

Р†РќРЎРЎР РЈРЉР†Р†РЇ

**РrР»CП РjРµРrРёC‡PSPsPiPs Р•Р°CГC,PsCГCрPIР°PSPSCП Р»C-РёР°CГCГЧРёPsPiPs
Р•Р°CГPsP±Cр**

Р”РЈРћР”РћР РЎ

В (DUODARTВ„Ў)

РЎРёР»Р°Рr:

*РrC-CГC‡C- CГРµC‡PsPIРёPSPё: РrCрC,Р°CГC,РµCГРёРr, C,Р°РjCГCрP»PsP•РёPSCр
PiC-РrCГPsC...P»PsCГРёРr;*

1 РёР°РёCГCрP»Р° РjC-CГC,РёC,ЧР РrCрC,Р°CГC,РµCГРёРrCр 0,5 РjPi C,Р°
C,Р°РjCГCрP»PsP•РёPSCр PiC-РrCГPsC...P»PsCГРёРrCр 0,4 РjPi;

***РrPsPiPsPjC-Р¶PSC- CГРµC‡PsPIРёPSPё: РjPsPSPsРrРёPiP»C-C‡РµCГРёРrРё
РёРёCГP»PsC,Рё РёР°РёCГРёР»PsPIPsC—,
Р±CрC,РёР»PiC-РrCГPsРёCГРёC,PsP»CрPsP» (Р• 321), Р¶РµР»Р°Р°Р°Р°Р°Р°Р°Р°,
PiP»C-C‡РµCГРёРrP, C,РёC,Р°PSCр РrC-PsРёCГРёРr (Р• 171), Р•Р°Р»C-Р•Р°
PsРёCГРёРr Р¶PsPIC,РёPNё (Р• 172), C,CГРёРiP»C-C‡РµCГРёРrРё
CГРµCГРµРrPSCЧPsPiPs Р»Р°PSC‡ЧРiР° C,Р° Р»РµC‡РёC,РёPS;
C‡РµР»CГР»PsP•Р° РjC-РёCГPsРёCГРёC,Р°Р»C-С‡PSP°,
РjРµC,Р°РёCГРёР»Р°C,PSPsPiPs CГPsPiPsP»C-РjРµCГCр РrРёCГРiРµCГC-СП,
C,Р°Р»ЧРё, C,CГРёРµC,РёР»C‡РёC,CГР°C,;***

***PsP±PsP»PsPSPёР° C,PIРµCГРrPsC— РёР°РёCГCзP»Рё: РёР°CГР°PiC-PSP°PS
(Р• 407), РёР°Р»C-СР С...P»PsCГРёРr, C,РёC,Р°PSCр РrC-PsРёCГРёРr (Р• 171),
FD&C Yellow 6 (E 110), PiC-PiCГPsPjРµP»PsP•Р°, PIC-CГРё
РёР°CГPSP°CрP±CГЧРёРёPNё, РёCГPsC...РjР°Р»ЧР
РёCрРёCГCГРrР•CПSPёPNё, Р•Р°Р»C-Р•Р° PsРёCГРёРr C‡РµCГPIPsPSPёPNё (Р•
172), SW-9008 Black Ink (СРµР»Р»Р°Рё, РiCГPsPiC-Р»РµPSPiP»C-РёPsP»ЧР,
Р•Р°Р»C-Р•Р° PsРёCГРёРr C‡PsCГPSPёPNё (Р• 172), РёР°Р»C-СР
PiC-РrCГPsРёCГРёРr).***

Р»C-РёР°CГCГЧРёР° C„PsCГРjР°. РЉР°РёCГCрP»Рё C,PIРµCГРrC—.

РћCГPSPsPIPsC- C„C-Р•РёРёPs-C...C-РjC-С‡PSC- PIР»Р°CГC,РёPIPsCГC,C-: C,PIРµCГРrР°

СГРµС±PSPiPs PjC-C...CpCП°. PµCПP±P»PëP·PSPs 75 % CПPµC†PµPïC,PsCПC-PI α₁ Cг PïPµCПPµPrPjC-C...CpCПPSPIC-PNë P·P°P»PsP·C- C” CПPµC†PµPïC,PsCПP°PjPë PïC-PrC,PëPïCг α_{1P°}.

