€PsCĤ PjC-CĤPsCĤ BĤ” CYP1A2.  
P”PµPiC-PrCĤPsPjPµC,P°P±PsP»C-C,, CµPePëPNë PjP°C” P°PSC,PëC,CĤPsPjP±PsC†PëC,P°CĤPSCf  
P°PeC,PëPIPsC-CÍC,CH Cf 4-7 CĤP°P·C-PI PIPëC%óCf, PSC-Pµ C†PëP»PsCÍC,P°P·PsP»,  
C-PjPsPIC-CĤPSPs, CfC,PIPsCĤCĤC”C,CHCÍCµ PiPsP»PsPIPSPëPj C†PëPSPsPj PíC-Pr PrC-C”CĤ  
CYP3A4. 4'-C,CĤP°PSCÍ-PiC-PrCĤPsPeCÍPëPjPµC,P°P±PsP»C-C, P· P°PeC,PëPIPsC-CÍC,CĤ,  
C%Ps PrPsCĤC-PIPsCĤC” 1/5 P°PeC,PëPIPsCÍC,C- C†PëP»PsCÍC,P°P·PsP»Cf,  
C-PjPsPIC-CĤPSPs,B CfC,PIPsCĤCĤC”C,CHCÍCµ P·P° PrPsPíPsPjPsPiPsCĤ CYP2C19.  
PŸP°PePëPj C†PëPSPsPj, P·P°CÍPsP±Pë, C%Ps PíCĤPëPiPSC-C†CfCĤC,CH CYP3A4

(PSP°PíCĤPëPeP»P°Pr, PrPµCĬPeC- PjP°PeCĤPsP»C-PrPë, P°P·PsP»PsPIC-  
PíCĤPsC,PëPíCĤPëP±PePsPIC- P·P°CíPSP±Pë, C-PSPiC-P±C-C,PsCĤPë PíCĤPsC,PµP°P·) P°P±Ps  
CYP2C19 (PSP°PíCĤPëPeP»P°Pr, C-PSPiC-P±C-C,PsCĤPë PíCĤPsC,PsPSPSPsPiPs PSP°CíPSPíP°),  
PíC-PrPIPëC%CrCĤC,CH P·P°PiP°P»CHPSCr C,,P°CĤPjP°PePsP»PsPiC-C±PSCr  
P°PeC,PëPIPSC-CíC,CH PSP° 32 % C,P° 42 % PIC-PrPíPSPIC-PrPSPs C,P° PjPsP¶CíC,CH  
PíC-PrPIPëC%CrPIP°C,Pë PíPSP±C-C±PSC- PµC,,PµPeC,Pë C†PëP»PsCíC,P°P·PsP»Cr. PĤPsP¶Pµ  
P±CíC,Pë PSPµPsP±C...C-PrPSPëPj P·PSPëP¶PµPSPSCĬ PrPsP·Pë C†PëP»PsCíC,P°P·PsP»Cr PrPs  
50 PjPi 2 CĤP°P·Pë PSP° PrPsP±Cí P·P°P»PµP¶PSPs PIC-Pr C-PSPPrPëPIC-PrCíP°P»CHPSPsC-  
PµC,,PµPeC,PëPIPSPsCíC,C- C,P° PíPµCĤPµPSPsCíPëPjPsCíC,C-.

PµCĤPëPµPsPj 100 PjPi C†PëP»PsCíC,P°P·PsP»Cr PSP° 7-PNë PrPµPSCĤ  
P·P°CíC,PsCíCíPIP°PSPSCĬ PµCĤPëC,CĤPsPjC-C†PëPSCr (PíPSPjC-CĤPSPëPNë  
C-PSPiC-P±C-C,PsCĤ CYP3A4) 500 PjPi 3 CĤP°P·Pë PSP° PrPsP±Cí PíCĤPëP·PIPSPPrPëP»Ps PrPs  
PíC-PrPIPëC%PµPSPSCĬ AUC C†PëP»PsCíC,P°P·PsP»Cr PrPs 74 %, CĬPePµ  
CíCíPíCĤPsPIPsPrP¶CíPIP°P»PsCíCĬ P·PSPëP¶PµPSPSCĬPj PSP° 24 % AUC PNëPsPiPs  
PrPµPiC-PrCĤPsPjPµC,P°P±PsP»C-C,Cr, P°P»Pµ P· PíPSPjC-C,PSPëPj PíC-PrPIPëC%PµPSPSCĬPj  
AUC 4'-C,CĤP°PSCí-PíC-PrCĤPsPeCíPëPjPµC,P°P±PsP»C-C,Cr. P— CíCĤP°C...CíPIP°PSPSCĬPj  
P·PSP°C±PµPSCĤ AUC P·P°PiP°P»CHPSP° C,,P°CĤPjP°PePsP»PsPiC-C±PSP°  
P°PeC,PëPIPSC-CíC,CH C†PëP»PsCíC,P°P·PsP»Cr P·P±C-P»CHC€CíC''C,CHCíCĬ PSP° 34 %  
PíCĤPë PsPrPSPsC±P°CíPSPsPjCr PNëPsPiPs PíCĤPëPNëPsPjC- P· PµCĤPëC,CĤPsPjC-C†PëPSPsPj.  
PáP° PíC-PrCíC,P°PIC- C†PëC... PrP°PSPëC... CĤPµPePsPjPµPSPPrPsPIP°PSP° PrPsP·P°  
C†PëP»PsCíC,P°P·PsP»Cr CíC,P°PSPsPIPëC,CH 50 PjPi 2 CĤP°P·Pë PSP° PrPsP±Cí PíCĤPë  
PsPrPSPsC±P°CíPSPsPjCr PíCĤPëPNëPsPjC- P· PµCĤPëC,CĤPsPjC-C†PëPSPsPj C,P° P·  
PíPSPPrC-P±PSPëPjPë P»C-PeP°CĤCíCĤPePëPjPë P·P°CíPSP±P°PjPë (PSP°PíCĤPëPeP»P°Pr, C-P·  
PeP»P°CĤPëC,CĤPsPjC-C†PëPSPsPj).

PŸCíPjC-CíPSPëPNë PíCĤPëPNëPsPj PsPrPSPsCĤP°P·PsPIPëC... PrPsP· PePµC,PsPePsPSP°P·PsP»Cr  
(CíPëP»CHPSPëPNë C-PSPiC-P±C-C,PsCĤ CYP3A4) 400 PjPi C,P° C†PëP»PsCíC,P°P·PsP»Cr 100  
PjPi PíCĤPëP·PIPSPPrPëPi PrPs PíC-PrPIPëC%PµPSPSCĬ AUC C†PëP»PsCíC,P°P·PsP»Cr PSP° 117  
, CĬPePµ CíCíPíCĤPsPIPsPrP¶CíPIP°P»PsCíCĤ P·PSPëP¶PµPSPSCĬPj PSP° 15 % AUC  
PrPµPiC-PrCĤPsPjPµC,P°P±PsP»C-C,Cr C,P° PíC-PrPIPëC%PµPSPSCĬPj PSP° 87 % AUC 4'-  
C,CĤP°PSCí-PíC-PrCĤPsPeCíPëPjPµC,P°P±PsP»C-C,Cr. P— CíCĤP°C...CíPIP°PSPSCĬPj  
P·PSP°C±PµPSCĤ AUC P·P°PiP°P»CHPSP° C,,P°CĤPjP°PePsP»PsPiC-C±PSP°  
P°PeC,PëPIPSC-CíC,CH C†PëP»PsCíC,P°P·PsP»Cr P·P±C-P»CHC€CíC''C,CHCíCĬ PSP°  
35 % PíCĤPë PsPrPSPsC±P°CíPSPsPjCr P·P°CíC,PsCíCíPIP°PSPSC- P·  
PePµC,PsPePsPSP°P·PsP»PsPj. PáP° PíC-PrCíC,P°PIC- C†PëC... PrP°PSPëC...  
CĤPµPePsPjPµPSPPrPsPIP°PSP° PrPsP·P° C†PëP»PsCíC,P°P·PsP»Cr CíC,P°PSPsPIPëC,CH  
50 PjPi 2 CĤP°P·Pë PSP° PrPsP±Cí PíCĤPë PsPrPSPsC±P°CíPSPsPjCr PíCĤPëPNëPsPjC- P·  
PePµC,PsPePsPSP°P·PsP»PsPj C,P° P· PíPSPPrC-P±PSPëPjPë P·P°CíPSP±P°PjPë  
(PSP°PíCĤPëPeP»P°Pr, P· C-C,CĤP°PePsPSP°P·PsP»PsPj).

