

$P''Ps \text{ Cl}'PeP \gg P^\circ PrCr' P \gg C-PeP^\circ C\bar{H}Cl'CHPePsPiPs \text{ } P \cdot P^\circ Cl'PsP \pm Cr' PIC \dots PsPrC\bar{H}C, CH_3 \text{ } PrC-C\bar{H}C \ddagger C-$
 $C\bar{H}P_{\mu}C \ddagger PsPIP\ddot{e}PSP\ddot{e} P \cdot P^\circ PSC, P\ddot{e}PiC-PiP_{\mu}C\bar{H}C, P_{\mu}PSP \cdot P\ddot{e}PIPSP\ddot{e}PjP\ddot{e} PIP \gg P^\circ Cl'C, P\ddot{e}PIPsCl'C, C\bar{H}PjP\ddot{e}:$
 $C\bar{H}P_{\mu}P \cdot P_{\mu}C\bar{H}PiC-PS, PeP \gg PsPiP^\circ PjC-Pr \text{ } C-PrP\ddot{e}PiC-PrC\bar{H}PsP_{\mu}C\bar{H}PiPsPeC\bar{H}P\ddot{e}Cl'C, P\ddot{e}PS.$

РμPсC”PrPSP°PSPSCЦ PиPsP·PёC,PePIPSPsC— PrC-C— CђPμP·PμCђPиC-PSCр,
PeP»PsPiP°PjC-PrCр C,P° PrPePiC-PrCђPsPμCђPiPsPeCђPeCрC,PёPSCр PиPsCрPёP»CђC”
PiC-PиPsC,PμPSP·PePIPSCr PrC-Cђ PiCђPμPiP°CђP°C,Cр C,P° PrP°C” PjPsP¶P»PePIC-CрC,Cђ
P·P°CрC,PsCрPsPICрPiP°C,Pё PjPμPSCёC- PrPsP·Pё PePsP¶PSPsPiPs P· PePsPjPiPsPSPμPSC,C-PI,
C%oPs P·PjPμPSCёCрC” CђPeP·PёPe PиPsP±C-CђPSPёC... PμC,,PμPeC,C-PI.

P PμP·PμCђPиC-PS Bђ” P°P»PeP°P»PsC—Pr CђP°CрPiPsP»CHC,,C-C— P·
PSPμPNёCђPsP»PμPiC,PёCђPSPsCђ C,P° PiC-PиPsC,PμPSP·PePIPSPsCђ PrC-CЦPjPё.
PќP°P»PμP¶PёC,Cђ PrPs PiCђCрPiPё P»C-PeP°CђCрCHPePёC... P·P°CрPsP±C-PI, CЦPeC-
P±P»PsPeCрCђC,Cђ P°PrCђPμPSPμCђPiC-CђPSC- PSPμPNёCђPsPSPё. PђPμC...P°PSC-P·Pj
PrC-C— PиPsP»CЦPiP°C” Cр PiPёPIC-P»CHPSPμPSPSC- PSPμPNёCђPsPjPμPrC-P°C,PsCђC-PI P·
P°PrCђPμPSPμCђPiC-CђPSPёC... CрPePSPsPiC,PёCђPSPёC... PiCђP°PSCрP»CЦCђPSPёC...
PrPμPiPs PiCђPμCрPёPSP°PiC,PёCђPSPёC... CрC,CђCрPeC,CрCђ. P PμP·PμCђPиC-PS
PiPμCђPμCёPePsPrP¶P°C” P·P°C...PsPiP»PμPSPSCђ C,P° PSP°PePsPiPёCђPμPSPSCђ
PeP°C,PμC...PsP»P°PjC-PSC-PI C- CрPμCђPsC,PsPSC-PSCр PI CрPμPSC,CђP°P»CHPSC-PNё C,P°
PiPμCђPёC,,PμCђPёCђPSC-PNё PSPμCђPiPsPIC-PNё CрPeCрC,PμPjC-, CрPiCђPёCђPePSCЦC”
CђPμC,,P»PμPeC,PsCђPSCр PiP°PiPsC,PsPSC-Cђ. P PsP·CёPeCђCђC” PeCђPsPiPsPSPsCрPSC-
CрCрPrPePSPё, CрPiPsPIC-P»CHPSCђC” CђP°CрC,PsC,Cр CрPμCђCрPμPiPёC...
CрPePsCђPsCђPμPSCђ C- CрPiCђPёCђPePSCЦC” C,CђPePiP°P»Pμ P·PSPёP¶PμPSPSCЦ
P°CђC,PμCђC-P°P»CHPSPsPiPs C,PёCрPeCр.

PђP»PsPiP°PjC-Pr Bђ” CђPμCђPsPiPёPSP° P· PiPsPjC-CђPSPsCђ C-PSC,PμPSCрPёPiPSC-CрC,Cђ
PrC-CрCђPμC,PёCђPSPsC— PrC-C—, C%oPs PiCђPёC,P°PjP°PSPSP° PiCђCрPiC-
C,C-P°P·PePrPSPёC... PrC-CрCђPμC,PёPeC-PI. PμCђPёPiPSC-CђCрC”
CђPμP°P±CрPsCђP±CрC-Cђ C-PsPSC-PI PSP°C,CђC-Cђ Cр PePsCђPePsPIC-PNё
PrC-P»CЦPSCрC- PiPёCрC...C-PrPSPsPiPs PIC-PrPrC-P»Cр PiPμC,P»C- P”PμPSP»Pμ, P°
C,P°PePsP¶ Cр P·PiPёPiPёCрC,PёC... PrPeCрC,P°P»CHPSPёC... PeP°PSP°P»CHCрCЦC..., C%oPs
CрPiCђPёCђPePSCЦC” PiPsCрPёP»PμPSPSCЦ PrC-CрCђPμP·Cр C,P° PSP°C,CђC-PNёCрCђPμP·Cр
C,P° P·PSPёP¶PμPSPSCЦ P°CђC,PμCђC-P°P»CHPSPsPiPs C,PёCрPeCр.
PμCђPePiPSC-CђPμPSPSCЦ P·PiPsCђPsC,PSPsPiPs PICрPjPsPeC,CрPiP°PSPSCЦ C-PsPSC-PI
PSP°C,CђC-Cђ PiPsCрPёP»CђC” PeP°P»C-PNёCрCђPμP· C- PjPsP¶Pμ PiCђPёP·PiPμCрC,Pё PrPs
PiC-PиPsPeP°P»C-C”PjC-C—.

P”PePiC-PrCђPsPμCђPiPsPeCђPeCрC,PёPS PSP°P»PμP¶PёC,Cђ PrPs PiCђCрPiPё
PiC-PrCђPsPiPμPSC-P·PsPiP°PSPёC... P°P»PeP°P»PsC—PrC-PI CђC-P¶PeC-PI. P’C-PS C”
P°PiPsPSC-CрC,PsPj PrPsC,,P°PjC-PSPμCђPiC-CђPSPёC... C,P°
CрPμCђPsC,PsPSC-PSPμCђPiC-CђPSPёC... CђPμCрPμPiC,PsCђC-PI C,P°
P°PSC,P°PiPsPSC-CрC,PsPj P°P»CHC,,P°-P°PrCђPμPSPsCђPμCрPμPiC,PsCђC-PI.
P—P°PiPrCЦPePё PrC-C— PSP° CрPμPSC,CђP°P»CHPSCр PSPμCђPiPsPICр CрPeCрC,PμPjCр
P·PSPёP¶CрC” C,PsPSCрCр CрC,C-PSPsPe PeCђPsPiPsPSPsCрPSPёC... CрCрPrPePS,
P·PjPμPSCёCрC” C,PёCрPe. PђCђC-Pj C,PsPiPs PrPePiC-PrCђPsPμCђPiPsPeCђPeCрC,PёPS
PiCђPёPiPSC-CђCрC” CђPμC,,P»PμPeCрPё P±P°CђPsCђPμCрPμPiC,PsCђC-PI C,P°
P·PjPμPSCёCрC” CђPμP°PeC,PёPiPSC- C,P°C...C-PeP°CђPrC-C—.

PαP°CђPjP°PePsPeC-PSPμC,PёPeP°.

PђP±CрPsCђP±CрC-CЦ C,P° CђPsP·PiPsPrC-P». PђPeCђPμPjC- PePsPjPiPsPSPμPSC,Pё
PќPsCђPjP°C,PμPSCрCр PICрPjPsPeC,CрCђC,CђCрCЦ P·C- CёP»CрPSPePsPiPs-
PePёCёPePsPiPs C,CђP°PeC,Cр PI C,P°PePёC... PeC-P»CHPePsCрC,CЦC...:
CђPμP·PμCђPиC-PS - PiCђPёP±P»PeP·PSPs 40 %, PeP»PsPiP°PjC-Pr - P±C-P»CHCёPμB B
90 %, PrPePiC-PrCђPsPμCђPiPsPeCђPeCрC,PёPS - PiCђPёP±P»PeP·PSPs 25 %.
PђP°PeCрPёPjP°P»CHPSCр PePsPSCрPμPSC,CђP°CрC-Cђ Cр PeCђPsPIC- PiC-CрP»CЦ

РiРµCђPсCђP°P»CђPSPsPiPs P·P°CГC,PsCГCГPIP°PSPSCЦ PIPsPSPë PrPsCГCЦPIP°CђC,CH CђPµCђPµP· 1-3 PiPsPrPëPSPë, 1-2 PiPsPrPëPSPë C,P° 0,6 PiPsPrPëPSPë PIC-PrPiPsPIC-PrPSPs.