PÿP°PjCГCзP»PsP·PëPS

PÿP°PjCГCгP»PsP·PëPS PïC-PrPIPëC%oCгC” PjP°PëCГPëPjP°P»CHBPSCr C€PIPëPrPëC-CГC,CH C,PsPëCг CГPµC±C- P·P° CПP°C...CгPSPsPë P·PSPëP¶PµPSPSCИ C,PsPSCгCГCг PïP»P°PrPëPëC... Pj'CИP·C-PI CгCПPµC,CПPë C- PïPµCПPµPrPjC-C...CpCПPSPiPsC— P·P°P»PsP·Pë, C%oPs CгCГCгPIP°C” PsP±CГC,CПCгPëC†C-CH. PµCПPµPïP°CПP°C, C,P°PëPsP¶ P·PSPëP¶CгC” PIPëCПP°P¶PµPSC-CГC,CH CГPëPjPïC,PsPjC-PI PïPsPrCПP°P·PSPµPSPSCИ C,P° PsP±CГC,CПCгPëC†C-C—, Cг CПPsP·PIPëC,PëCг CИPëPëC... C-CГC,PsC,PSCr CПPsP»CH PIC-PrC-PiCПP°C” PSPµC,CПPëPjP°PSPSCИ CГPµC±C- C,P° CГPëPsCПPsC±PµPSPSCИ PïP»P°PrPëPëC... PjBП™ CИP·C-PI PSPëP¶PSC-C... PIC-PrPrC-P»C-PI CГPµC±PSPiPIC-PrPSPëC... C€P»CИC...C-PI. PÿP°PëPëPNë PµC,,PµPëC, PrPsCГCИPïP°C”C,CHCГCИ PïC-Pr C±P°CГ PrPsPIPiPsC,CПPëPIP°P»PsC— C,PµCПP°PïC-C—. PкPµPsP±C...C-PrPSC-CГC,CH PsPïPµCПP°C,PëPIPSPiPs PIC,CПCгC±P°PSPSCИ P°P±Ps PëP°C,PµC,PµCПPëP·P°C†C-C— P·PSP°C±PSPs P·PSPëP¶CгC”C,CHCГCИ.

PПPSC,P°PiPsPSC-CГC,Pë α₁-P°PrCПPµPSPsCПPµC†PµPïC,PsCПC-PI PjPsP¶CгC,CH P·PjPµPSC€CгPIP°C,Pë PëCПPsPiBП™ CИPSPëPNë C,PëCГPë C€P»CИC...PsPj P·PSPëP¶PµPSPSCИ P·P°PiP°P»CHBSPSPiPs PïPµCПPëC,,PµCПPëC±PSPSPiPs PsPïPsCПCг. PµC-Pr C±P°CГ PrPsCГP»C-PrP¶PµPSPSCИ PIPïP»PëPICг C,P°PjCГCгP»PsP·PëPSCг PSPµ CГPiPsCГC,PµCПC-PiP°P»PsCГCH PëP»C-PSC-C±PSPs P·PSP°C±CгC%oPsPiPs P·PSPëP¶PµPSPSCИ PëCПPsPiBП™ CИPSPSPiPs C,PëCГPëCг.

PαP°CПPjP°PëPsPëC-PSPµC,PëPëP°.

PПC-P¶ PIPiPµPrPµPSPSCИPj PëPsPjP±C-PSP°C†C-C— PrCгC,P°CГC,PµCПPëPr- C,P°PjCГCгP»PsP·PëPS C- PsPrPSPsC±P°CГPSPëPj PIPiPµPrPµPSPSCИPj PrPsP· PrCгC,P°CГC,PµCПPëPrCг C- C,P°PjCГCгP»PsP·PëPSCг PI PëP°PïCГCгP»P°C... PsPëCПPµPjPs P±CгP»P° PïCПPsPrPµPjPsPSCГC,CПPsPIP°PSP° P±C-PsPµPëPIC-PIP°P»PµPSC,PSC-CГC,CH.