P—P°CíC,PsCíCíPIP°PSPSCĬ 100 PjPi C†PëP»PsCíC,P°P·PsP»Cr 2 CĤP°P·Pë PSP° PrPsP±Cí P·  
PrPëP»C,C-P°P·PµPjPsPj (CíP»P°P±PePëPNë C-PSPiC-P±C-C,PsCĤ CYP3A4) 180 PjPi 1 CĤP°P·  
PSP° PrPsP±Cí PíCĤPëP·PIPSPPrPëP»Ps PrPs PíC-PrPIPëC%PµPSPSCĬ AUC  
C†PëP»PsCíC,P°P·PsP»Cr PSP° 44 %, CĬPePµ CíCíPíCĤPsPIPsPrP¶CíPIP°P»PsCíCĬ  
P·P±C-P»CHC€PµPSPSCĬPj AUC PrPµPiC-PrCĤPsPjPµC,P°P±PsP»C-C,Cr PSP° 4 % C,P°  
P·P±C-P»CHC€PµPSPSCĬPj AUC 4'-C,CĤP°PSCí-PíC-PrCĤPsPeCíPëPjPµC,P°P±PsP»C-C,Cr PSP°  
43 %. P— CíCĤP°C...CíPIP°PSPSCĬPj P·PSP°C±PµPSCĤ AUC P·P°PiP°P»CHPSP°  
C,,P°CĤPjP°PePsP»PsPiC-C±PSP° P°PeC,PëPIPSC-CíC,CH C†PëP»PsCíC,P°P·PsP»Cr  
P·P±C-P»CHC€CíC''C,CHCíCĬ PSP° 19 % PíCĤPë PsPrPSPsC±P°CíPSPsPjCr PíCĤPëPNëPsPjC- P·

PrPëP»C,C-P°P·PµPjPsPj. P“CḡCfPSC,CfCḡCfPëCfCḡ PSP° C†PëC... PrP°PSPëC...,  
PePsCḡPëPiCfPIP°PSPSCḡ PrPsP·Pë PSPµ PîPsC,CḡC-P±PSPs.

PµCḡPëPNëPsPj PsPrPSPsCḡP°P·PsPIPSC— PrPsP·Pë 100 PjPi C†PëP»PsCfC,P°P·PsP»Cf P· 240 PjP»  
PiCḡPµPNëPiC,,CḡCfC,PsiPisPiPs CfPsPeCf (C-PSPiC-P±C-C,PsCḡ PePëCëPePsPisPiPs CYP3A4)  
PSPµ PIPëCḡPIP»CḡPI PîPsPjC-C,PSPSiPs PµC,,PµPeC,Cf PSP°  
C,,P°CḡPjP°PePsPeC-PSPµC,PëPeCf C†PëP»PsCfC,P°P·PsP»Cf. Pj P·PiBḡ™ CḡP·PeCf P· C†PëPj  
PePsCḡPëPiCfPIP°PSPSCḡ PrPsP·Pë PSPµ PîPsC,CḡC-P±PSPs. PµCḡPsC,Pµ PIPḡPëPIP°PSPSCḡ  
PIPµP»PëPePëC... PsP±CfCḡPiC-PI PiCḡPµPNëPiC,,CḡCfC,PsiPisPiPs CfPsPeCf PjPsPḡPµ  
PjP°C,Pë PeP»C-PSC-C†PSPs P·PSP°C†CfC°PëPNë PIPiP»PëPI PSP°  
C,,P°CḡPjP°PePsPeC-PSPµC,PëPeCf C†PëP»PsCfC,P°P·PsP»Cf.

P—P°CfC,PsCfCfPIP°PSPSCḡ C†PëP»PsCfC,P°P·PsP»Cf P· PsPjPµPiCḡP°P·PsP»PsPj  
(C-PSPiC-P±C-C,PsCḡPsPj CYP2C19) PiCḡPëP·PIPsPrPëP»Ps PrPs P·P±C-P»CḡCëPµPSPSCḡ AUC  
C†PëP»PsCfC,P°P·PsP»Cf PSP° 22 %, CḡPePµ CfCfPiCḡPsPisPrPḡCfPIP°P»PsCfCḡ  
P·P±C-P»CḡCëPµPSPSCḡPj AUC PrPµPiC-PrCḡPsPjPµC,P°P±PsP»C-C,Cf PSP° 68 % C-  
P·PSPëPḡPµPSPSCḡPj AUC 4'C,CḡP°PSCf-PiC-PrCḡPsPeCfPëPjPµC,P°P±PsP»C-C,Cf PSP° 36 %.  
P— CfCḡP°C...CfPIP°PSPSCḡPj P·PSP°C†PµPSCḡ AUC P·P°PiP°P»CḡPSP°  
C,,P°CḡPjP°PePsP»PsPiC-C†PSP° P°PeC,PëPIPSC-CfC,Cḡ C†PëP»PsCfC,P°P·PsP»Cf  
P·P±C-P»CḡCëCfC”C,CḡCfCḡ PSP° 47 % PiCḡPë PsPrPSPsC†P°CfPSPsPjCf PiCḡPëPNëPsPjC- P·  
PsPjPµPiCḡP°P·PsP»PsPj. Pk° PiC-PrCfC,P°PIC- C†PëC... PrP°PSPëC...  
CḡPµPePsPjPµPSPPrPsPIP°PSP° PrPsP·P° C†PëP»PsCfC,P°P·PsP»Cf CfC,P°PSPsPIPëC,Cḡ 50 PjPi 2  
CḡP°P·Pë PSP° PrPsP±Cf PiCḡPë PsPrPSPsC†P°CfPSPsPjCf PiCḡPëPNëPsPjC- P·  
PsPjPµPiCḡP°P·PsP»PsPj.

PŸCáP±CfC,CḡP°C,Pë C,,PµCḡPjPµPSC,Cá C†PëC,PsC...CḡPsPjCá P 450. P°CfP»Ps  
PIC-PrP·PSP°C†PµPSPs PiC-PrPIPëC°PµPSPSCḡ C†PëP»PsCfC,P°P·PsP»PsPj AUC  
P»PsPIP°CfC,P°C,PëPSCf (C†CfC,P»PëPIPëPNë CfCfP±CfC,CḡP°C, CYP3A4) C,P° PNëPsPiPs  
β-PiC-PrCḡPsPeCfPëPePëCfP»PsC,Pë PrPs 70 %. PŸP»C-Pr PrPsC,CḡPëPjCfPIP°C,PëCfCḡ  
PsP±PµCḡPµPḡPSPsCfC,C- PiCḡPë CfCfPjC-CfPSPsPjCf PiCḡPëPNëPsPjC-  
C†PëP»PsCfC,P°P·PsP»Cf P·C- CfCfP±CfC,CḡP°C,P°PjPë CYP3A4 P· PICfP·CḡPePëPj  
C,PµCḡP°PiPµPIC,PëC†PSPëPj C-PSPPrPµPeCfPsPj (C,P°PePëPjPë CḡPe C†PëP·P°PiCḡPëPr,  
PiP°P»PsC,,P°PSC,CḡPëPS, PiC-PjPsP·PëPr, PîPsC...C-PrPSC- CḡC-PḡPeC-PI). PµPsC,CḡC-P±PSPs  
PrPsC,CḡPëPjCfPIP°C,PëCfCḡ PsP±PµCḡPµPḡPSPsCfC,C- PiCḡPë PsPrPSPsC†P°CfPSPsPjCf  
PiCḡPëPNëPsPjC- C†PëP»PsCfC,P°P·PsP»Cf P·C- CfC,P°C,PëPSP°PjPë, C°P°Ps  
PjPµC,P°P±PsP»C-P·CfCḡC,CḡCfCḡ C,,PµCḡPjPµPSC,P°PjPë C†PëC,PsC...CḡPsPjCf CYP3A4,  
PSP°PiCḡPëPeP»P°Pr P·C- CfPëPjPIP°CfC,P°C,PëPSPsPj, P°C,PsCḡPIP°CfC,P°C,PëPSPsPj C-  
P»PsPIP°CfC,P°C,PëPSPsPj.

P†PSPrCáPeC,PsCḡPë C,,PµCḡPjPµPSC,Cá C†PëC,PsC...CḡPsPjCá P 450. P°PiP»PëPI  
C-PSPPrCfPeC,PsCḡC-PI CYP3A4 C,P° CYP2C19 (C,P°PePëC... CḡPe PeP°CḡP±P°PjP°P·PµPiC-PS,  
C,,PµPSC-C,PsC—PS, CḡPëC,,P°PjPiC-C†PëPS C,P° PiCḡPµPiP°CḡP°C,Pë P·PIC-CḡPsP±PsCḡ)  
PSP° C,,P°CḡPjP°PePsPeC-PSPµC,PëPeCf C†PëP»PsCfC,P°P·PsP»Cf PSPµ P±CfPI  
PrPsCfP»C-PrPḡPµPSPëPNë. PŸPµPsCḡPµC,PëC†PSPs P°PSC,PëC,CḡPsPjP±PsC,PëC†PSP° PrC-Cḡ  
PjPsPḡPµ P±CfC,Pë P·PjC-PSPµPSP°, C,PsPjCf PSPµPsP±C...C-PrPSPs PiCḡPsPisPrPëC,Pë  
PjPsPSC-C,PsCḡPëPSPi Cf CḡP°P·C- P·P°CfC,PsCfCfPIP°PSPSCḡ C†PëP»PsCfC,P°P·PsP»Cf P·  
C-PSPPrCfPeC,PsCḡP°PjPë CYP3A4 C,P° CYP2C19.