P PµP·PµCђPiC-PS PSPµ P·PIБђ™ CЦP·CГC”C,CHCГCЦ P· P±C-P»PeP°PjPë PiP»P°P·PjPë PeCђPsPIC-, P° PeP»PsPiP°PjC-Pr C,P° PrPëPiC-PrCђPsPµCђPiPsPeCђPëCГC,PëPS P·PIБђ™ CЦP·CГCђC,CHCГCЦ P· P±C-P»PeP°PjPë PiP»P°P·PjPë PeCђPsPIC- PSP° 46 % C,P° 68 % PIC-PrPiPsPIC-PrPSPs. PhP±Бђ™ C”Pj CђPsP·PiPsPrC-P»CГ PeP»PsPiP°PjC-PrCГ CГC,P°PSPsPiPëC,CH 1,5 P»/PePi, CГ C,PsP№ CђP°CГ CЦPe PrPëPiC-PrCђPsPµCђPiPsPeCђPëCГC,PëPSCГ - 16 P»/PePi.

P PµP·PµCђPiC-PS C,P° PrPëPiC-PrCђPsPµCђPiPsPeCђPëCГC,PëPS PiCђPsPSPëPeP°CђC,CH PeCђC-P·CH PiP°PjP°C,PsPµPSC†PµC,,P°P»C-CђPSPëP№ P±P°CђБђ™ C”Cђ C,P° PiP»P°C†PµPSC,CГ.

PћPµC,P°P±PsP»C-P·Pj C,P° PIPëPIPµPrPµPSPSCЦ. P’CГC- PePsPjPiPsPSPµPSC,Pë PiCђPµPiP°CђP°C,CГ PjPµC,P°P±PsP»C-P·CГCђC,CHCГCЦ CГ PiPµCђC-PSC†C- PrPs PSPµP°PeC,PëPIPSPëC... PjPµC,P°P±PsP»C-C,C-PI (PiCђPëCђPsPjCГ PeP»PsPiP°PjC-Pr CГ PjPµPSC€C-P№ PjC-CђC-, PSC-P¶ CђPµP·PµCђPiC-PS C,P° PrPëPiC-PrCђPsPµCђPiPsPeCђPëCГC,PëPS).

P PµP·PµCђPiC-PS PI PsCГPSPsPIPSPsPjCГ PIPëPIPsPrPëC,CHCГCЦ CГ PIPëPiP»CЦPrC- PjPµC,P°P±PsP»C-C,C-PI C-P· PeP°P»PsPj C,P° CГPµCђPµCђ (PjPµPSC€Pµ 1 % PIPëPIPsPrPëC,CHCГCЦ C-P· CГPµCђPµCђ). PµPµCђC-PsPr PSP°PiC-PICђPsP·PiP°PrCГ CГC,P°PSPsPiPëC,CH PiCђPëP±P»PëP·PSPs 4,5 PiPsPrPëPSPë CГ P°P»CHC,,P°-C,,P°P·C- C,P° PiCђPëP±P»PëP·PSPs 271 PiPsPrPëPSCГ CГ P±PµC,P°-C,,P°P·C-.

P”PëPiC-PrCђPsPµCђPiPsPeCђPëCГC,PëPS PIPëPIPsPrPëC,CHCГCЦ P· PsCђPiP°PSC-P·PjCГ PI PsCГPSPsPIPSPsPjCГ P· PeP°P»PsPj; PjPµPSC€Pµ 1 % PIPëPrC-P»CЦC”C,CHCГCЦ C-P· CГPµCђPµCђ. PµPµCђC-PsPr PSP°PiC-PICђPsP·PiP°PrCГ CГC,P°PSPsPiPëC,CH PiCђPëP±P»PëP·PSPs 2 PiPsPrPëPSPë PI P°P»CHC,,P°-C,,P°P·CГ C,P° PiCђPëP±P»PëP·PSPsB B B B B B B 14 PiPsPrPëPS PI P±PµC,P°-C,,P°P·CГ.

PљP»PsPiP°PjC-Pr PIPëPIPsPrPëC,CHCГCЦ, PiPsP»PsPIPSPëPj CђPëPSPsPj, CђPµCђPµP· PSPëCђPePë (P±C-P»CHC€Pµ 30 % PIC-Pr P·P°CГC,PsCГPsPIP°PSPsC— PrPsP·Pë). PµPµCђC-PsPr PSP°PiC-PICђPsP·PiP°PrCГ CГC,P°PSPsPiPëC,CH PiCђPëP±P»PëP·PSPs 6 PiPsPrPëPS.

РљP»C-PSC-CђPSC- C...P°CђP°PeC,PµCђPëCГC,PëPePë.

PµPsPeP°P·P°PSPSCЦ.

P·CГPµPSC†C-P°P»CHPSP° P°CђC,PµCђC-P°P»CHPSP° PiC-PiPµCђC,PµPSP·C-CЦ C,P° PICГC- C,,PsCђPjPë CГPëPjPiC,PsPjP°C,PëCђPSPëC... P°CђC,PµCђC-P°P»CHPSPëC... PiC-PiPµCђC,PµPSP·C-P№, PjPsPSPsC,PµCђP°PiC-CЦ CЦPePëC... PSPµPµC,,PµPeC,PëPIPSP°.

PµCђPsC,PëPiPsPeP°P·P°PSPSCЦ.

PµC-PrPIPëC%PµPSP° CђCГC,P»PëPIC-CГC,CH PrPs CГCГP»CHC,,P°PSC-P»P°PjC-PrC-PI, CђPµP·PµCђPiC-PSCГ C,P° P°P»PeP°P»PsC—PrC-PI CђC-P¶PeC-PI C,P° C-PSC€PëC...

PePsPjPiPsPSPμPSC,C-PI P»C-PeP°CḡCfCHPePsPiPs P·P°CfPsP±Cf. P°C-PiPsPeP°P»C-C°PjC-Cḡ.

PμPμCḡC-PsPr PIP°PiC-C,PSPsCfC,C- C,P° PiPsPrCfPIP°PSPSCḡ PiCḡCfPrPrCḡ.

P PμP·PμCḡPiC-PS.

P°PμPiCḡPμCfC-Cḡ (PSP° PrP°PSPëPμ C±P°Cf P°P±Ps PI P°PSP°PjPSPμP·C-), C...PIPsCḡPsP±P° PμP°CḡPeC-PSCfPsPSP°, PμPiC-P»PμPiCfC-Cḡ, PμP»PμPeC,CḡPsCfCfPrPsPjPSP° C,PμCḡP°PiC-Cḡ. P°PëCḡP°P·PePsPIP° C...PIPsCḡPsP±P° CëP»CfPSPeP° C,P° PrPIP°PSP°PrC†CḡC,PëPiP°P»PsC— PePëCëPePë Cf C„P°P·C- P·P°PiPsCfC,CḡPμPSPSCḡ, PIPëCḡP°P·PePsPIPëPμ PePsP»C-C,; C„PμPsC...CḡPsPjPsC†PëC,PspjP°, CfCfPiCḡPsPIC-PrPSPμ P°P±Ps PSPμPrP°PIpsc” P»C-PeCfPIP°PSPSCḡ C-PSPiC-P±C-C,PscḡP°PjPë PjPsPSPsP°PjC-PSPsPeCfPëPrP°P·Pë.

PḡC-P°P·PëPrPSC- PrC-CfCḡPμC,PëPePë.

PḡCḡP¶PeP° PSPëCḡPePsPIP° PSPμPrPsCfC,P°C,PSC-CfC,Cḡ (PeP»C-CḡPμPSCf PeCḡPμP°C,PëPSC-PSCf PjPμPSCëPμ 30 PjP»/C...PI); PiPμC±C-PSPePsPIP° PSPμPrPsCfC,P°C,PSC-CfC,Cḡ; PiC-PiPsPSP°C,CḡC-C°PjC-Cḡ; PiC-PiPμCḡCfCḡPëPePμPjC-Cḡ.

P°PëPiC-PrCḡPsPμCḡPiPsPeCḡPëCfC,PëPS.

PḡPrPSPsC±P°CfPSPμ P·P°CfC,PscfCfPIP°PSPSCḡ P· CfPëP»CḡPSPsPrC-CḡC±PëPjPë C-PSPiC-P±C-C,PscḡP°PjPë C-P·PsC„PμCḡPjPμPSC,C-PI CYP3A4, C,P°PePëC..., CḡPe C-PSPiC-P±C-C,PscḡPë PiCḡPsC,PμP°P·Pë, PjP°PeCḡPsP»C-PrPSC- P°PSC,PëP±C-PsC,PëPePë C,P° P°P·PsP»CḡPSC- PiCḡPsC,PëPiCḡPëP±PePsPIC- P·P°CfPsP±Pë (PrPëPI. CḡPsP·PrC-P» B«P·P·P°C°PjPsPrC-Cḡ P· C-PSCëPëPjPë P»C-PeP°CḡCfCHPePëPjPë P·P°CfPsP±P°PjPë C,P° C-PSCëC- PIPëPrPë PIP·P°C°PjPsPrC-PμB»).