P”PsCГP»C-PrP¶PµPSPSCИ P±C-PsPµPëPIC-PIP°P»PµPSC,PSPsCГC,C- PsPrPSPsCПP°P·PsPIPëC... PrPsP· P±CгP»Ps PïCПPsPIPµPrPµPSPs CИPë PSP°C,C%oPµ, C,P°Pë C- PïC-CГP»CИ PïCПPëPNëPsPjCг C—P¶C-. Pj PïPsCПC-PIPSCИPSPSC- P·C- CГC,P°PSPsPj PSP°C,C%oPµ, Cг CПP°P·C- P·P°CГC,PsCГCгPIP°PSPSCИ PïC-CГP»CИ PïCПPëPNëPsPjCг C—P¶C- CГPiPsCГC,PµCПC-PiP°P»PsCГCИ 30 % P·PSPëP¶PµPSPSCИ C_{max} (PjP°PëCГPëPjP°P»CHBSPSPsC— PëPsPSC†PµPSC,CПP°C†C-C—) C,P°PjCГCгP»PsP·PëPSCг Cг CГPëP»P°PrC- PëPsPjP±C-PSP°C†C-C— PrCгC,P°CГC,PµCПPëPr-C,P°PjCГCгP»PsP·PëPS. P±P¶P° PSPµ PIPïP»PëPIP°P»P° PSP° AUC (PïP»PsC%oCг PïC-Pr C,,P°CПPjP°PëPsPëC-PSPµC,PëC±PSPsCП PëCПPëPIPsCH) C,P°PjCГCгP»PsP·PëPSCг.

P’CГPjPsPëC,CзPIP°PSPSCИ

P”CгC,P°CГC,PµCПPëPr

PµC-CГP»CИ PïPµCПPsCПP°P»CHBSPSPiPs PIPiPµPrPµPSPSCИ PsPrPSPsCПP°P·PsPIPsC— PrPsP·Pë 0,5 PjPi PrCгC,P°CГC,PµCПPëPrCг C±P°CГ PrPs PrPsCГCИPiPSPµPSPSCИ PNëPsPiPs

$$\text{Pj} \cdot \text{C} \ddagger \text{PsP} \gg \text{PsPIC} - \text{PeC} - \text{PI} \cdot \text{C}, \text{P}^\circ \text{Pj} \text{C} \dot{\text{I}} \text{C} \dot{\text{r}} \text{P} \gg \text{PsP} \cdot \text{P} \ddot{\text{e}} \text{PS} \cdot \text{P} \cdot \text{PI} \text{B} \text{H}^{\text{TM}} \text{C} \dot{\text{I}} \text{P} \cdot \text{C} \dot{\text{r}} \text{C}'' \text{C}, \text{C} \text{H} \text{C} \dot{\text{I}} \text{C} \dot{\text{I}} \text{P} \cdot \text{P} \pm \text{C} - \text{P} \gg \text{PeP}^\circ \text{Pj} \text{P} \ddot{\text{e}}$$

PiP»P°P·PjPë PiCTjPëP±P»PëP·PSPs PSP° 99 %. PhP±Bj™ C”Pj CTjPsP·PiPsPrC-P»Cr C”
PSPµPIPµP»PëPePëPj (PiCTjPëP±P»PëP·PSPs 0,21/PePi).

PhPµC, P°P±PsP»C-P·Pj

P”CrC, P°CrC, PµCTjPëPr

P”CrC, P°CrC, PµCTjPëPr P°PeC, PëPIPSPs PjPµC, P°P±PsP»C-P·CrC”C, CHCrCj in vivo. P’
CrPjPsPIP°C... in vitro PrCrC, P°CrC, PµCTjPëPr PjPµC, P°P±PsP»C-P·CrC”C, CHCrCj
C†PëC, PsC...CTjPsPjPsPjB P450 3A4 C- 3A5, CrC, PIPsCTjCTjCTjC†Pë C, CTjPë
PjPsPSPsPiC-PrCTjPsPeCrPëP»CHPsPIP°PSPëC... PjPµC, P°P±PsP»C-C, Pë C- PsPrPëPS
PrPëPiC-PrCTjPsPecPëP»CHPsPIP°PSPëPNë PjPµC, P°P±PsP»C-C,.