PµC-Pr C†P°Cf PrPsCfP»C-PrPḡPµPSCḡ C,CḡC,CḡPSPsPiP°P»C-PSPSCḡ (CḡPePµ  
C-PSPPrCfPeCfC” CYP1A2) P·PSPëPḡCfPIP°P»Ps PePsPSC†PµPSC,CḡP°C†C-Cḡ  
C†PëP»PsCfC,P°P·PsP»Cf Cf PiP»P°P·PjC- PeCḡPsPIC- PSP° 18 %.



P†PSC€C- PìPsC, PµPSC†C- PN₆PSC- PIP·P°C”PjPsPrC-C—

PŸP»C-Pr PrPsC, CĤPĕPjC†PIP°C, PĕC†CĬ PsP±PµCĤPµP¶PSPsC†C, C- PìCĤPĕ  
PsPrPSPsC†P°C†PSPsPjC† P·P°C†C, PsC†C†PIP°PSPSC- C†PĕP»PsC†C, P°P·PsP»C† P· P±C†PrCĤ-  
CĬPePĕPj C-PSC€PĕPj P·P°C†PsP±PsPj, P·PrP°C, PSPĕPj P·PSPĕP¶C†PIP°C, Pĕ  
P°CĤC, PµCĤC-P°P»CĤPSPĕPN₆ C, PĕC†Pe, C†PµCĤPµP· PjPsP¶P»PĕPIC-C†C, CĤ  
CĤPsP·PIPĕC, PeC† P°PrPĕC, PĕPIPSPsPiPs PiC-PìPsC, PµPSP·PĕPIPSPsPiPs PµC,, PµPeC, C†, C%₀Ps  
C†C†PìCĤPsPIPsPrP¶C†C”C, CĤC†CĬ CĤPµC,, P»PµPeC, PsCĤPSPsCĤ C, P°C...C-PeP°CĤPrC-C”CĤ.

**PĥC†PsP±P»PĕPIP sC†C, C- P·P°C†C, PsC†CžPIP°PSPSCĬ.**

PµCĤPĕPN₆PSCĬC, PSC-C†C, CĤ P»C-PeC†PIP°PSPSCĬ C†PĕP»PsC†C, P°P·PsP»PsPj  
C†P»C-Pr CĤPµC, PµP»CĤPSPs PsC†C-PSCĤPIP°C, Pĕ PìPsCĤCĬPr P· C-PSC€PĕPjPĕ  
PIP°CĤC-P°PSC, P°PjPĕ P»C-PeC†PIP°PSPSCĬ, C, P°PePĕPjPĕ CĬPe  
CĤPµPIP°C†PeC†P»CĬCĤPĕP·P°C†C-CĬ.

P’ C†PĕP»C† C†PIP sPiPs PjPµC...P°PSC-P·PjC† PrC-C— C†PĕP»PsC†C, P°P·PsP» PjPsP¶Pµ  
C†PìCĤPĕC†PĕPSPĕC, Pĕ C, P°C...C-PeP°CĤPrC-CĤ, PIC-PrC†C†C, C, CĬ C†PµCĤC†PµP±PĕC, C, CĬ,  
C, P°C...C-P°CĤPĕC, PjC-CĤ C, P°/P°P±Ps P°CĤC, PµCĤC-P°P»CĤPSC† PiC-PìPsC, PµPSP·C-CĤ.  
P—P±C-P»CĤC€PµPSPSCĬ C†P°C†C, PsC, Pĕ C†PµCĤC†PµPIPĕC... C†PePsCĤPsC†PµPSCĤ,  
P°C†PsC†C-PN₆PsPIP°PSPµ P· C†PĕP»PsC†C, P°P·PsP»PsPj, C†C, P°PSPsPIPĕC, CĤ  
PìCĤPĕP±P»PĕP·PSPs PIC-Pr 5 PrPs 7 C†PrP°CĤC-PI PSP° C...PIPĕP»PĕPSC†, C, PsPjC† C†  
PìP°C†C-C”PSC, C-PI C-P· CĤPĕP·PĕPePsPj PrP°PSPsPiPs CĬPIPĕC%₀P° PìCĤPµPìP°CĤP°C,  
PjPsP¶Pµ C†PìCĤPĕC†PĕPSCĬC, Pĕ C†C, PµPSPsPeP°CĤPrC-CĤ.

P—P°B PìP°C†C-C”PSC, P°PjPĕ, CĬPeC- PjPsP¶C†C, CĤ P±C†C, Pĕ C†C...PĕP»CĤPSC- PrPs  
PìC-PrPIPĕC%₀PµPSPsPiPs CĤPĕP·PĕPeC† C†PµCĤPµPN₆PsP·PSPĕC... PìPsP±C-C†PSPĕC...  
PµC,, PµPeC, C-PI P· P±PsPeC† C†PµCĤC†CĬ PI CĤPµP·C†P»CĤC, P°C, C-  
P·P±C-P»CĤC€PµPSPSCĬ C†P°C†C, PsC, Pĕ C†PµCĤC†PµPIPĕC... C†PePsCĤPsC†PµPSCĤ,  
PSP°PìCĤPĕPeP»P°Pr P·P° PìP°C†C-C”PSC, P°PjPĕ P·C- C†C, P°P±C-P»CĤPSPsCĤ  
C-C€PµPjC-C†PSPsCĤ C...PIP sCĤPsP±PsCĤ C†PµCĤC†CĬ, C†P»C-Pr CĤPµC, PµP»CĤPSPs  
C†PìPsC†C, PµCĤC-PiP°C, Pĕ PìC-Pr C†P°C† P»C-PeC†PIP°PSPSCĬ C†PĕP»PsC†C, P°P·PsP»PsPj;  
PìPsC, CĤC-P±PSPs PICĤP°C...PsPIC†PIP°C, Pĕ, C%₀Ps C†PĕP»PsC†C, P°P·PsP»  
PìCĤPsC, PĕPìPsPeP°P·P°PSPĕPN₆ PìP°C†C-C”PSC, P°Pj, CĬPeC- PìPµCĤPµPSPµC†P»Pĕ  
PSPµC†C, P°P±C-P»CĤPSC† C†C, PµPSPsPeP°CĤPrC-CĤ P°P±Ps C-PSC,, P°CĤPeC,  
PjC-PsPeP°CĤPrP°/PePsCĤPsPSP°CĤPSPµ PIC, CĤC†C†P°PSPSCĬ PìCĤPsC, CĬPiPsPj  
PsC†C, P°PSPSC-C... 6 PjC-C†CĬC†C-PI, C, P° PìP°C†C-C”PSC, P°Pj P· C, CĬP¶PePsCĤ  
C, P°C...C-P°CĤPĕC, PjC-C”CĤ PI P°PSP°PjPSPµP·C-.

PŸP»C-Pr PrPsC, CĤPĕPjC†PIP°C, PĕC†CĬ PsP±PµCĤPµP¶PSPsC†C, C- PìCĤPĕ  
PìCĤPĕP·PSP°C†PµPSPSC- C†PĕP»PsC†C, P°P·PsP»C† PìP°C†C-C”PSC, P°Pj P· PµPeC, PsPìC-C”CĤ  
PìPµCĤPµPrC†PµCĤPrCĬ C- C€P»C†PSPsC†PeC-PI, P° C, P°PePsP¶ PìP°C†C-C”PSC, P°Pj C-P·  
C,, C-P±CĤPĕP»CĬC†C-C”CĤ P°P±Ps C, CĤC-PìPsC, C-PSPSCĬPj PìPµCĤPµPrC†PµCĤPrCĬ.

PµP°C†C-C”PSC, C-PI PSPµPsP±C...C-PrPSPs PìPsPìPµCĤPµPrPĕC, Pĕ PìCĤPs  
PSPµPsP±C...C-PrPSC-C†C, CĤ P·PIPµCĤPSC†C, PĕC†CĬ PrPs P»C-PeP°CĤCĬ C† PIPĕPìP°PrPeC†  
PìPsCĬPIPĕ PeCĤPsPIPsC, PµC† P°P±Ps C†PĕPSC†C-PI PìC-Pr C†P°C† C, PµCĤP°PìC-C—. Pj  
CĤP°P·C- PìPsCĬPIPĕ PsC†PSPĕC... PeCĤPsPIPsC, PµC† PìCĤPĕPN₆PsPj C†PĕP»PsC†C, P°P·PsP»C†  
PSPµPsP±C...C-PrPSPs PìCĤPĕPìPĕPSPĕC, Pĕ.

PhCГPeC-P»CHPePe PиCTHPIPI°CTHP°C, P·PrP°C, PSPePNe PиCTHPePIpsc-C†CPIPI°C, Pe  
P°PiCTHPPIPI°CTC-CTH C, CTHPSPJ±PSc†Pec, C-PI, P·P±C-P»CHC€Crc”C, CHCГCИ CTHPeP·PePe  
PeCTHPSPIPSc, PИC† PиC-Pr C†P°CГ C...C-CTHCrcCTHPiC-C†PSPeC... PIC, CTHCrcC†P°PSCн  
(PIPeP»CTHC†P°CTHC†Pe PSPPI·PSP°C†PSC- PIC, CTHCrcC†P°PSPSCИ, PSP°PиCTHPePeP»P°Pr  
PIPePrP°P»PИPSPSCИ P·CPI±P°). PИPeC%Ps PиP°CTC-C”PSC, CГ PSPPIPs±C...C-PrPSPs  
PиCTHPSPIPИCГC, Pe C...C-CTHCrcCTHPiC-C†PSPИ PIC, CTHCrcC†P°PSPSCИ C-  
P°PSC, PeP°PiCTHPPIPI°CTC-PNеPSPePNe PИC,, PИPeC, C” PSPPI±P°P¶P°PSPePj, PиCTHPePNePsPj  
C†PeP»PScГC, P°P·PsP»CГ CГP»C-Pr PиCTHPePIPePSPeC, Pe P·P° 5 PrPSC-PI PrPs  
PsPIPICTHP°CTC-C—.