P°P·P°C°PjPsPrC-Cḡ P· C-PSCëPëPjPë P»C-PeP°CḡCfCHPePëPjPë P·P°CfPsP±P°PjPë C,P° C-PSCëC- PIPëPrPë PIP·P°C°PjPsPrC-PμB.

PḡPëCfC,PμPjPSC- PePsCḡC,PëPePsCfC,PμCḡPsC—PrPë C,P° PSPμCfC,PμCḡPsC—PrPSC- PiCḡPsC,PëP·P°PiP°P»CḡPSC- PiCḡPμPiP°CḡP°C,Pë PiPsCfP»P°P±P»CḡCḡC,Cḡ P°PSC,PëPiC-PiPμCḡC,PμPSP·PëPIPSCf PrC-Cḡ PiCḡPμPiP°CḡP°C,Cf.

PμCḡPë PsPrPSPsC±P°CfPSPsPjCf P·P°CfC,PscfCfPIP°PSPSC- P· PiCḡPμPiP°CḡP°C,P°PjPë, C%Ps P·PSPëP¶CfCḡC,Cḡ P°CḡC,PμCḡC-P°P»CḡPSPëPμ C,PëCfPe, PjPsP¶P»PëPIC- PiPsCfPëP»PμPSPSCḡ PiC-PiPsC,PμPSP·PëPIPSPsPiPs PμC„PμPeC,Cf PkPsCḡPjP°C,PμPSCfCf.

PμCḡPμPiP°CḡP°C, P·PjPμPSCëCfC” PeP»C-CḡPμPSCf P»C-C,C-Cḡ PSPëCḡPeP°PjPë, C%Ps PiCḡPëP·PIPsPrPëC,Cḡ PrPs PiC-PrPIPëC%PμPSPSCḡ CḡC-PIpsc-PI Cf PeCḡPsPIC-. PμCḡPë C—C... PsPrPSPsC±P°CfPSPsPjCf P·P°CfC,PscfCfPIP°PSPSC- PSPμPsP±C...C-PrPSPs PePsPSC,CḡPsP»CḡPIP°C,Pë CḡC-PIPμPSCḡ P»C-C,C-Cḡ Cf PeCḡPsPIC- C,P° PiCḡPë PSPμPsP±C...C-PrPSPsCfC,C- PIC-PrPePsCḡPëPiCfPIP°C,Pë PrPsP·Pë.

PḡP»PsPiP°PjC-Pr, C%Ps PIC...PsPrPëC,Cḡ PrPs CfPeP»P°PrCf PiCḡPμPiP°CḡP°C,Cf, P·PSPëP¶CfC” PrC-Cḡ PiPμCḡPsCḡP°P»CḡPSPëC... P°PSC,PëPePsP°PiCfP»CḡPSC,C-PI.

P·C„PμPeC,Pë P»C-PeP°CḡCfCHPePëC... P·P°CfPsP±C-PI, CḡPeC- PrPμPiCḡPμCfPëPIPSPs PrC-CḡC,Cḡ PSP° C†PμPSC,CḡP°P»CḡPSCf PSPμCḡPIPsPICf CfPëCfC,PμPjCf

(P°P»PePsPiPsP»CH C,P° P°PSC,PëPrPµPîCḡPµCḡP°PSC,Pë), Cḡ PîPsC”PrPSP°PSPSC- P· PḡPsCḡPjP°C,PµPSCḡPsPj PjPsPḡCḡC,CH PîPsCḡPëP»CḡPIP°C,PëCḡCḡ. PhCḡCH CḡPsPjCḡ PîP°CḡC-C”PSC,Pë PSPµ PîPsPIPëPSPSC- PIPḡPëPIP°C,Pë P°P»PePsPiPsP»CH C,P° P· PsP±PµCḡPµPḡPSC-CḡC,Cḡ P·P°CḡC,PsCḡPsPICḡPIP°C,Pë P°PSC,PëPrPµPîCḡPµCḡP°PSC,Pë PîCḡPë C,PµCḡP°PîC-C— PḡPsCḡPjP°C,PµPSCḡPsPj.

P PµP·PµCḡPîC-PS PîPsCḡPëP»CḡC” PîCḡPëPîPSC-CḡCḡPIP°P»CHPSPëPNḡ PIPîP»PëPI PSP° CḡPµPSC,CḡP°P»CHPSCḡ PSPµCḡPIPḡPICḡ CḡPëCḡC,PµPjCḡ P·P°CḡPsP±C-PI PrP»Cḡ PSP°CḡPePsP·Cḡ, PrPµCḡPePëC... P°PSC,PëPîC-CḡC,P°PjC-PSPSPëC... PîCḡPµPîP°CḡP°C,C-PI C,P° P±P°CḡP±C-C,CḡCḡP°C,C-PI. P PµP·PµCḡPîC-PS PjPsPḡPµ PîPsCḡPëP»CḡPIP°C,Pë PµC,,PµPeC,Pë P°PrCḡPµPSP°P»C-PSCḡ P°P±Ps C-PSCëPëC... CḡPëPjPîP°C,PsPjC-PjPµC,PëCḡPSPëC... CḡPµCḡPsPIPëPS.

PḡPµ CḡPµPePsPjPµPSPḡCḡC,CHCḡCḡ PîPsC”PrPSCḡPIP°C,Pë P·P°CḡC,PsCḡCḡPIP°PSPSCḡ PîCḡPµPîP°CḡP°C,Cḡ P· C-PSPîC-P±C-C,PsCḡP°PjPë PjPsPSPsP°PjC-PSPsPeCḡPëPrP°P·Pë CḡPµCḡPµP· PîCḡPëPîPSC-CḡPµPSPSCḡ CḡPµPSC,CḡP°P»CHPSPsC— PSPµCḡPIPḡS— CḡPëCḡC,PµPjPë, PîC-PrPIPëC%PµPSPSCḡ P°CḡC,PµCḡC-P°P»CHPSPsPiPs C,PëCḡPeCḡ C,P° P·PSP°CḡPSPµ P·PSPëPḡPµPSPSCḡ C,PµPjPîPµCḡP°C,CḡCḡPë C,C-P»P°. PḡCḡC-Pj C,PsPiPs, C-PSC,PµCḡPIP°P» PjC-Pḡ P·P°CḡC,PsCḡCḡPIP°PSPSCḡPj CḡPëC... PîCḡPµPîP°CḡP°C,C-PI PjP°C” CḡC,P°PSPsPIPëC,Pë PSPµ PjPµPSCëPµ 2 C,PëPḡPSC-PI.

PḡCḡPµPîP°CḡP°C, P·PjPµPSCëCḡC” PµC,,PµPeC,PëPIPSC-CḡC,CH P»PµPIPḡPrPsPîPë.

PḡCḡPë PîPsC”PrPSP°PSPSC- CḡPµP·PµCḡPîC-PSCḡ P· PrC-CḡCḡPµC,PëPSP°PjPë PîPsCḡPëP»CḡC”C,CHCḡCḡ CḡPëP·PëPe PîC-PîPsC,PµPSP·C-C—. PḡCḡPë PîPsC”PrPSP°PSPSC- P· C-PSCëPëPjPë PîC-PîPsC,PµPSP·PëPIPSPëPjPë P·P°CḡPsP±P°PjPë Bḡ” PîPsCḡPëP»PµPSPSCḡ PîC-PîPsC,PµPSP·PëPIPSPsC— PrC-C—.