PµC-CrP»Cj PiPµCTjPsCTjP°P»CHPsPSPsPiPs PIPiPµPrPµPSPSCj PrCrC, P°CrC, PµCTjPëPrCr PI
PrPsP·C- 0,5 PjPi/PrPsP±Cr PrPs PrPsCrCjPiPSPµPSPSCj CTjC-PIPSPsPIP°P¶PSPsC—
PePsPSC†PµPSC, CTjP°C†C-C— 1,0Bj”15,4 % (CrPµCTjPµPrPSC” P·PSP°C†PµPSPSCj Bj” 5,4 %)
PIPiPµPrPµPSPsC— PrPsP·Pë PrCrC, P°CrC, PµCTjPëPrCr PIPëPrC-P»CjC”C, CHCrCj Cr PeP°P»
CrB PSPµP·PjC-PSPµPSPsPjCrB PIPëPiP»CjPrC-.B P PµC€C, P° PIPëPrC-P»CjC”C, CHCrCj Cr
PeP°P» Cr PIPëPiP»CjPrC- 4 PsCrPSPsPIPSPëC... PjPµC, P°P±PsP»C-C, C-PI, C%oPs
PjC-CrC, CjC, CH 39 %, 21 %, 7 % C- 7 % PePsP¶PSPsC— P· CTjPµC†PsPIPëPS,
PiPsPiBj™ CjP·P°PSPëC... P· P»C-PeP°CTjCrCHPePëPj P·P°CrPsP±PsPj, C- 6
PrCTjCrPiPsCTjCjPrPSPëC... PjPµC, P°P±PsP»C-C, C-PI (PjPµPSC€Pµ 5 % PePsP¶PµPS). Pj
CrPµC†C- P»CHPrPëPSPë PIPëCjPiP»PµPSPs P»PëC€Pµ PSPµP·PSP°C†PSC-
PeC-P»CHPePsCrC, C- PSPµP·PjC-PSPµPSPsPiPs PrCrC, P°CrC, PµCTjPëPrCr (PjPµPSC€Pµ PSC-P¶
0,1 % PrPsP·Pë).

PjP°PjCrCrP»PsP·PëPS

P·PSP°PSC, C-PsPjPµCTjPSP° P±C-PsPePsPSPIPµCTjCrC-Cj P· C, P°PjCrCrP»PsP·PëPSCr
PiC-PrCTjPsC...P»PsCTjPëPrCr [R(-) C-P·PsPjPµCTj] Cr S() C-P·PsPjPµCTj Cr P»CHPrPëPSPë PSPµ
PIC-PrP±CrPIP°C”C, CHCrCj. PjP°PjCrCrP»PsP·PëPSCr PiC-PrCTjPsC...P»PsCTjPëPr
P°PeC, PëPIPSPs PjPµC, P°P±PsP»C-P·CrC”C, CHCrCj C,, PµCTjPjPµPSC, P°PjPë
C†PëC, PsC...CTjPsPjCr P450 Cr PiPµC†C-PSC†C-, C- PjPµPSC€Pµ 10 % PrPsP·Pë
PIPëPrC-P»CjC”C, CHCrCj Cr CrPµC†Cr PI PSPµP·PjC-PSPµPSPsPjCr PIPëPiP»CjPrC-. PhP»Pµ
C,, P°CTjPjP°PePsPeC-PSPµC, PëC†PSPëPNë PiCTjPsC,, C-P»CH PjPµC, P°P±PsP»C-C, C-PI Cr
P»CHPrPëPSPë PSPµ PICrC, P°PSPsPIP»PµPSPëPNë. P PµP·CrP»CHC, P°C, Pë
PrPsCrP»C-PrP¶PµPSCH in vitro PIPeP°P·CrCTjC, CH PSP° C, Pµ, C%oPs PrPs
PjPµC, P°P±PsP»C-P·PjCr C, P°PjCrCrP»PsP·PëPSCr P·P°P»CrC†P°CTjC, CHCrCj
C,, PµCTjPjPµPSC, Pë CYP3A4 C- CYP2D6, P° C, P°PePsP¶ PSPµP·PSP°C†PSPsCTj C” CrC†P°CrC, CH
C-PSC€PëC... C-P·PsC,, PµCTjPjPµPSC, C-PI CYP.

PµCTjPëPiPSC-C†PµPSPSCj P°PeC, PëPIPSPsCrC, C- C,, PµCTjPjPµPSC, C-PI, CjPeC-
P±PµCTjCrC, CH CrC†P°CrC, CH Cr PiPµC†C-PSPePsPIPsPjCr PjPµC, P°P±PsP»C-P·PjC-, PjPsP¶Pµ
PiCTjPëP·PIPµCrC, Pë PrPs PiPsCrPëP»PµPSPsC— PrC-C— C, P°PjCrCrP»PsP·PëPSCr. PµPµCTjPµPr
PIPëPiPµPrPµPSPSCjPj P· CrPµC†PµCTj PjPµC, P°P±PsP»C-C, Pë C, P°PjCrCrP»PsP·PëPSCr
PiC-PrCTjPsC...P»PsCTjPëPrCr PiC-PrPrP°CTjC, CHCrCj C€PëCTjPsPePsPjCr
P·PiBj™ CjP·CrPIP°PSPSCr P· PiP»CHPeCrCTjPsPSC-PrPsPj P°P±Ps CrCrP»CHC,, P°C, PsPj.