PkP°PrC...PsPrPeP»Pe PиPsPsPrPePSPsPeC- PиPsPIC-PrPsPjP»PИPSPSCИ PиCTHPs  
PiPIPIP°C, PsP»PsPiC-C†PSC- PIC-PrC...PeP»PИPSPSCИ, PIPeP»CTHC†P°CTHC†Pe  
C, CTHPSPJ±PSc†Pec, PsPIPIpsc-CTH, P»PИPNePePsPIPIpsc-CTH, P°PiCTHP°PSCrP»PSc†Pec, PsP·,  
PIP°PSC†Pec, PsPIPIpsc-CTH C, P° P°PIP»P°CГC, Pec†PScГ P°PSPPIjC-CTH. P°C-P»CHC€C-CTC, CH  
PIP°CTC-C”PSC, C-PI PIPePrCГP¶CГPIPI°P»Pe PиC-CГP»CИ PиCTHPePIPePSPPIPSPSCИ  
PиCTHPePNePsPjCГ C†PeP»PScГC, P°P·PsP»CГ. PИCTHPsC, PИ PrPIPeC-P»CHPeP° PIPePIP°PrPeC-PI  
PIP°PSC†Pec, PsPIPIpsc-C— C, P° P°PIP»P°CГC, Pec†PSPsC— P°PSPPIjC-C— PjP°P»Pe  
P»PIPI, P°P»CHPSPePNe PSP°CГP»C-PrPsPe.

PИP°CTC-C”PSC, C-PI CГP»C-Pr PиPsPIPICTHPPIPrPeC, Pe PиCTHPs PSPPIPs±C...C-PrPSC-CTC, CH  
PSPPIPI°PNePSPs PиPsPIC-PrPsPjP»CИC, Pe PиCTHPs P±CГPrCH-CTPeC- PsP·PSP°PePe, CTPeC-  
PjPsP¶CГC, CH CГPIC-PrC†Pec, Pe PиCTHPs CTHP°PSPSC-PNе CTHPSP·PIPeC, PsPe  
PIP°C, PsP»PsPiC-C†PSPeC... P·PjC-PS PeCTHPsPIC-, C, P°PePeC... CИPe PиC-PIPICTHC, PИCTHPjC-CИ  
C, P° P±C-P»CH CГ PиPsCTHP»C-. PkPIPsP±C...C-PrPSPs PиCTHPsPIPICTC, Pe PиPsPIPSPePNe  
P°PSP°P»C-P· PeCTHPsPIC-, CИPeC%Ps C” PиC-PrPsP·CTHP° PSP° C-PSC,, PИPeC†C-CTH P°P±Ps  
PSP°CИPIpsc- P±CГPrCH-CTPeC- C-PSC€C- PeP»C-PSC-C†PSC- PsP·PSP°PePe  
PIP°C, PsP»PsPiC-C†PSPeC... P·PjC-PS PeCTHPsPIC-. PИCTHPePNePsPj C†PeP»PScГC, P°P·PsP»CГ  
CГP»C-Pr PиCTHPePIPePSPeC, Pe, CИPeC%Ps C” PeP»C-PSC-C†PSC- P°P±Ps  
P»P°P±PsCTHP°C, PsCTHPsC- PrPsPeP°P·Pe PИP°C, PsP»PsPiC-C†PSPeC... P·PjC-PS PeCTHPsPIC-.

PJ PиP°CTC-C”PSC, C-PI, CИPeC- PиCTHPePNePjP°P»Pe PиPsC, CГP¶PSC- C-PSPiC-P±C-C, PsCTHPe  
CYP3A4 P°P±Ps CYP2C19, CГPIPsCГC, PИCTHC-PIP°P»PScГCИ P·P±C-P»CHC€PИPSPSCИ  
CTHC-PIpscИ C†PeP»PScГC, P°P·PsP»CГ CГ PиP»P°P·PjC- PeCTHPsPIC-. PJ C, P°PePeC...  
PIPePIP°PrPeP°C... CTHPИPePsPjPИPSPrPsPIPI°PSP° PrPsP·P° C†PeP»PScГC, P°P·PsP»CГ  
CГC, P°PSPsPIPeC, CH 50 PjPi 2 CTHP°P·Pe PSP° PrPsP±CГ.

PkPIPsP±C...C-PrPSP° PsP±PICTHPPI¶PSC-CTC, CH PиCTHPe CГCГPjC-CГPSPsPjCГ  
P·P°CГC, PsCГCГPIPI°PSPSC- C†PeP»PScГC, P°P·PsP»CГ P· C-PSPiC-P±C-C, PsCTHP°PjPe P°P±Ps  
C-PSPrCГPeC, PsCTHP°PjPe CYP 3A4, CYP 2C19 P°P±Ps CГCГP±CГC, CTHP°C, P°PjPe CYP 3A4.

PkPIPsP±C...C-PrPSPs P· PsP±PICTHPPI¶PSC-CTC, CH PиCTHPeP·PSP°C†P°C, Pe PиCTHPPIPI°CTHP°C,  
PIP°CTC-C”PSC, P°Pj C-P· PИPICTHPPIPrCГPICTHPPrPSPsCTH P°P±Ps C€P»CГPSPsC†PePsPIPsCTH  
PИPeC, PsPiC-C”CTH, C,, C-P±CTHPeP»CИC†C-C”CTH P°P±Ps C, CTHC-PиPsC, C-PSPSCИPj  
PIPICTHPPIPrCГPICTHPPrCH.

PkPIPsP±C...C-PrPSP° PsP±PICTHPPI¶PSC-CTC, CH PиCTHPe CГCГPjC-CГPSPsPjCГ  
PиCTHPePNePsPjC- C†PeP»PScГC, P°P·PsP»CГ P· P±CГPrCH-CTPePePjPe C-PSC€PePjPe  
P»C-PeP°CTHCГCHPePePjPe P·P°CГPsP±P°PjPe, C%Ps PjPsP¶CГC, CH P·PSPeP¶CГPIPI°C, Pe  
P°CTHC, PИCTHC-P°P»CHPSPePNe C, PecГPe, PsCГPeC-P»CHPePe C-CГPSCrc” CTHPeP·PePe  
P°PrPeC, PePIPSPsPIPs PiC-PиPsC, PИPSP·PePIPSPsPIPs PИC,, PИPeC, CГ P·  
CTHPИC,, P»PИPeC, PsCTHPSPsCTH C, P°C...C-PeP°CTHPrc-C”CTH.

РҮР»С-Рr PrPsC,СҲРөРјСrPIP°С,РөСГCЦ PsP±PμСҲРμP¶PSPsCГC,С- PиСҲРө  
PиСҲРөP·PSP°С±PμPSPSC- C†PөP»PsCГC,Р°P·PsP»Cр P· P±CрPrСҲ-СЦPөPөPjPө C-PSC€PөPjPө  
P°PSC,РөC,СҲPsPjP±PsC,РөC±PSPөPjPө P·P°CГPsP±P°PjPө.

P·C,,PμPөC, PiP°P»СҲРјСrPIP°PSPSCЦ PSP°PiP°PrCр C-PSCГCр»СҲC,Cр, C%oPs  
PиСҲРөPсЦPIP»CЦC” C†PμPNө PiСҲРμPiP°СҲР°C,, PSPμ PrPsCГP»C-PrP¶CрPIP°PICГCЦ PиСҲРө  
P°CГPөPjPиC,PsPjP°С,РөC±PSPsPjCр C-C€PμPjC-C±PSPsPjCр C-PSCГCр»СҲC,С-.

*P—P°CГC,PsCГCзPIP°PSPSCЦ Cз PiPμСҲC-PsPr PIP°PiC-C,PSPsCГC,С- P°P±Ps  
PiPsPrCзPIP°PSPSCЦ PiСҲCзPrPrСҲ.*

*P’P°PiC-C,PSC-CГC,СҲ*

P’C-PrCГCрC,PSC- PиC-PrC,PIPμСҲPrP¶PμPSC- PrP°PSC- C%oPsPrPs  
P·P°CГC,PsCГCрPIP°PSPSCЦ C†PөP»PsCГC,Р°P·PsP»Cр PIP°PiC-C,PSPөPj,  
PiPsC,PμPSC†C-PNөPSPөPNө СҲРөP·PөPө PSPμPIC-PrPsPjPөPNө. P”PsCГP»C-PrP¶PμPSPSCЦ PSP°  
C,PIP°СҲРөPSP°С... PiPsPөP°P·P°P»Pө СҲРμPiСҲPsPrCрPөC,РөPIPSCр  
C,PsPөCГPөC±PSC-CГC,СҲ. PμPsC,PμPSC†C-PNөPSPөPNө СҲРөP·PөPө PSPμPIC-PrPsPjPөPNө,  
C,PsPjCр C†PөP»PsCГC,Р°P·PsP» PSPμ PjPsP¶PSP° PиСҲРөPNөPjP°С,Рө Cр PiPμСҲC-PsPr  
PIP°PiC-C,PSPsCГC,С-.