PhPrPSPsCḡP°CḡPSPµ P·P°CḡC,PsCḡCḡPIP°PSPSCḡ P·P°CḡPsP±C-PI P· PIPjC-CḡC,PsPj PîPsC...C-PrPSPëC... CḡC-PḡPeC-PI P· C-PSPîC-P±C-C,PsCḡP°PjPë C-P·PsC,,PµCḡPjPµPSC,C-PI CYP3A4, C,P°PePëC..., CḡPe PjP°PeCḡPsP»C-PrPSC- P°PSC,PëP±C-PsC,PëPePë, C-PSPîC-P±C-C,PsCḡPë PîCḡPsC,PµP°P·Pë P°P±Ps P°P·PsP»CHPSC- PîCḡPsC,PëPîCḡPëP±PePsPIC- P·P°CḡPsP±Pë PîCḡPsC,PëPîPsPeP°P·P°PSPµ (PrPëPI. CḡPsP·PrC-P» B«PḡCḡPsC,PëPîPsPeP°P·P°PSPSCḡB»), PsCḡPeC-P»CHPePë PI C,P°PePëC... PIPëPîP°PrPeP°C... PjPsPḡP»PëPIPëPNḡ PîPsCḡC,CḡPëPNḡ C,PsPeCḡPëCḡPSPëPNḡ PIPîP»PëPI P°P»PeP°P»PsC—PrC-PI CḡC-PḡPeC-PI (PµCḡPiPsC,PëP·Pj), CḡPePëPNḡ C...P°CḡP°PeC,PµCḡPëP·CḡC”C,CHCḡCḡ PIP°P·Ps CḡPîP°P·PjPsPj C,P° C-CëPµPjC-C”Cḡ PeC-PSCḡC-PIPsPe, C%Ps C-PSPëPsP»Pë PîCḡPëP·PIPḡPrPëC,CH PrPs PSPµPsP±C...C-PrPSPsCḡC,C- P°PjPîCḡC,P°CḡC-C—. PḡCḡPëPeP»P°PrP°PjPë CḡPëP»CHPSPsPrC-CḡCḡPëC... C-PSPîC-P±C-C,PsCḡC-PI C-P·PsC,,PµCḡPjPµPSC,C-PI CYP3A4 C” PîCḡPsC,PëPîCḡPëP±PePsPIC- P·P°CḡPsP±Pë PePµC,PsPePsPSP°P·PsP» C,P° C-C,CḡP°PePsPSP°P·PsP», C-PSPîC-P±C-C,PsCḡPë PîCḡPsC,PµP°P·Pë CḡPëC,PsPSP°PIC-Cḡ, PSPµP»C,,C-PSP°PIC-Cḡ C,P° C-PSPḡPëPSP°PIC-Cḡ, P° C,P°PePsPḡ PjP°PeCḡPsP»C-PrPSC- P°PSC,PëP±C-PsC,PëPePë PµCḡPëC,CḡPsPjC-CḡPëPS, PeP»P°CḡPëC,CḡPsPjC-CḡPëPS C,P° C,CḡPsP»PµP°PSPḡPsPjC-CḡPëPS. PḡPSCëC- PjPµPSCë PîPsC,CḡPḡPSC- C-PSPîC-P±C-C,PsCḡPë C-P·PsC,,PµCḡPjPµPSC,C-PI CYP3A4 CḡP»C-Pr P·P°CḡC,PsCḡPsPICḡPIP°C,Pë P· PsP±PµCḡPµPḡPSC-CḡC,Cḡ. P”Ps PjPµPSCë PîPsC,CḡPḡPSPëC... C-PSPîC-P±C-C,PsCḡC-PI PSP°P»PµPḡP°C,CH CḡP°PePIC-PSP°PIC-Cḡ, PSPµC,,P°P·PsPrPsPS, C,,P»CḡPePsPSP°P·PsP», PîCḡPµPNḡPîC,,CḡCḡC,PsPIPëPNḡ CḡC-Pe, C,,P»CḡPsPeCḡPµC,PëPS, C,,P»CḡPIPḡPeCḡP°PjC-PS, P·PëP»PµCḡC,PsPS C,P° PeP»PsC,CḡPëPjP°P·PsP».

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

вҖґ PґC-Pr CґP°CÍ P»C-PeCÍPIP°PSPSCÍ PSPґ CÍP»C-Pr PIPґPëPIP°C,Pë CÍPіPëCґC,PSC-
PSP°PіPsC—.

вҖґ PґP»C-Pr PіP°PґвҖґ™ CІC,P°C,Pë, C%Ps PeP»C-PSC-CґPSPëPNґ PґC,,PґPeC, PIC-Pr
P·P°CÍC,PsCÍCґPIP°PSPSCÍ PќPsCґPґP°C,PґPSCÍCґ PSP°CÍC,P°C” CґPґCґPґP·
PrPґPeC-P»CHPeP° PrPSC-PI, P° C-PSPsPrC- - CґPґCґPґP· 4 C,PëPґPSC- PіC-CÍP»CІ
PіPsCґP°C,PeCґ P»C-PeCґPIP°PSPSCÍ. PґPґCґPґP· CґPґ PSPґ CÍP»C-Pr
PіC-PrPIPëC%CґPIP°C,Pë PrPsP·CґPIP°PSPSCÍ, CІPeC%Ps PSP° PіPsCґP°C,PeCґ
P»C-PeCґPIP°PSPSCÍ PSPґ PIC-PrP±CґP»PsCÍCІ P·PSPëPґPґPSPSCÍ
P°CґC,PґCґC-P°P»CHPSPsPiPs C,PëCÍPeCґ.

вҖґ PJ P·PiбҖ™ CІP·PeCґ P· PґPsPґP»PëPIPëPґ PіC-PrPIPëC%PґPSPSCІPґ
PePsPSCґPґPSC,CґP°CґC-C— PiP»CHPePsP·Pë PI PeCґPsPIC- PіP°CґC-C”PSC,P°Pґ P·
CґCґPeCґPsPIPëPґ PrC-P°P±PґC,PsPґ P°P±Ps PіPsCґCґPґPSPSCІPґ PґPґC,P°P±PsP»C-P·PґCґ
PіPґCґC-PsPrPëCґPSPs CÍP»C-Pr PePsPSC,CґPsP»CHPIP°C,Pë PiP»C-PePґPґC-CH P°P±Ps, Cґ
CґP°P·C- PSPґPsP±C...C-PrPSPsCÍC,C-, PіC-PrC-P±CґP°C,Pë P°PrPґPePIP°C,PSCґ PrPsP·Cґ
PiC-PіPsPiP»C-PePґPґC-CґPSPsPiPs P·P°CÍPsP±Cґ.

вҖґ PґCґPґPіP°CґP°C, CÍP»C-Pr P·P°CÍC,PsCÍPsPICґPIP°C,Pë P· PsP±PґCґPґPґPSC-CÍC,CH
C...PIPsCґPëPґ PSP° PіPsPrP°PiCґCґ, PsCÍPeC-P»CHPePë PіCґPґPіP°CґP°C, PґPsPґPґ
PіC-PrPIPëC%CґPIP°C,Pë CґC-PIPґPSCґ CÍPґCґPsPiPsC— PePëCÍP»PsC,Pë PI
CÍPëCґPsPIP°C,CґC- PeCґPsPIC-.

вҖґ PhCÍPeC-P»CHPePë PќPsCґPґP°C,PґPSCІ PґPsPґPґ CÍPіCґPëCґPëPSCІC,Pë
PiC-PіPsPeP°P»C-C”PґC-CH, PіP°CґC-C”PSC,P°Pґ PI C...PsPrC- P»C-PeCґPIP°PSPSCІ
CґPґPePsPґPґPSPґCґC,CHCÍCІ PIPґPëPIP°C,Pë C—PґCґ P· PIPëCÍPsPePëPґ PIPґC-CÍC,PsPґ
PeP°P»C-CH (C,,CґCґPeC,Pë, PsPiPsCґC-, CґPëP±P°) C- PіPґCґC-PsPrPëCґPSPs
PePsPSC,CґPsP»CHPIP°C,Pë CґC-PIPґPSCґ PeP°P»C-CH Cґ PeCґPsPIC-.

вҖґ P»C-PeP°CґCґCHPePëPNґ P·P°CÍC-P± CÍP»C-Pr P·P°CÍC,PsCÍPsPICґPIP°C,Pë P·
PsCÍPsP±P»PëPIPsCH PsP±PґCґPґPґPSC-CÍC,CH PіP°CґC-C”PSC,P°Pґ P· PSPëCґPePsPiPsCH
PSPґPrPsCÍC,P°C,PSC-CÍC,CH, PsCÍPeC-P»CHPePë PIC-PS PґPsPґPґ PіPsCÍPëP»PëC,Pë
PSPëCґPePsPICґ PSPґPrPsCÍC,P°C,PSC-CÍC,CH P· PsPrPSPsPiPs P±PsPeCґ, P° P· C-PSCëPsPiPs
- PNґPsPiPs PґC,,PґPeC,PëPIPSC-CÍC,CH PґPsPґPґ P·PSPëP·PëC,PëCÍCІ. PќPґ CÍP»C-Pr
P·P°CÍC,PsCÍPsPICґPIP°C,Pë PіCґPґPіP°CґP°C, PіP°CґC-C”PSC,P°Pґ C-P· PIPëCґP°PґPґPSPsCH
PіPґCґC-PSPePsPiPsCH C,P° PSPëCґPePsPiPsCH PSPґPrPsCÍC,P°C,PSC-CÍC,CH.

вҖґ PJ P·PiбҖ™ CІP·PeCґ P· PSP°CІPIPSC-CÍC,CH Cґ PіCґPґPіP°CґP°C,C- CÍP°C...P°CґPsP·Pë
PіP°CґC-C”PSC,P°Pґ P· CґCґPsPrPґPґPSPëPґPë PіPsCґCґPґPSPSCІPґPë,
PіPsPiбҖ™ CІP·P°PSPëPґPë P· PSPґPіPґCґPґPSPsCÍPëPґC-CÍC,CH C,,CґCґPeC,PsP·Pë,
CÍPëPSPґCґPsPґPґ PґP°P»CHP°P±CÍPsCґP±CґC-C— PiP»CHPePsP·Pë-PiP°P»P°PeC,PsP·Pë
P°P±Ps PrPґC,,C-CґPëC,PsPґ CÍP°C...P°CґP°P·Pë-C-P·PsPґP°P»CHC,P°P·Pë PSPґ CÍP»C-Pr
P·P°CÍC,PsCÍPsPICґPIP°C,Pë CґPґPNґ PіCґPґPіP°CґP°C,.