P’PëPIPµPrPµPSPSCj

P”CrC, P°CrC, PµCTjPëPr

P’PëPIPµPrPµPSPSCj PrCrC, P°CrC, PµCTjPëPrCr P·P°P»PµP¶PëC, CH PIC-Pr PrPsP·Pë, C-

C†PμPNε PιCΉPsC†PμCΓ CΓP»C-Pr PsPιPεCΓCρPIP°C,Pε C΀Pε C,P°PePεPNε, C%Ps
PIC-PrP±CρPIP°C”C,CHCΓC΀ PrPIPsPjP° PιP°CΉP°P»PμP»CHΉPSPεPjPε CεP»C΀C...P°PjPε:
PsPrPεPS C” PSP°CΓPεC†CρPIP°PSPεPj PιCΉPε PeP»C-PSC-C†PSPs P·PSP°C†CρC%PεC...
PePsPSC†PμPSC,CΉP°C†C-C΀C..., P° PrCΉCρPiPεPNε BΉ” PSPμPSP°CΓPεC†CρPIP°PSPεPj. PμCΉPε
PSPεP·CHΉPePεC... PePsPSC†PμPSC,CΉP°C†C-C΀C... Cρ CΓPεCΉPsPIP°C,C†C- PeCΉPsPIC-
(PjPμPSCεPμ 3 PSPI/PjP») PrCρC,P°CΓC,PμCΉPεPr CεPIPεPrPePs PIPεPIPsPrPεC,CHCΓC΀ C΀Pε
P·P°P»PμP¶PSPεPj PIC-Pr PePsPSC†PμPSC,CΉP°C†C-C—, C,P°Pe C- PSPμP·P°P»PμP¶PSPεPj
PIC-Pr PePsPSC†PμPSC,CΉP°C†C-C— CεP»C΀C...PsPj. PμCΉPε P·P°CΓC,PsCΓCρPIP°PSPSC-
PsPrPSPsCΉP°P·PsPIPεC... PrPsP· 5 PjPi P°P±Ps PjPμPSCεPεC... PrPsP· P±CρP»Ps
PIPεC΀PIP»PμPSPs PsP·PSP°PePε CεPIPεPrPePsPiPs PeP»C-CΉPμPSCΓCρ C-
PICΓC,P°PSPsPIP»PμPSPs PιPμCΉC-PsPr PSP°PιC-PIPIPεPIPμPrPμPSPSC΀, C΀PεPεPNε
C,CΉPεPIP°C” PIC-Pr 3 PrPs 9 PrPSC-PI.

PμCΉPε C,PμCΉP°PιPμPIC,PεC†PSPεC... PePsPSC†PμPSC,CΉP°C†C-C΀C... PιC-CΓP»C΀
PιPsPIC,PsCΉPSPsPiPs PIPIPμPrPμPSPSC΀ PrPsP·Pε 0,5 PjPi/PrPsP±Cρ PrPsPjC-PSCΓC”
PιPsPIC-P»CHΉPSC-CεPεPNε, P»C-PSC-PNεPSPεPNε CεP»C΀C... PIPεPIPμPrPμPSPSC΀, P°
PιPμCΉC-PsPr PSP°PιC-PIPIPεPIPμPrPμPSPSC΀ CΓC,P°PSPsPIPεC,CH PιCΉPεP±P»PεP·PSPs
3BΉ”5 C,PεP¶PSC-PI.

PϋP°PjCΓCρP»PsP·PεPS

PϋP°PjCΓCρP»PsP·PεPS C- PNεPsPiPs PjPμC,P°P±PsP»C-C,Pε PIPεPIPsPrC΀C,CHCΓC΀
PιPμCΉPμPIP°P¶PSPs P· CΓPμC†PμCΉ, PI C΀PεC-PNε PιCΉPεP±P»PεP·PSPs 9 % PrPsP·Pε
PιCΉPεCΓCρC,PSC- Cρ PIPεPiP»C΀PrC- PSPμP·PjC-PSPμPSPsC— P°PeC,PεPIPSsC—
CΉPμC†PsPIPεPSPε.