*PμPμСҲC-PsPr PiPsPrCзPIP°PSPSCЦ PiСҲCзPrPrСҲ*

P”PsCГP»C-PrP¶PμPSPSCЦ PSP° C,PIP°СҲРөPSP°С... PiPsPөP°P·P°P»Pө, C%oPs  
C†PөP»PsCГC,Р°P·PsP» PjPsP¶Pμ PiСҲPsPSPөPөP°С,Рө Cр PiСҲCрPrPSPμ PjPsP»PsPөPs.  
PҮPsC±PSC- PrP°PSC- PиСҲРөP PиСҲPsPSPөPөPSPμPSPSCЦ C†PөP»PsCГC,Р°P·PsP»Cр Cр  
PиСҲCрPrPSPμ PjPsP»PsPөPs P¶C-PSPөPө PIC-PrCГCрC,PSC-. P’СҲР°C...PsPICрCҲC±Pө  
PjPsP¶P»PөPIPөPNө PSPμPiP°С,РөPIPSPөPNө PIPиP»PөPI PSP° PrPөC,РөPSCр,  
P·P°CГC,PsCГCрPIP°PSPSCЦ PиСҲРμPiP°СҲР°C,Cр Cр PiPμСҲC-PsPr PiPsPrCрPIP°PSPSCЦ  
PиСҲCрPrPrСҲ PSPμ СҲРμPөPsPjPμPSPrCрC”C,СҲCГCЦ. Pj СҲР°P·C-  
PSPμPsP±C...C-PrPSPsCГC,С- P»C-PөCрPIP°PSPSCЦ C†PөP»PsCГC,Р°P·PsP»PsPj CГP»C-Pr  
PиСҲРөPiPөPSPөC,Рө PiPsPrCрPIP°PSPSCЦ PиСҲCрPrPrСҲ.

*PαPμСҲC,PөP»СҲPSC-CГC,СҲ*

P”PsCГP»C-PrP¶PμPSPSCЦ PSP° C,PIP°СҲРөPSP°С... PiPsPөP°P·P°P»Pө, C%oPs  
C†PөP»PsCГC,Р°P·PsP» PSPμ PIPиP»PөPIP°С” PSP° C,,PμСҲC,РөP»СҲPSC-CГC,СҲ.

*P—PrP°С,PSC-CГC,СҲ PIPиP»PөPIP°С,Рө PSP° C€PIPөPrPөC-CГC,СҲ СҲРμP°PөC†C-C—  
PиСҲРө PөPμСҲCзPIP°PSPSC- P°PIC,PsC,СҲР°PSCГPиPsСҲC,PsPj P°P±Ps C-PSC€PөPjPө  
PjPμC...P°PSC-P·PjP°PjPө.*

РҮР»С-Рr PrPsC,СҲРөРјСrPIP°С,РөCГCЦ PsP±PμСҲРμP¶PSPsCГC,С-, PsCГPөC-P»СҲPөPө  
PиСҲРө PиСҲРөPNөPsPjC- PиСҲРμPiP°СҲР°C,Cр PjPsP¶P»PөPIPμ P·P°PiP°PjPsСҲPsC±PμPSPSCЦ.

***PҮPiPsCГC-P± P·P°CГC,PsCГCзPIP°PSPSCЦ C,P° PrPsP·Pө.***

*P”PsP·CзPIP°PSPSCЦ*

P PμPePsPjPμPSPPrPsPIP°PSP° PrPsP·P° PiCḡPμPiP°CḡP°C,Cr Bḡ“ PiPs 100 PjPi 2 CḡP°P·Pë PSP° PrPsP±Cr. PÿP°P±P»PμC,PePë PiCḡPëPNëPjP°CḡC,CH P·P° 30 C...PIPëP»PëPS PrPs PIP¶PëPIP°PSPSCḡ C—P¶C- P°P±Ps C±PμCḡPμP· 2 PiPsPrPëPSPë PiC-ClP»Cḡ PIP¶PëPIP°PSPSCḡ C—P¶C-, PICḡP°PSC†C- C,P° PIPμC±PμCḡC-.

PμCḡPëPNëPsPj PiCḡPμPiP°CḡP°C,Cr PiC-Pr C±P°Cl C—PrPë PjPsP¶Pμ PiC-PrPIPëC%CrPIP°C,Pë PNëPsPiPs PjP°PeClPëPjP°P»CHPSCr PePsPSC†PμPSC,CḡP°C†C-CH PI PiP»P°P·PjC- PeCḡPsPIC-, C%Ps P·P±C-P»CHC€CrC” CḡPëP·PëPe PIPëPSPëPePSPμPSPSCḡ PiPsP±C-C±PSPëC... CḡPμP°PeC†C-PNë. P—PSP°C±PSPμ PiPsPeCḡP°C%PμPSPSCḡ ClC,P°PSCr PiP°C†C-C”PSC,C-PI ClPiPsClC,PμCḡC-PiP°C”C,CHClCḡ PiC-ClP»Cḡ PiCḡPëPNëPsPjCr PiCḡPμPiP°CḡP°C,Cr PiCḡPsC,CḡPiPsPj 16-24 C,PëP¶PSC-PI, C-PSPPrC- PiPsPeCḡP°C%PμPSPSCḡ PIC-PrP·PSP°C±P°P»Pë PIP¶Pμ PiC-ClP»Cḡ P»C-PeCrPIP°PSPSCḡ PiCḡPsC,CḡPiPsPj 4-12 C,PëP¶PSC-PI. PiPeC%Ps PiCḡPsC,CḡPiPsPj 6 PjC-ClCḡC†C-PI P»C-PeCrPIP°PSPSCḡ PSPμ P±CrP»Ps PμC,,PμPeC,PëPIPSPëPj, P»C-PeP°Cḡ PiPsPIPëPSPμPS PiCḡPëP·PSP°C±PëC,Pë C-PSC€Cr C,PμCḡP°PiC-CH. PμP°C†C-C”PSC,P°Pj, CḡPeC- PsC,CḡPëPjCrCḡC,CH P»C-PeCrPIP°PSPSCḡ C†PëP»PsClC,P°P·PsP»PsPj, ClP»C-Pr PrPsC,CḡPëPjCrPIP°C,PëClCḡ P·PjC-PS Cr ClPiPsClPsP±C- P¶PëC,C,Cḡ (PIC-PrPjPsPIP° PIC-Pr PeCrCḡC-PSPSCḡ C,P° C,,C-P·PëC±PSC- PIPiCḡP°PIPë) C- PiCḡPsPrPsPIP¶CrPIP°C,Pë C,,P°CḡPjP°PePsP»PsPiC-C±PSC- PIC,CḡCrC±P°PSPSCḡ (PSP°PiCḡPëPeP»P°Pr, PiCḡPëPNëPsPj P»C-PiC-PrPsP·PSPëP¶CrPIP°P»CHPSPëC... C,P° P°PSC,PëC,CḡPsPjP±PsC†PëC,P°CḡPSPëC... P·P°ClPsP±C-PI), C%PsP± P·PjPμPSC€PëC,Pë CḡPëP·PëPe CḡPsP·PIPëC,PeCr ClPμCḡC†PμPIP- ClCrPrPëPSPSPëC... CḡPIPëC%. PiPëP»PsClC,P°P·PsP» PSPμ P·P°PjC-PSCḡC” PrP°PSC- PIPëPrPë P»C-PeCrPIP°PSPSCḡ.

P—PjPμPSC€PμPSPSCḡ PrPsP·Pë PrPs 50 PjPi 2 CḡP°P·Pë PSP° PrPsP±Cr CḡPμPePsPjPμPSPPrPsPIP°PSPμ PrP»Cḡ PiP°C†C-C”PSC,C-PI, CḡPeC- PsC,CḡPëPjCrCḡC,CH P»C-PeP°CḡClCHPeC- P·P°ClPsP±Pë, C%Ps ClPëP»CHPSPs C-PSPiC-P±CrCḡC,CH CYP3A4, PSP°PiCḡPëPeP»P°Pr PrPμCḡPeC- PjP°PeCḡPsP»C-PrPë, P°P·PsP»CHPSC- C,,CrPSPiC-C†PëPrPë, C-PSPiC-P±C-C,PSCḡPë PiCḡPsC,PμP°P·Pë, P°P±Ps P·P°ClPsP±Pë, CḡPeC- ClPëP»CHPSPs C-PSPiC-P±CrCḡC,CH CYP2C19, PSP°PiCḡPëPeP»P°Pr PsPjPμPiCḡP°P·PsP» (PrPëPI. CḡPsP·PrC-P» B«P’P·P°C”PjPsPrC-Cḡ P· C-PSC€PëPjPë P»C-PeP°CḡClCHPePëPjPë P·P°ClPsP±P°PjPë C,P° C-PSC€C- PIPëPrPë PIP·P°C”PjPsPrC-PNëB»).