өҖґ PґCґPґPіP°CґP°C, PґC-CÍC,PëC,CH P»P°PeC,PsP·Cґ, C,PsPґCґ PSPґ CÍP»C-Pr PNґPsPiPs
P·P°CÍC,PsCÍPsPICґPIP°C,Pë PіP°CґC-C”PSC,P°Pґ P· CґCґPsPrPґPґPSPëPґPë
PіPsCґCґCґPґPSPSCІPґPë, PіPsPiбҖ™ CІP·P°PSPëPґPë P· PSPґPіPґCґPґPSPsCÍPëPґC-CÍC,CH
PіP°P»P°PeC,PsP·Pë, PrPґC,,C-CґPëC,PsPґ P»P°PeC,P°P·Pë P»P°PіPіP° P°P±Ps
CÍPëPSPґCґPsPґPґ PґP°P»CHP°P±CÍPsCґP±CґC-C— PiP»CHPePsP·Pë-PiP°P»P°PeC,PsP·Pë.

*PePμCTĤCžPIP°PSPSC- P°PIC,PsC,CTĤP°PSCÍPĩPsCTĤC,PsPj P°P±Ps C-PSC€PěPjPě
PjPμC...P°PSC-P·PjP°PjPě.*

*PkPsCTĤPjP°C,PμPSCÍ PIPĩP»PěPIP°C” PSP° P·PrP°C,PSC-CÍC,CH PePμCTĤCÍPIP°C,Pě
P°PIC,PsPjPsP±C-P»PμPj C,P° C-PSC€PěPjPě PjPμC...P°PSC-P·PjP°PjPě.*

*P—PsPeCTĤPμPjP°, PSP° PĩPsC†P°C,PeCÍ P»C-PeCÍPIP°PSPSCĬ C,P° PĩC-CÍP»CĬ
P·P±C-P»CHC€PμPSPSCĬ PrPsP·Pě PĩCTĤPμPĩP°CTĤP°C,CÍ PjPsP¶CÍC,CH PIPěPSPěPeP°C,Pě
P°CÍC,PμPSC-CĬ C,P° PsCTĤC,PsCÍC,P°C,PěC†PSP° PĩC-PĩPsC,PμPSP·C-CĬ, C%oPs
PsP±PjPμP¶CÍC” P·PrP°C,PSC-CÍC,CH PePμCTĤCÍPIP°C,Pě P°PIC,PsPjPsP±C-P»PμPj C,P°
C-PSC€PěPjPě PjPμC...P°PSC-P·PjP°PjPě.*

PŸPĩPsCÍC-P± P·P°CÍC,PsCÍCžPIP°PSPSCĬ C,P° PrPsP·Pě.

*PμCTĤPμPĩP°CTĤP°C, PjPsP¶PSP° P·P°CÍC,PsCÍPsPICÍPIP°C,Pě PI PePsPjP±C-PSP°C†C-C— P·
C-PSC€PěPjPě P°PSC,PěPĩC-PĩPμCTĤC,PμPSP·PěPIPSPěPjPě P·P°CÍPsP±P°PjPě, C,P°PePěPjPě
CĬPe P±PμC,P°-P±P»PsPeP°C,PsCTĤPě C,P° PIP°P·PsPrPěP»P°C,P°C,PsCTĤPě.*

*P”PsP·CÍPIP°PSPSCĬ PIPěP·PSP°C†P°C” P»C-PeP°CTĤ C-PSPrPěPIC-PrCÍP°P»CHPSPs PrP»CĬ
PePsP¶PSPsPiPs C...PIPsCTĤPsPiPs.*

*P PμPePsPjPμPSPrCÍC”C,CHCÍCĬ PĩPsC†PěPSP°C,Pě P»C-PeCÍPIP°PSPSCĬ P· 1
C,P°P±P»PμC,PePě 1 CTĤP°P· PSP° PrPsP±CÍ. P—P°P»PμP¶PSPs PIC-Pr
PμC,,PμPeC,PěPIPSsCÍC,C- P»C-PeCÍPIP°PSPSCĬ C,P° C,CĬP¶PePsCÍC,C- PĩPsP±C-C†PSPěC...
CTĤPμP°PeC†C-PNē PrPsP·P° PjPsP¶Pμ P±CÍC,Pě P·P±C-P»CHC€PμPSP° PrPs 3 C,P°P±P»PμC,PsPe
1 CTĤP°P· PSP° PrPsP±CÍ PĩCTĤPě P·P°CÍC,PsCÍCÍPIP°PSPSC- CÍ 3 PĩCTĤPěPNēPsPjPě. PŸPěPj PSPμ
PjPμPSC€ C,CTĤPμP±P° PĩP°PjBĤ™ CĬC,P°C,Pě, C%oPs, P·PIP°P¶P°CTĤC†Pě PSP° PjPμC...P°PSC-P·Pj
PrC-C— C,P° C,,P°CTĤPjP°PePsP»PsPiC-C†PSC- PIP»P°CÍC,PěPIPsCÍC,C- PsPeCTĤPμPjPěC...
P°PeC,PěPIPSpěC... CTĤPμC†PsPIPěPS PĩCTĤPμPĩP°CTĤP°C,CÍ, P»C-PeCÍPIP°PSPSCĬ PjPsP¶Pμ
PIPIP°P¶P°C,PěCÍCĬ PSPμPμC,,PμPeC,PěPIPSpěPj P»PěC€Pμ C†PμCTĤPμP·
PIC-PrCÍCÍC,PSC-CÍC,CH PĩPsP·PěC,PěPIPSsPiPs PμC,,PμPeC,CÍ C†PμCTĤPμP· 14 PrPSC-PI
PĩC-CÍP»CĬ PĩPsC†P°C,PeCÍ C,PμCTĤP°PiC-C—. PŸP°PePěPj C†PěPSPsPj, PrPsP·P° PSPμ PjPsP¶Pμ
P±CÍC,Pě P·P±C-P»CHC€PμPSP° P±PμP· PSP°PiP°P»CHPSPsC— PĩPsC,CTĤPμP±Pě.*

*PkC†C-PeCÍPIP°PSPěPNē C,PμCTĤP°PĩPμPIC,PěC†PSPěPNē PμC,,PμPeC,
CÍPĩPsCÍC,PμCTĤC-PiP°C”C,CHCÍCĬ PĩCTĤPěP±P»PěP·PSPs C†PμCTĤPμP· 1BĤ”4 C,PěP¶PSC-.*

P”PsP·CÍPIP°PSPSCĬ CÍP»C-Pr P·P±C-P»CHC€CÍPIP°C,Pě PĩPsCÍC,CÍPĩPsPiPs.

*P—P°PiP°P»CHPSP° PμC,,PμPeC,PěPIPSp° PrPsP·P° CÍC,P°PSPsPIPěC,CH 1 C,P°P±P»PμC,PeP° 1
CTĤP°P· PSP° PrPsP±CÍ. PhPrPSP°Pe CÍ PrPμCĬPePěC... PĩP°C†C-C”PSC,C-PI P·P°PiP°P»CHPSP°
PμC,,PμPeC,PěPIPSp° PrPsP·P° CÍC,P°PSPsPIPěC,CH 1 C,P°P±P»PμC,PeP° C†PμCTĤPμP·
PrPμPSCĤ. PμCTĤPě PIC-PrCÍCÍC,PSPsCÍC,C- P·P°PrPsPIC-P»CHPSPsPiPs
PiC-PĩPsC,PμPSP·PěPIPSsPiPs PμC,,PμPeC,CÍ PrPsP·CÍ PĩCTĤPμPĩP°CTĤP°C,CÍ C,CTĤPμP±P°
P·P±C-P»CHC€PěC,Pě PrPs 1 C,P°P±P»PμC,PePě PrPIC-C†C- PSP° PrPsP±CÍ, P° CÍ
PIPěPSCĬC,PePsPIPěC... PIPěPĩP°PrPeP°C... BĤ” PrPs 1 C,P°P±P»PμC,PePě C,CTĤPěC†C- PSP°
PrPsP±CÍ (PSPμ CÍP»C-Pr P·P°CÍC,PsCÍPsPICÍPIP°C,Pě PIPěC%oC- PrPsP·Pě).*

*PŸP°P±P»PμC,PeCÍ PSPμPsP±C...C-PrPSPs PĩCTĤPěPNēPjP°C,Pě PĩC-Pr C†P°CÍ P°P±Ps
PsPrCTĤP°P·CÍ PĩC-CÍP»CĬ C—PrPě.*

$P''PsP \cdot C \acute{z} PIP^{\circ} PSPSC \sqcup C \acute{z} P \ddot{I} P^{\circ} C \dagger C - C''PSC, C - PI P \gg C - C, PSCHPsPiPs PIC - PeC \acute{z}$. $P \acute{k} P_{\mu} P_j P^{\circ} C''$
 $PSP_{\mu} PsP \pm C \dots C - PrPSPsC \acute{I} C, C - C \acute{r} PePsC \ddot{H} P_{\mu} PeC \dagger C - C - PrPsP \cdot C \acute{r} PIP^{\circ} PSPSC \sqcup$. $P \acute{h} PrPSP^{\circ} Pe$
 $C \acute{I} P \gg C - Pr P \pm C \acute{r} C, P \ddot{e} PsP \pm P_{\mu} C \ddot{H} P_{\mu} P \P PSP \ddot{e} P_j C \ddagger P_{\mu} C \ddot{H} P_{\mu} P \cdot P \ddot{i} C - PrPIP \ddot{e} C \% P_{\mu} PSP \ddot{e} P N_{\circ} C \ddot{H} P \ddot{e} P \cdot P \ddot{e} Pe$
 $C \ddot{H} PsP \cdot PIP \ddot{e} C, PeC \acute{r} P^{\circ} C \ddot{H} C, P_{\mu} C \ddot{H} C - P^{\circ} P \gg CHPsPsC - P \ddot{i} C - P \ddot{i} PsC, P_{\mu} PSP \cdot C - C - C, P^{\circ}$
 $P \ddot{i} PsC \ddot{H} C \acute{r} C \epsilon P_{\mu} PSPSC \sqcup C \ddot{H} C - PIPSC \sqcup P_{\mu} P \gg P_{\mu} PeC, C \ddot{H} PsP \gg C - C, C - PI C \acute{r} PeC \ddot{H} PsPIC -$.