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

PkPμPjP°C” PSPμPsP±C...C-PrPSPsClC,C- Cr PePsCḡPμPeC†C-C— PrPsP·Pë PrP»Cḡ C†C-C”C— PeP°C,PμPiPsCḡC-C— PiP°C†C-C”PSC,C-PI.

*PμP°C†C-C”PSC,Pë P· PSPëCḡPePsPIPsCḡ PSPμPrPsClC,P°C,PSC-ClC,CH*

P”P»Cḡ PiP°C†C-C”PSC,C-PI C-P· PeP»C-CḡPμPSClPsPj PeCḡPμP°C,PëPSC-PSCr > 25 PjP»/C...PI ClPiPμC†C-P°P»CHPSP° PePsCḡPμPeC†C-Cḡ PrPsP·CrPIP°PSPSCḡ PSPμ PiPsC,CḡC-P±PSP°. PiPëP»PsClC,P°P·PsP» PiCḡPsC,PëPiPsPeP°P·P°PSPëPNë PiP°C†C-C”PSC,P°Pj C-P· PeP»C-CḡPμPSClPsPj PeCḡPμP°C,PëPSC-PSCr ≤ 25 PjP»/C...PI.

*PμP°C†C-C”PSC,Pë P· PiPμC±C-PSPePsPIPsCḡ PSPμPrPsClC,P°C,PSC-ClC,CH*

P”P»Cḡ PiP°C†C-C”PSC,C-PI P· P»PμPiPePsCḡ C,,PsCḡPjPsCḡ P·P°C...PIPsCḡCḡPIP°PSPSCḡ PiPμC±C-PSPePë PrPsP·Cr PSPμ PiPsC,CḡC-P±PSPs P·PjC-PSCḡPIP°C,Pë. P’C-PrClCrC,PSC- PrP°PSC- C%PsPrPs PiP°C†C-C”PSC,C-PI P· PiPsPjC-CḡPSPsCḡ P°P±Ps C,CḡP¶PePsCḡ PiPμC±C-PSPePsPIPsCḡ PSPμPrPsClC,P°C,PSC-ClC,CH. PhClPeC-P»CHPePë C†PëP»PsClC,P°P·PsP» P°PeC,PëPIPSPs PjPμC,P°P±PsP»C-P·CrC”C,CHClCḡ

C,,PμCḡPjPμPSC,P°PjPë PīPμC‡C-PSPePë, PIC-PS PīCḡPsC,PëPīPsPeP°P°PSPëPNë  
PīP°C‡C-C" PSC,P°Pj P· PīPsPjC-CḡPSPsCḡ P°P±Ps C,CḡPḡPēPsCḡ PīPμC‡C-PSPePsPīPsCḡ  
PSPμPrPsCḡC,P°C,PSC-CḡC,Cḡ.

*PμP°C‡C-C" PSC,Pë P»C-C,PSCHḡPsPīPs PIC-PeCž*

PḡPμPjP°C" PSPμPsP±C...C-PrPSPsCḡC,C- PI PePsCḡPμPeC‡C-C— PrPsP·Pë PrP»Cḡ C‡C-C" C—  
PeP°C,PμPīPsCḡC-C— PīP°C‡C-C" PSC,C-PI.

*P" C-C,Pë.*

P»C-PeP°CḡCḡCḡPePëPNë P·P°CḡC-P± PSPμ CḡPμPePsPjPμPSPPrCḡC" C,CḡCḡCḡ  
PīCḡPëP·PSP°C‡P°C,Pë PrC-C,CḡPj C‡PμCḡPμP· PIC-PrCḡCḡC,PSC-CḡC,Cḡ PrP°PSPëC...  
C%oPsPrPs P±PμP·PīPμPePë C,P° PμC,,PμPeC,PëPīPSPsCḡC,C- PNëPsPīPs  
P·P°CḡC,PscḡCḡPīP°PSPSCḡ.

***PμPμCḡPμPrPsP·CžPIP°PSPSCḡ.***

P‡PSC,,PsCḡPjP°C‡C-Cḡ PīCḡPs PīPsCḡC,CḡPμ PīPμCḡPμPrPsP·CḡPIP°PSPSCḡ  
PsP±PjPμPḡPμPSP°. PḡPsPḡP»PëPIC- CḡPëP»CḡPSPëPNë PīPsP»PsPīPSPëPNë P±C-P»Cḡ,  
PrC-P°CḡPμCḡ, C,P°C...C-PeP°CḡPrC-Cḡ C,P° CḡPμCḡC‡PμPīP° P°CḡPëC,PjC-Cḡ. P—P°  
PīP°C‡C-C" PSC,P°PjPë PSPμPsP±C...C-PrPSPs CḡPīPsCḡC,PμCḡC-PīP°C,Pë C,P°  
PīCḡPsPīPsPrPëC,Pë PīC-PrC,CḡPëPjCḡPīP°P»CḡPSCḡ C,PμCḡP°PīC-Cḡ. PḡPμPsP±C...C-PrPSPs  
CḡPīPsCḡPsPḡPSPëC,Pë CḡP»CḡPSPsPe CḡP»CḡC...PsPj C-PSPrCḡPeC‡C-C—  
P±P»CḡPIP°PSPSCḡ P°P±Ps PīCḡPsPjPëPīP°PSPSCḡ CḡP»CḡPSPeP°.

***PμPsP±C-C‡PSC- CḡPμP°PeC‡C-C—.***

PḡP°PNëP±C-P»CḡCḡ PīPsCḡPëCḡPμPSC- PīPsP±C-C‡PSC- CḡPμP°PeC‡C-C—, PīCḡPs CḡPēC-  
PīPsPIC-PrPsPjP»CḡP»Pë Cḡ PeP»C-PSC-C‡PSPëC... PrPsCḡP»C-PrPḡPμPSPSCḡC...  
PīCḡPμPīP°CḡP°C,Cḡ, P±CḡP»Pë PīPsP»PsPīPSPëPNë P±C-P»Cḡ (>30 %), PrC-P°CḡPμCḡ (>15 %)  
C,P° PīPsCḡCḡCḡPμPSPSCḡ CḡPsP±PsC,Pë PePëCḡPμCḡPSPëPeCḡ (>15 %). PīC-  
CḡPμP°PeC‡C-C— P·P°P·PīPëC‡P°PNë PjP°P»Pë P»PμPīPeCḡ P°P±Ps PīPsPjC-CḡPSCḡ  
C-PSC,PμPSCḡPëPīPSC-CḡC,Cḡ C,P° C-PSPrC- PīPsP»PμPīCḡCḡPīP°P»PëCḡCḡ P·P°  
CḡP°C...CḡPSPsPe P·PSPëPḡPμPSPSCḡ PrPsP·Pë.

PḡPμP±P°PḡP°PSC- PμC,,PμPeC,Pë, CḡPēC- C-PSPrC- PjPsPḡCḡC,Cḡ PīPëPSPëPePSCḡC,Pë  
PīC-Pr C‡P°Cḡ P·P°CḡC,PscḡCḡPīP°PSPSCḡ C‡PëP»PsCḡC,P°P·PsP»Cḡ, P·P°P·PSP°C‡PμPSC- Cḡ  
C,P°P±P»PëC‡C- PSPëPḡC‡Pμ.

PḡP°CḡC,Psc,P° PīPsP±C-C‡PSPëC... CḡPμP°PeC‡C-PNë: PrCḡPḡPμ C‡P°CḡC,Ps (≥1/10);  
C‡P°CḡC,Ps (≥1/100, <1/10); PSPμC‡P°CḡC,Ps (≥1/1000, <1/100); CḡC-PrPePs (≥1/10000,  
<1/1000); PrCḡPḡPμ CḡC-PrPePs (<1/10000); C‡P°CḡC,Psc,P° PSPμPIC-PrPsPjP°  
(PSPμPjPsPḡP»PëPīPs PsC‡C-PSPëC,Pë PSP° PsCḡPSPsPIC- PSP°CḡPīPSPëC... PrP°PSPëC...).  
PḡP°CḡC,Psc,P° PīPsP±C-C‡PSPëC... CḡPμP°PeC‡C-PNë, CḡPēC-  
CḡPīPsCḡC,PμCḡC-PīP°P»PëCḡCḡ Cḡ PīPμCḡC-PsPr  
PīC-CḡP»CḡCḡPμC" CḡC,CḡP°C‡C-PNëPSPsPīPs P·P°CḡC,PscḡCḡPīP°PSPSCḡ,

PIPIP°P¶P°C”C,CHCÍCÍCÍ PSpµPIC-PrPsPjPsCñ (PSPµPjPsP¶P»PëPIPs PsC†C-PSPëC,Pë PSP° PsCÍPSPsPIC- PSP°CÍPIPSPëC... PrP°PSPëC...).

P— P±PsPeCž PeCñPsPIPsPSPsCÍPSPsC— C,P° P»C-PjC,,P°C,PëC†PSPsC— CÍPëCÍC,PµPjPë	C†P°CÍC,Ps	PµPeC...C-PjPsP·
	PSPµC†P°CÍC,Ps	P°PSPµPjC-CÍ
	CñC-PrPePs	PiCñPsPrPsPIP¶PµPSPSCÍ C†P°CÍCí PeCñPsPIPsC,PµC†C-, C,CñPsPjP±PsC†PëC,PµPjC-CÍ
P— P±PsPeCž C-PjCžPSPSPsC— CÍPëCÍC,PµPjPë	C†P°CÍC,PsC,P° PSPµPIC-PrPsPjP°	C,PµPSPPrPµPSC†C-CÍ PrPs PeCñPsPIPsC,PµC†, C,CñPsPjP±PsC†PëC,PsPiPµPSC-CÍ, PiCñP°PSC†P»PsC†PëC,PsPiPµPSC-CÍ, P°PiCñP°PSC†P»PsC†PëC,PsP·, P»PµPNëPePsPiPµPSC-CÍ, PiP°PSC†PëC,PsPiPµPSC-CÍ, P°PiP»P°CÍC,PëC†PSP° P°PSPµPjC-CÍ
	PSPµC†P°CÍC,Ps	P°P»PµCñPiC-C†PSP° CñPµP°PeC†C-CÍ
P— P±PsPeCž PsP±PjC-PSCž CñPµC†PsPIPëPS C,P° C...P°CñC†CžPIP°PSPSCÍ	C†P°CÍC,Ps	PiPµCñPëC,,PµCñPëC†PSC- PSP°P±CñCÍPePë, PSP°P±CñCÍPePë PsP±P»PëC†C†CÍ, P°PSPsCñPµPeCÍC-CÍ
	PSPµC†P°CÍC,Ps	PiC-PiPµCñPiP»C-PePµPjC-CÍ, C†CíPeCñPsPIPëPNë PrC-P°P±PµC,
P— P±PsPeCž PiCÍPëC...C-PePë	PSPµC†P°CÍC,Ps	C,CñPëPIPsP¶PSC-CÍC,CH
P— P±PsPeCž PSPµCñPIPsPIPsC— CÍPëCÍC,PµPjPë	PrC†P¶Pµ C†P°CÍC,Ps	PiPsP»PsPIPSPëPNë P±C-P»CH
	C†P°CÍC,Ps	P·P°PiP°PjPsCñPsC†PµPSPSCÍ
	PSPµC†P°CÍC,Ps	P±PµP·CÍPsPSPSCÍ, PSC-C†PSC- P¶P°C...Pë
	C†P°CÍC,PsC,P° PSPµPIC-PrPsPjP°	PiP°CñPµP·, PiC-PiPµCÍC,PµP·C-CÍ
P— P±PsPeCž PsCñPiP°PSC-PI P·PsCñCž	C†P°CÍC,PsC,P° PSPµPIC-PrPsPjP°	PePsPSBñ™ CñPSPeC,PëPIC-C,
P— P±PsPeCž PsCñPiP°PSC-PI CÍP»Cž...Cž	C†P°CÍC,PsC,P° PSPµPIC-PrPsPjP°	PrP·PIC-PS C- CëC†Pj C† PIC†C...P°C...
P— P±PsPeCž CÍPµCñC†PµPIPsC— CÍPëCÍC,PµPjPë	C†P°CÍC,Ps	PIC-PrC†C†C,C,CÍ CÍPµCñC†PµP±PëC,C,CÍ, C,P°C...C-PeP°CñPrC-CÍ, CÍC,PµPSPsPeP°CñPrC-CÍ, P°CñPëC,PjC-CÍ, CëP»C†PSPsC†PePsPIP° PµPeCÍC,CñP°CÍPëCÍC,PsP»C-CÍ
	PSPµC†P°CÍC,Ps	C-PSC,,P°CñPeC, PjC-PsPeP°CñPrP°, C,,C-P±CñPëP»CÍC†C-CÍ PiPµCñPµPrCÍPµCñPrCH, P·P°CÍC,C-PNëPSP° CÍPµCñC†PµPIP° PSPµPrPsCÍC,P°C,PSC-CÍC,CH, PSP°PrCëP»C†PSPsC†PePsPIP° C,P°C...C-PeP°CñPrC-CÍ, CëP»C†PSPsC†PePsPIP° C,P°C...C-PeP°CñPrC-CÍ, PSPµPiCñPëC,PsPjPSC-CÍC,CH

P— P±PsPeCž CÍCžPrPëPSPSPsC— CÍPëCÍC,PµPjPë	PSPµC±P°CÍC,Ps	PъCђPsPIPsPIPëP»PëPI PI PsPePs, PSPsCÍPsPIC- PeCђPsPIPsC,PµC±C-, CëP»CíPSPePsPIPs-PePëCëPePsPIC- PeCђPsPIPsC,PµC±C-, PSPµP·Bђ™ CИCÍPsPIP°PSC- PeCђPsPIPsC,PµC±C-, PsCђC,PsCÍC,P°C,PëC±PSP° PiC-PiPsC,PµPSP·C-CИ
	C±P°CÍC,PsC,P° PSPµPIC-PrPsPjP°	PiCђPëPiP»PëPIPë, PiC-PiPµCђC,PµPSP·C-CИ, PiC-PiPsC,PµPSP·C-CИ, PeCђPsPIPsPIPëP»PëPIPë PI PjPsP·PsPe, P»PµPiPµPSC-, PjBђ™ CИP·Pë, PrPëC...P°P»CђPSC- CëP»CИC...Pë; PiC-PrCëPeC-CђPSC- PeCђPsPIPsPIPëP»PëPIPë
P— P±PsPeCž PrPëC...P°P»CђPSPsC— CÍPëCÍC,PµPjPë	C±P°CÍC,Ps	CђPëPSC-C,, C,,P°CђPëPSPiC-C,
	PSPµC±P°CÍC,Ps	P·P°PrPëCëPeP°, PiPSPµPIPjPsPSC-CИ, PeP°CëPµP»Cђ
	C±P°CÍC,PsC,P° PSPµPIC-PrPsPjP°	C-PSC,PµCђCÍC,PëC±C-P°P»CђPSP° PiPSPµPIPjPsPSC-CИ
P— P±PsPeCž CëP»CžPSPePsPIPs- PePëCëPePsPIPsPiPs C,CђP°PeC,Cž	PrCíP¶Pµ C±P°CÍC,Ps	PrC-P°CђPµCИ, PiPsCђCíCëPµPSPSCИ PIPëPiPsCђPs¶PSPµPSPSCИ
	C±P°CÍC,Ps	PSCíPrPsC,P° C- P±P»CђPIP°PSPSCИ, PrPëCÍPiPµPiCÍC-CИ, PjPµC,PµPsCђPëP·Pj, P±C-P»Cђ Cí P¶PëPIPsC,C-
	PSPµC±P°CÍC,Ps	PiP°CÍC,CђPëC,
P— P±PsPeCž PiPµPiP°C,PsP±C-P»C-P°CђPSPsC— CÍPëCÍC,PµPjPë	C±P°CÍC,PsC,P° PSPµPIC-PrPsPjP°	PiPµPiP°C,PëC,, P°PSPsPjP°P»C-C— PiPµC±C-PSPePsPIPsC— C,,CíPSPeC±C-C—, P¶PsPIC,CИPSPëC±CИ
	C±P°CÍC,Ps	PIPëCÍPëPiP°PSPSCИ, CÍPIPµCђP±C-P¶
P— P±PsPeCž CëPeC-CђPë C- PiC-PrCëPeC-CђPSPëC... C,PeP°PSPëPs	C±P°CÍC,PsC,P° PSPµPIC-PrPsPjP°	PµPeP·PµPjP°, CëPeC-CђPSC- PIPëCÍPëPiP°PSPSCИ, CÍPëPSPPrCђPsPj PŸC,C-PIPµPSCÍP°-P°P¶PsPSCÍPsPSP°, C,PsPeCÍPëC±PSPëP№ PµPiC-PrPµCђPjP°P»CђPSPëP№ PSPµPeCђPsP»C-P·, PeCђPsPiPëPiBђ™ CИPSPeP°
	PSPµC±P°CÍC,Ps	PjC-P°P»PiC-CИ
P— P±PsPeCž CÍPePµP»PµC,PSPs- PjBђ™ CИP·PsPIPsC— CÍPëCÍC,PµPjPë C,P° CÍPiPsP»CžC±PSPsC— C,PeP°PSPëPSPë	CђC-PrPePs	PSPëCђPePsPIP° PSPµPrPsCÍC,P°C,PSC-CÍC,Cђ, PiPsCђCíCëPµPSPSCИ C,,CíPSPeC±C-C— PSPëCђPsPe
	C±P°CÍC,PsC,P° PSPµPIC-PrPsPjP°	PiPµPjP°C,CíCђC-CИ, PiPsP»P°PeC-CíCђC-CИ
P—P°PiP°P»CђPSC- CђPsP·P»P°PrPë	C±P°CÍC,Ps	P±C-P»Cђ Cí PiCђCíPrCИC..., P°CÍC,PµPSC-CИ
	PSPµC±P°CÍC,Ps	PsP·PSPsP±, PSPµP·PrCíP¶P°PSPSCИ
	C±P°CÍC,PsC,P° PSPµPIC-PrPsPjP°	PiC-CђPµPeCÍC-CИ, P±C-P»Cђ
P»P°P±PsCђP°C,PsCђPSC- PrPsCÍP»C-PrP¶PµPSPSCИ	C±P°CÍC,PsC,P° PSPµPIC-PrPsPjP°	PiC-PrPIPëC%PµPSPSCИ CђC-PIPSCИ CÍPµC±PsPIPsC— PePëCÍP»PsC,Pë, CÍPµC±PsPIPëPSPë Cí PeCђPsPIC-, PeCђPµP°C,PëPSC-PSCí PI PeCђPsPIC-