$P''PsP \cdot C \acute{z} PIP^{\circ} PSPSC \sqcup PrP \gg C \sqcup P \ddot{I} P^{\circ} C \dagger C - C''PSC, C - PI P \cdot PSP_{\mu} PrPsC \acute{I} C, P^{\circ} C, PSC - C \acute{I} C, C \ddot{H}$
 $P \ddot{i} P_{\mu} C \dagger C - PSPeP \ddot{e} C, P^{\circ} PSP \ddot{e} C \ddot{H} PsPe$. $P \acute{k} P_{\mu} C \acute{I} P \gg C - Pr P \cdot P^{\circ} C \acute{I} C, PsC \acute{I} PsPIC \acute{r} PIP^{\circ} C, P \ddot{e}$
 $P \ddot{i} C \ddot{H} P_{\mu} P \ddot{i} P^{\circ} C \ddot{H} P^{\circ} C, P \ddot{I} P^{\circ} C \dagger C - C''PSC, P^{\circ} P_j C - P \cdot C, C \sqcup P \P PePsC \ddot{H}$
 $PSP_{\mu} PrPsC \acute{I} C, P^{\circ} C, PSC - C \acute{I} C, C \ddot{H} P \ddot{i} P_{\mu} C \dagger C - PSPeP \ddot{e} C, P^{\circ} PSP \ddot{e} C \ddot{H} PsPe (PeP \gg C - C \ddot{H} P_{\mu} PSC \acute{I}$
 $PeC \ddot{H} P_{\mu} P^{\circ} C, P \ddot{e} PSC - PSC \acute{r} PSP \ddot{e} P \P C \ddagger P_{\mu} 30 P_j P \gg / C \dots PI)$.

$P''C - C, P \ddot{e}$. $P \acute{k} P_{\mu} P \cdot P^{\circ} C \acute{I} C, PsC \acute{I} PsPIC \acute{r} PIP^{\circ} C, P \ddot{e}$.

$P_{\mu} P_{\mu} C \ddot{H} P_{\mu} PrPsP \cdot C \acute{z} PIP^{\circ} PSPSC \sqcup$.

$P \ddot{Y} P \ddot{e} P_j P \ddot{i} C, PsP_j P \ddot{e} P \ddot{i} P_{\mu} C \ddot{H} P_{\mu} PrPsP \cdot C \acute{r} PIP^{\circ} PSPSC \sqcup P \gg C - PeP^{\circ} C \ddot{H} C \acute{I} C \ddot{H} PePsPiPs P \cdot P^{\circ} C \acute{I} PsP \pm C \acute{r}$:
 $PSC \acute{r} PrPsC, P^{\circ}, P \pm P \gg C \ddot{H} PIP^{\circ} PSPSC \sqcup, PrC - P^{\circ} C \ddot{H} P_{\mu} C \sqcup, P \ddot{i} PsP \gg PsPIPSP \ddot{e} P N_{\circ} P \pm C - P \gg CH,$
 $PIC - PrC \dagger C \acute{r} C, C, C \sqcup P \P P^{\circ} C \ddot{H} C \acute{r}, C \acute{I} P \ddot{i} C \ddot{H} P^{\circ} P \ddot{i} P^{\circ}, P \ddot{i} C - P \ddot{i} PsPeP^{\circ} P \gg C - C''P_j C - C \sqcup, P_j C - P^{\circ} C \acute{I} C, P_{\mu} PSC - C \sqcup,$
 $P^{\circ} C \ddot{H} C, P_{\mu} C \ddot{H} C - P^{\circ} P \gg CHPsP^{\circ} P \ddot{i} C - P \ddot{i} PsC, P_{\mu} PSP \cdot C - C \sqcup, P \pm C \ddot{H} P^{\circ} PrP \ddot{e} PeP^{\circ} C \ddot{H} PrC - C \sqcup,$
 $P \ddot{i} PsC \ddot{H} C \acute{r} C \epsilon P_{\mu} PSPSC \sqcup C \ddot{H} P \ddot{e} C, P_j C \acute{r} C \acute{I} P_{\mu} C \ddot{H} C \dagger C \sqcup, PrP_{\mu} P \ddot{i} C \ddot{H} P_{\mu} C \acute{I} C - C \sqcup,$
 $P \cdot P^{\circ} P \ddot{i} P^{\circ} P_j PsC \ddot{H} PsC \ddagger P_{\mu} PSPSC \sqcup, PePsP_j P^{\circ}, P \ddot{i} PsP \gg PsPIPSP \ddot{e} P N_{\circ} P \pm C - P \gg CH,$
 $C \acute{I} P \ddot{i} P \gg C \acute{r} C, P^{\circ} PSC - C \acute{I} C, CH C \acute{I} PIC - PrPsP_j PsC \acute{I} C, C -, P_j \mathbb{B} \ddot{H}^{\mathbb{T}} C \sqcup P \cdot PsPIP^{\circ} C \acute{I} P \gg P^{\circ} P \pm PeC - C \acute{I} C, CH$.

$P_{\mu} C - C \acute{I} P \gg C \sqcup P \ddot{i} P_{\mu} C \ddot{H} P_{\mu} PrPsP \cdot C \acute{r} PIP^{\circ} PSPSC \sqcup PSP_{\mu} PsP \pm C \dots C - PrPSPs PSP_{\mu} P \ddot{i} P^{\circ} P N_{\circ} PSPs$
 $P \cdot C \ddot{H} PsP \pm P \ddot{e} C, P \ddot{e} P \ddot{i} C \ddot{H} PsP_j P \ddot{e} PIP^{\circ} PSPSC \sqcup C \epsilon P \gg C \acute{r} PSPeP^{\circ} C, P^{\circ} P \cdot P^{\circ} C \acute{I} C, PsC \acute{I} PsPIC \acute{r} PIP^{\circ} C, P \ddot{e}$
 $P^{\circ} PeC, P \ddot{e} PIPsPIP^{\circ} PSP_{\mu} PIC \acute{r} P \ddot{i} C - P \gg P \gg C \sqcup$. $P \acute{k} P_{\mu} PsP \pm C \dots C - PrPSPs PePsPSC, C \ddot{H} PsP \gg C \ddot{H} PIP^{\circ} C, P \ddot{e}$
 $C, C \acute{r} PSPeC \dagger C - C \ddot{H} C \acute{I} P_{\mu} C \ddot{H} C \dagger C \sqcup, P^{\circ} C \ddot{H} C, P_{\mu} C \ddot{H} C - P^{\circ} P \gg CHPsP \ddot{e} P N_{\circ} C, P \ddot{e} C \acute{I} Pe C, P^{\circ} PIPsPrPSPs -$
 $P_{\mu} P \gg P_{\mu} PeC, C \ddot{H} PsP \gg C - C, PSC \acute{r} C \ddot{H} C - PIPSPsPIP^{\circ} P \ddot{i} C \acute{r}$. $P_{\mu} C \ddot{H} P \ddot{e} PSP_{\mu} PsP \pm C \dots C - PrPSPsC \acute{I} C, C -$
 $P \cdot P^{\circ} C \acute{I} C, PsC \acute{I} PsPIC \acute{r} PIP^{\circ} C, P \ddot{e} C \acute{I} P \ddot{e} P_j P \ddot{i} C, PsP_j P^{\circ} C, P \ddot{e} C \ddagger PSP_{\mu} P \gg C - PeC \acute{r} PIP^{\circ} PSPSC \sqcup$.

$P_{\mu} PsP \pm C - C \dagger PSC - C \ddot{H} P_{\mu} P^{\circ} PeC \dagger C - C -$.

$P \grave{L} C \ddot{H} P \ddot{e} C, P_{\mu} C \ddot{H} C - C - PsC \dagger C - PSPeP \ddot{e} C \ddagger P^{\circ} C \acute{I} C, PsC, P \ddot{e} C \ddot{H} PsP \cdot PIP \ddot{e} C, PeC \acute{r}$
 $P \ddot{i} PsP \pm C - C \dagger PSP \ddot{e} C \dots C \ddot{H} P_{\mu} P^{\circ} PeC \dagger C - P N_{\circ} P \ddot{i} C \ddot{H} P \ddot{e} P \cdot P^{\circ} C \acute{I} C, PsC \acute{I} C \acute{r} PIP^{\circ} PSPSC -$
 $P \gg C - PeP^{\circ} C \ddot{H} C \acute{I} C \ddot{H} PePsPiPs P \cdot P^{\circ} C \acute{I} PsP \pm C \acute{r}$:

$P''C \acute{z} P \P P_{\mu}$	$\geq 1/10$
$C \ddagger P^{\circ} C \acute{I} C, Ps:$	
$P \S P^{\circ} C \acute{I} C, Ps:$	$\geq 1/100 \text{ б} \ddot{H}'' < 1/10$
$P \acute{k} P_{\mu} C \ddagger P^{\circ} C \acute{I} C, Ps:$	$\geq 1/1000 \text{ б} \ddot{H}'' < 1/100$
$P C - PrPePs:$	$\geq 1/10000 \text{ б} \ddot{H}'' < 1/1000$
$P''C \acute{z} P \P P_{\mu}$	$< 1/10000$
$C \ddot{H} C - PrPePs:$	
$P \acute{k} P_{\mu} PIC - PrPsP_j Ps:$	$C \ddagger P^{\circ} C \acute{I} C, PsC, C \acute{r} PIP \ddot{e} P \ddot{i} P^{\circ} PrPeC - PI PSP_{\mu} P_j PsP \P PSP^{\circ}$ $PsC \dagger C - PSP \ddot{e} C, P \ddot{e} P \cdot P^{\circ} PSP^{\circ} C \sqcup PIPSP \ddot{e} P_j P \ddot{e} PrP^{\circ} PSP \ddot{e} P_j P \ddot{e}$

$P - P \pm PsPeC \acute{z} C \epsilon P \gg C \acute{z} PSPePsPIPs - PeP \ddot{e} C \epsilon PePsPIPsPiPs \ C, C \eta P^{\circ} PeC, C \acute{z}$: $PSP_{\mu}PIC - PrPsPjPs$ бҗ“ $PSCrPrPsC, P^{\circ} \ C, P^{\circ} \ P \pm P \gg C \eta PIP^{\circ} PSPSC \sqcup$.

$P - P \pm PsPeC \acute{z} PjP_{\mu}C, P^{\circ} P \pm PsP \gg C - P \cdot PjC \acute{z}$: $PSP_{\mu}PIC - PrPsPjPs$ бҗ“ $PiC - P \ddot{i}PsPeP^{\circ}P \gg C - C''PjC - C \sqcup$.

$P - P \pm PsPeC \acute{z} C \acute{I}P_{\mu}C \eta C \dagger P_{\mu}PIPs - C \acute{I}C \acute{z} PrP \ddot{e} PSPSPsC - C \acute{I}P \ddot{e} C \acute{I}C, P_{\mu}PjP \ddot{e}$: $PSP_{\mu}PIC - PrPsPjPs$ бҗ“ $PsC \eta C, PsC \acute{I}C, P^{\circ}C, P \ddot{e}C \ddagger PSP^{\circ} \ PiC - P \ddot{i}PsC, P_{\mu}PSP \cdot C - C \sqcup$.

$P - P \pm PsPeC \acute{z} PSP_{\mu}C \eta PIPsPIPsC - C \acute{I}P \ddot{e} C \acute{I}C, P_{\mu}PjP \ddot{e}$: $PSP_{\mu}PIC - PrPsPjPs$ бҗ“ $PjC - P^{\circ}C \acute{I}C, P_{\mu}PSC - C \sqcup$.

$P - P \pm PsPeC \acute{z} C \eta P_{\mu}C \acute{I}P \ddot{i}C - C \eta P^{\circ}C, PsC \eta PSPsC - C \acute{I}P \ddot{e} C \acute{I}C, P_{\mu}PjP \ddot{e}$: $PSP_{\mu}PIC - PrPsPjPs$ бҗ“ $C \eta P \ddot{e}PSC - C, .$

$P \triangleright P^{\circ}P \pm PsC \eta P^{\circ}C, PsC \eta PSC - P \ddot{i}PsPeP^{\circ}P \cdot PSP \ddot{e}PeP \ddot{e}$: $PSP_{\mu}PIC - PrPsPjPs$ бҗ“ $PiC - P \ddot{i}P_{\mu}C \eta CrC \eta P \ddot{e}PeP_{\mu}PjC - C \sqcup \ C, P^{\circ} \ PiC - P \ddot{i}P_{\mu}C \eta PiP \gg C - PeP_{\mu}PjC - C \sqcup$.

$P - P^{\circ}PiP^{\circ}P \gg CHbPSC - P \ddot{i}PsC \eta C \acute{z} C \epsilon P_{\mu}PSPSC \sqcup$: $PSP_{\mu}PIC - PrPsPjPs$ бҗ“ $P^{\circ}C \acute{I}C, P_{\mu}PSC - C \sqcup$.

$P - P \pm PsPeC \acute{z} C \acute{I}P \ddot{e} C \acute{I}C, P_{\mu}PjP \ddot{e} \ PeC \eta PsPIPsC, PIPsC \eta P_{\mu}PSPSC \sqcup$: $PSP_{\mu}PIC - PrPsPjPs$ бҗ“ $C, C \eta PsPjP \pm PsC \dagger P \ddot{e}C, PsP \ddot{i}P_{\mu}PSC - C \sqcup$.

$P - P \pm PsPeC \acute{z} PiC \acute{I}P \ddot{e}C \dots C - PeP \ddot{e}$: $PSP_{\mu}PIC - PrPsPjPs$ бҗ“ $PrP_{\mu}P \ddot{i}C \eta P_{\mu}C \acute{I}C - C \sqcup$.

$P \ddot{y}P^{\circ}PePsP \P PjPsP \P CrC, CHb \ C \acute{I}P \ddot{i}PsC \acute{I}C, P_{\mu}C \eta C - PiP^{\circ}C, P \ddot{e}C \acute{I}C \sqcup \ P \ddot{i}PsP \pm C - C \ddagger PSC - C \eta P_{\mu}P^{\circ}PeC \dagger C - C -, \ C \dots P^{\circ}C \eta P^{\circ}PeC, P_{\mu}C \eta PSC - PrP \gg C \sqcup \ C \eta P_{\mu}P \cdot P_{\mu}C \eta P \ddot{i}C - PSCr$.

$P - P \pm PsPeC \acute{z} C \epsilon P \gg C \acute{z} PSPePsPIPs - PeP \ddot{e} C \epsilon PePsPIPsPiPs \ C, C \eta P^{\circ}PeC, C \acute{z}$: $PrC - P^{\circ}C \eta P_{\mu}C \sqcup, C \acute{I}CrC \dots C - C \acute{I}C, CHb \ Cr \ C \eta PsC, C -, \ PiC - PrPIP \ddot{e}C \%_{\circ} P_{\mu}PSPSC \sqcup \ C \acute{I}P_{\mu}PeC \eta P_{\mu}C \dagger C - C - C \epsilon P \gg CrPSPePsPIPsPiPs \ C \acute{I}PsPeCr, \ PiC - PrPIP \ddot{e}C \%_{\circ} P_{\mu}PSP_{\mu}C \acute{I}P \gg P \ddot{e}PSPsPIP \ddot{e}PrC - P \gg P_{\mu}PSPSC \sqcup, \ PiC - PrPIP \ddot{e}C \%_{\circ} P_{\mu}PSPSC \sqcup \ P^{\circ}P \ddot{i}P_{\mu}C, P \ddot{e}C, Cr, PIP \ddot{e}C \eta P^{\circ}P \cdot PePsPIP^{\circ} \ C \dots PIPsC \eta PsP \pm P^{\circ}, \ C \epsilon P \gg CrPSPePsPIPs - PeP \ddot{e} C \epsilon PePsPIP^{\circ} \ PeC \eta PsPIPsC, P_{\mu}C \ddagger P^{\circ}, \ P \pm C - P \gg CHb \ Cr \ P \P P \ddot{e}PIPsC, C -.$

$P - P \pm PsPeC \acute{z} PjP_{\mu}C, P^{\circ}P \pm PsP \gg C - P \cdot PjC \acute{z}$: $PiC - P \ddot{i}PsPeP^{\circ}P \gg CHbC \dagger C - C''PjC - C \sqcup, PjP_{\mu}C, P^{\circ}P \pm PsP \gg C - C \ddagger PSP \ddot{e}PN_{\circ} \ P^{\circ}P \gg PeP^{\circ}P \gg PsP \cdot$.

$P - P \pm PsPeC \acute{z} C \acute{I}P_{\mu}C \eta C \dagger P_{\mu}PIPs - C \acute{I}C \acute{z} PrP \ddot{e} PSPSPsC - C \acute{I}P \ddot{e} C \acute{I}C, P_{\mu}PjP \ddot{e}$: $C \acute{I}P \ddot{e}PSCrC \acute{I}PsPIP^{\circ} \ P \pm C \eta P^{\circ}PrP \ddot{e}PeP^{\circ}C \eta PrC - C \sqcup, \ P^{\circ}C \eta P \ddot{e}C, PjC - C \sqcup \ C \acute{I}P_{\mu}C \eta C \dagger C \sqcup, \ C \acute{I}P \ddot{e}PjPiC, PsPjP \ddot{e}C \acute{I}C, P_{\mu}PSPsPeP^{\circ}C \eta PrC - C -, \ P^{\circ}C \eta C, P_{\mu}C \eta C - P^{\circ}P \gg CHbPSP^{\circ} \ PiC - P \ddot{i}PsC, P_{\mu}PSP \cdot C - C \sqcup, \ P \ddot{i}C \eta P \ddot{e}P \ddot{i}P \gg P \ddot{e}PIP \ddot{e}, \ PIC, C \eta P^{\circ}C, P^{\circ} \ C \acute{I}PIC - PrPsPjPsC \acute{I}C, C -, \ C \acute{I}P_{\mu}C \eta C \dagger P_{\mu}PIP^{\circ} \ PSP_{\mu}PrPsC \acute{I}C, P^{\circ}C, PSC - C \acute{I}C, CHb, \ C \dagger P_{\mu}C \eta P_{\mu}P \pm C \eta PsPIP^{\circ}C \acute{I}PeCrP \gg C \sqcup C \eta PSC - P \ddot{i}PsC \eta CrC \epsilon P_{\mu}PSPSC \sqcup$.