P—P±C-P»CђCëPµPSPSCИ PeC-P»CђPePsCÍC,C- PIPëPiP°PrPeC-PI PIC-PrC±CíC,C,CИ

СГРµСЧС+РµР±РёС,С,СЦ С,Р° РїРµСЧРёС,,РµСЧРёС±РSPёС... РSP°Р±СЧСЦРёС-РІ СГРїРсСГС,РµСЧС-РїР°Р»РsСГСЦ, РёРsР»Рё С+РёР»РsСГС,Р°Р·РsР» Р·Р°СГС,РsСГРsРІСРїР°Р»Рё СГСРјС-СГРSPs Р· С-РSCёРёРјРё РІР°Р·РsРРРёР»Р°С,Р°С,РsСЧР°РјРё, СЦРёС- РјРsР¶СРС,СН СГРїСЧРёС±РёРSPёС,Рё СЧРµС,,Р»РµС,РsСЧРSPСР С,Р°С...С-РёР°СЧРРС-СН, РSP°РїСЧРёРёР»Р°Р»Р°Р Р· РРРёРіС-РРСЧРsРіС-СЧРёРёРРёРSPsРІРёРјРё Р±Р»РsРёР°С,РsСЧР°РјРё РёР°Р»СНС+С-С"РІРёС... РёР°РSP°Р»С-РІ.

Р,,РРРёРSPёРј РїРsР±С-С±РSPёРј РµС,,РµС,РsРј, РїСЧРё СЦРёРsРјСР РІРёРјР°Р°Р»Рё РїСЧРёРёРіРёРSPµРSPSCЦ Р»С-РёСРРІР°Р»РSPSCЦ СР ≥ 3 % РїР°С+С-С"РSC,С-РІ, СЦРёС- РsС,СЧРёРёРјСРРІР°Р»Рё С+РёР»Р»РsСГС,Р°Р·РsР», Р±СРРІ РїРsР»РsРІРSPёР№ Р±С-Р»СН. Р+РSCёС- РїРsСёРёСЧРµРSC- РїСЧРёС±РёРSPё РїСЧРёРёРіРёРSPµРSPSCЦ Р»С-РёСРРІР°Р»РSPSCЦ РІРёР»Р»СНС+Р°Р»Р»ё СГРРёР»»СНРSPµ СГРµСЧС+РµР±РёС,С,СЦ С,Р° РРС-Р°СЧРµС (СЦРёС- РІРёРSPёРёР°Р»Р»Р Р· С±Р°СГС,РsС,РsСН 1,1 %).

Р—Р°СГС,РsСГСРІР°Р»РSPSCЦ С+РёР»»РsСГС,Р°Р·РsР»СР РјРsР¶Рµ Р±СРС,Рё РїРsРІВГ™ СЦР·Р°РSPµ Р· РїС-РРРІРёС%РµРSPёРј СЧРёР·Р·РёРёРsРј РёСЧРsРІРsС,РµС±С-, С- С+РµР№ СЧРёР·Р·РёРё РјРsР¶Рµ Р·Р±С-Р»СНСёСРРІР°Р»Р°С,РёСГСЦ РїСЧРё РsРРSPsС±Р°СГРSPsРјСР РїСЧРёРёР№РsРјС- Р»С-РёР°СЧРСГСНРёРsРіРs Р·Р°СГРsР±СР Р· Р±СРРСН-СЦРёРёРј С-РSCёРёРј РїСЧРµРїР°СЧР°С,РsРј, С%Рs С±РёРSPёС,СН С,Р°РёСР Р¶ РРС-СН.

Р РёР·РёРё РІРSCРС,СЧС-СёРSCНРsРsС±РSPёС... РёСЧРsРІРsС,РµС± РјРsР¶Рµ Р±СРС,Рё РІРёС%РёРј СР РїР°С+С-С"РSC,С-РІ С-Р· С+СРРёСЧРsРІРёРј РРС-Р°Р±РµС,РsРј.

Рј РїР°С+С-С"РSC,С-РІ РІС-РёРsРј РІС-РР 70 СЧРsРёС-РІ СГРРїРsСГС,РµСЧС-РїР°Р»Р°СГСЦ РїС-РРРІРёС%РµРSP° С±Р°СГС,РsС,Р° РІРёРіРРёРёС-РІ РРС-Р°СЧРµС— С,Р° СГРРёР»»СНРSPsРіС СГРµСЧС+РµР±РёС,С,СЦ.

РµРsРІС-РРРsРјР»РµРSPSCЦ РїСЧРs РїС-РРРsР·СЧСНРІР°Р»РSPSC- РїРsР±С-С±РSC- СЧРµР°РёС

РµРsРІС-РРРsРјР»РµРSPSCЦ РїСЧРs РїС-РРРsР·СЧСНРІР°Р»РSPSC- РїРsР±С-С±РSC- СЧРµР°РёС-С— РїС-СГР»СЦ СЧРµС"СГС,СЧР°С+С-С— Р»С-РёР°СЧРСГСНРёРsРіРs Р·Р°СГРsР±СР С" РІР°Р»Р»Р»Р»Р»Р» РїСЧРsС+РµРРРСЧРsСН. Рј РРРsР·РІРsР»СЦС" РїСЧРsРРРsРІР°Р»Р°С,Рё РјРsРSC-С,РsСЧРёРSPi СГРРІС-РІРІС-РРРSPsСёРµРµРSPSCЦ РёРsСЧРёСГС,СН/СЧРёР·Р·РёРё РРР»СЦ РІС-РРРїРsРІС-РРРSPsРіРs Р»С-РёР°СЧРСГСНРёРsРіРs Р·Р°СГРsР±СР. РµРµРРРёС±РSPёРј РїСЧР°С+С-РІРSPёРёРј РSPµРsР±С...С-РРРSPs РїРsРІС-РРРsРјР»СЦС,Рё РїСЧРs Р±СРРРСН-СЦРёС- РїС-РРРsР·СЧСНРІР°Р»РSPSC- РїРsР±С-С±РSC- СЧРµР°РёС-С— С±РµСЧРµР· РSP°С+С-РsРSP°Р»СНРSCР СГРРёСГС,РµРјСР РїРsРІС-РРРsРјР»РµРSPСН.

**РµРµСЧРС-РS РїСЧРёРРР°С,РSPsСГС,С-. 3 СЧРsРёРё.**

**РјРјРsРІРё Р·Р±РµСЧС-РїР°РSPSCЦ.**

Р—Р±РµСЧС-РїР°С,Рё РїСЧРё С,РµРјРјРµРµСЧР°С,СРСЧС- РSPµ РІРёС%РµРµ 25 В°РŸ РІ РsСЧРёРіС-РSP°Р»СНРSC-Р№ СРРїР°РёРsРІС+С-.



P—P±PμCḡC—PiP°C, Pë Cí PSPμPrPsCíC, CíPíPSPsPjCí PrP»Cḡ PrC—C, PμPNḡ PjC—CíC†C—.

**PJḡP°PePsPIPeP°.**

PμPs 10 C, P°P±P»PμC, PsPe Cí P±P»C—CíC, PμCḡC—; PíPs 3 P°P±Ps 6 P±P»C—CíC, PμCḡC—PI Cí PeP°CḡC, PsPSPSC—PNḡ CíPíP°PePsPIC†C—.

**PḡP°C, PμPiPsCḡC—Cḡ PIC—PrPíCíCíP°eCí.** P—P° CḡPμC†PμPíC, PsPj.

**P°PëCḡPsP±PSPëPe.**

PḡPḡ B«PḡPrP°PjPμPr PḡP°CḡPjP°B».

**PḡC—CíC†PμP·PSP°C...PsPrPḡPμPSPSCḡ PIPëCḡPsP±PSPëPeP° C, P° P°PrCḡPμCíP° PjC—CíC†Cḡ PíCḡPsPIP°PrPḡPμPSPSCḡ PNḡPsPiPs PrC—CḡP»CḡPSPsCíC, C—.**

P°CíP». PḡP°CḡC€. P°Pḡ. PμC—P»CíCíPrCíCḡPePsPiPs 5, 95-200, PμP°P±Bḡ™ CḡPSC—C†Pμ, PμPsP»CḡC%oP°.

**P—P°CḡPIPSPëPe.**

PḡP·Pḡ PḡP°CḡPjP°CíCḡCḡC, C—PeP°P»Cí P>C—PjC—C, PμPr.

**PḡC—CíC†PμP·PSP°C...PsPrPḡPμPSPSCḡ P·P°CḡPIPSPëPeP°.**

PḡPμPSPSPμPrC— PḡPIPμPSCḡ 89/201, 1077 PḡC—PePsCíC—Cḡ, PḡC—PíCḡ.