$P - P \pm PsPeC \acute{z} C \dagger P_{\mu}PSC, C \eta P^{\circ}P \gg CHbPSPsC - PSP_{\mu}C \eta PIPsPIPsC - C \acute{I}P \ddot{e} C \acute{I}C, P_{\mu}PjP \ddot{e}$: $P \cdot P^{\circ}P \ddot{i}P^{\circ}PjPsC \eta PsC \ddagger P_{\mu}PSPSC \sqcup, \ PrC \eta P^{\circ}C, C - PIP \gg P \ddot{e}PIC - C \acute{I}C, CHb, \ PePsC \epsilon PjP^{\circ}C \eta PSC - C \acute{I}PSPsPIP \ddot{e}PrC - PSPSC \sqcup, \ PiC - PrPIP \ddot{e}C \%_{\circ} P_{\mu}PSP^{\circ} \ C \acute{I}C, PsPjP \gg C \eta PIP^{\circ}PSC - C \acute{I}C, CHb, \ P_{\mu}PeC \acute{I}C, C \eta P^{\circ}PiC - C \eta P^{\circ}PjC - PrPSC - P \ddot{i}PsC \eta CrC \epsilon P_{\mu}PSPSC \sqcup \ (PIPeP \gg CHbC \ddagger P^{\circ}CHbC \ddagger P \ddot{e}P \ddot{i}P^{\circ}C \eta PeC - PSC \acute{I}PsPSC - P \cdot Pj), \ PiPsP \gg PsPIPSP \ddot{e}PN_{\circ} \ P \pm C - P \gg CHb, \ C \acute{I}C, P^{\circ}PS \ C, C \eta P \ddot{e}PIPsPiP \ddot{e},$

PiPsC7CfC€PµPSPSC¶ PePsPSC†PµPSC,C7P°C†C-C— C†PIP°PiPë, C†C,C†PiPsC7,
PiPsC7CfC€PµPSPSC¶ PsC7C-C''PSC,P°C†C-C—, PSP°P±C7C¶Pe PiPsP»PsPIPSPsPiPs
PjPsP·PeC†, C†PsPSP»PëPIC-C†C,CH, P±PµP·C†PsPSPSC¶.

P— P±PsPeCž C7PµC†PïC-C7P°C,PsC7PSPsC— C†PëC†C,PµPjPë: C7PëPSC-C,, PSP°P±C7C¶Pe
C†P»PëP·PsPIPsC— PsP±PsP»PsPSPePë PiPsC7PsP¶PSPëPSPë PSPsC†P°, P·P°PrPëC€PeP°,
PSPsC†PsPIP° PeC7PsPIPsC,PµC±P°.

P— P°PiP°P»CHBPSC- PiPsC7CžC€PµPSPSC¶: PSP°P±C7C¶PePë, C†P»P°P±PeC-C†C,CH.

P— P±PsPeCž PeC7PsPIC- C,P° P»C-PjC,,P°C,PëC±PSPsC— C†PëC†C,PµPjPë: P°PSPµPjC-C¶.

P— P±PsPeCž PµPSPPrPsPeC7PëPSPSPsC— C†PëC†C,PµPjPë: PiC-PrPIPëC%PµPSPSC¶
C†PµPeC7PµC†C-C— PiC7PsP»P°PeC,PëPSC†, PiP°P»P°PeC,PsC7PµC¶,
PiC-PSPµPePsPjP°C†C,C-C¶, PSP°P±C†C...P°PSPSC¶ PjPsP»PsC±PSPëC... P·P°P»PsP·.

P— P±PsPeCž PsP±PjC-PSCž C7PµC±PsPIPëPS C,P° C...P°C7C±CžPIP°PSPSC¶:
P·P±C-P»CHC€PµPSPSC¶ PjP°C†Pë C,C-P»P°.

P— P±PsPeCž PsC7PiP°PSC-PI C±CžC,C,C¶: PSPµC±C-C,PeC-C†C,CH P·PsC7C†,
PiC-PiPµC7PµPjC-C¶ PePsPSB7™ C7PSPeC,PëPIPë, C†P»CHPsP·PsC,PµC±P°,
PiPsC7CfC€PµPSPSC¶ C†P»C†C...C†.

P— P±PsPeCž C€PeC-C7Pë C,P° PiC-PrC€PeC-C7PSPsC— PeP»C-C,PePsPIPëPSPë: PµPeP·PµPjP°,
C†PIPµC7P±C-P¶, PiC†C7PiC†C7P°, PIPëC†PëPiP°PSPSC¶.

P— P±PsPeCž PSPëC7PsPe C,P° C†PµC±PsPIPëPIC-PrPSPëC... C€P»C¶C...C-PI: PrPëP·C†C7C-C¶,
PiP»PsPjPµC7C†P»PsPSPµC,,C7PëC.,

P— P±PsPeCž C7PµPiC7PsPrCžPeC,PëPIPSPsC— C†PëC†C,PµPjPë: P·PSPëP¶PµPSPSC¶
P»C-P±C-PrPs, PiPsC7CfC€PµPSPSC¶ PiPsC,PµPSC†C-C— C,P° PµC¶PeC†P»C¶C†C-C—.

P— P±PsPeCž PsPiPsC7PSPs-C7CžC...PsPIPsPiPs P°PiP°C7P°C,Cž C,P° C†PiPsP»CžC±PSPsC—
C,PeP°PSPëPSPë: PiPsC†C,C†C7P°P»CHBPSC- PiPsC7CfC€PµPSPSC¶.

P±C-P±C7PsP·PSC- P·PjC-PSPë C†PiPsC†C,PµC7C-PiP°P»Pë C† PrPµC¶PePëC...
PiP°C†C-C''PSC,C-PI, C¶PeC- C7PµPiC†P»C¶C7PSPs PiC7PëPµPjP°P»Pë PiC7PµPiP°C7P°C,Pë
P°P»PeP°P»PsC—PrC-PI C7C-P¶PeC-PI PiC7PsC,C¶PiPsPj PeC-P»CHPePsC... C7PsPeC-PI,
P·PsPeC7PµPjP°, C† PiP»PµPIC7C- C,P° C7PµC,C7PsPiPµC7C-C,PsPSPµP°P»CHPSPëC...
C,PeP°PSPëPSP°C..., C,P°PePsP¶ PI PrPµC¶PePëC... PIPëPiP°PrPeP°C...
C†PiPsC†C,PµC7C-PiP°P»Pë C,,C-P±C7PsP·PSC- P·PjC-PSPë PeP»P°PiP°PSC-PI C†PµC7C†C¶
(PrPëPI. C,P°PePsP¶ C7PsP·PrC-P» B«PhC†PsP±P»PëPIPsC†C,C-
P·P°C†C,PsC†C†PIP°PSPSC¶B»).

PjPµC7PjC-PS PiC7PëPrP°C,PSPsC†C,C-. 5 C7PsPeC-PI.

PjPjPsPIPë P·P±PµC7C-PiP°PSPSC¶. P—P±PµC7C-PiP°C,Pë PI PsC7PëPiC-PSP°P»CHBPSC-Pµ
C†PiP°PePsPIC†C- C† P·P°C...PëC%PµPSPsPjC† PIC-Pr C†PIC-C,P»P° C,P° PIPsP»PsPiPë
PjC-C†C†C- PiC7Pë C,PµPjPiPµC7P°C,C†C7C- PSPµ PIPëC%Pµ 25 B°PŸ. P—P±PµC7C-PiP°C,Pë

Cř PSPµPrPsCřC,CřPİPSPsPjCř PrP»Cİ PrC-C,PµPN PjC-CřC†C-.

PJPİP°PePsPIPeP°. PµPs 20 C,P°P±P»PµC,PsPe Cř P±P»C-CřC,PµCřC-; PİPs 1
P±P»C-CřC,PµCřCř PI CřPİP°PePsPIC†C-.

PљP°C,PµPiPsCřC-Cİ PIC-PrPİCřCřPeCř. P—P° CřPµC†PµPİC,PsPj.

P’PëCřPsP±PSPëPe.

PђPN-PŸC--P•PS PµPsP»CHC„P° P-PµC€CřPI PђPŸ / ICN Polfa Rzeszow S.A..

**PњC-CřC†PµP•PSP°C...PsPrP¶PµPSPSCİ PIPëCřPsP±PSPëPeP° C,P° PNPsPiPs
P°PrCřPµCřP° PjC-CřC†Cİ PİCřPsPIP°PrP¶PµPSPSCİ PrC-CİP»CHPSPsCřC,C-.**

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