PμC-CΓP»C΀ PIPSCρC,CΉC-CεPSCΉPsPIPμPSPSPsPiPs P°P±Ps PιPμCΉPsCΉP°P»CHΉPSPsPiPs
PIPIPμPrPμPSPSC΀ PI P»C-PeP°CΉCΓCHΉPeC-PNε C„PsCΉPjC- P· PSPμPiP°PNεPSPεPj
PIPεPIC-P»CHΉPSPμPSPSC΀Pj PιPμCΉC-PsPr PSP°PιC-PIPIPεPIPμPrPμPSPSC΀
C,P°PjCΓCρP»PsP·PεPSCρ, C%Ps PjC-CΓC,PεC,CHCΓC΀ Cρ PιP»P°P·PjC- PeCΉPsPIC-,
PePsP»PεPIP°C”C,CHCΓC΀ PI PrC-P°PιP°P·PsPSC- PIC-Pr 5 PrPs 7 PiPsPrPεPS. P\$PμCΉPμP·
C„P°CΉPjP°PePsPeC-PSPμC,PεPeCρ, CΉPμPiCρP»CHΉPsPIP°PSCρ CεPIPεPrPeC-CΓC,CH
PιPsPiP»PεPSP°PSPSC΀, Cρ PIPεPιP°PrPeCρ P· C,P°PjCΓCρP»PsP·PεPSPsPj Cρ PeP°PιCΓCρP»P°C...
P· PjPsPrPεC„C-PePsPIP°PSPεPj PIPεPIC-P»CHΉPSPμPSPSC΀Pj, CΓPιCΉP°PIP¶PSC-PNε
PιPμCΉC-PsPr PSP°PιC-PIPIPεPIPμPrPμPSPSC΀ C,P°PjCΓCρP»PsP·PεPSCρ,
PιCΉPεPNεPSC΀C,PsPiPs PιC-CΓP»C΀ C—PrPε, CΓC,P°PSPsPIPεC,CH PιCΉPεP±P»PεP·PSPs 10
PiPsPrPεPS, P° PI CΉC-PIPSsPIP°P¶PSC-PNε PePsPSC†PμPSC,CΉP°C†C-C— Cρ
PιP°C†C-C”PSC,C-PI BΉ” PιCΉPεP±P»PεP·PSPs 13 PiPsPrPεPS.

PμP°C†C-C”PSC,Pε P»C-C,PSCHΉPsPiPs PIC-PeCé

P”CρC,P°CΓC,PμCΉPεPr

PμP°CΉPjP°PePsPeC-PSPμC,PεPeP° PrCρC,P°CΓC,PμCΉPεPrCρ PsC†C-PSCΉPIP°P»P°CΓC΀ Cρ 36
P·PrPsCΉPsPIPεC... C†PsP»PsPIC-PeC-PI Cρ PIC-C†C- PIC-Pr 24 PrPs 87 CΉPsPeC-PI PιC-CΓP»C΀
PIPIPμPrPμPSPSC΀ PsPrPSPsCΉP°P·PsPIPsC— PrPsP·Pε 5 PjPi. P—PSP°C†PSPsC—
P·P°P»PμP¶PSPsCΓC,C- PIPιP»PεPICρ PrCρC,P°CΓC,PμCΉPεPrCρ PIC-Pr PIC-PeCρ
PιP°C†C-C”PSC,P° PSPμ CΓPιPsCΓC,PμCΉC-PιP°P»PsCΓC΀, P°P»Pμ PιPμCΉC-PsPr
PSP°PιC-PIPIPεPIPμPrPμPSPSC΀ P±CρPI PePsCΉPsC,CεPεPj Cρ C†PsP»PsPIC-PeC-PI PIC-PePsPj
PrPs 50 CΉPsPeC-PI. PϋC,P°C,PεCΓC,PεC†PSPεC... PIC-PrPjC-PSPSPsCΓC,PμPNε Cρ
PιPμCΉC-PsPrC- PSP°PιC-PIPIPεPIPμPrPμPSPSC΀ PSPμ P±CρP»Ps PιCΉPε
PιPsCΉC-PIPSsPIPSsC- PιCΉCρPιPε 50BΉ”69-CΉC-C†PSPεC... PrPsCΓP»C-PrP¶CρPIP°PSPεC... P·

PiC₇Cr₇PiPsC₇ PrPsC₇P»C-PrP₇Cr₇PIP°PSPëC... PIC-PePsPj PiPsPSP°Pr 70 C₇PsPeC-PI.

PÿP°PjC₇Cr₇P»PsP·PëPS

P₇P₇C₇P₇C...C₇P₇C₇PSP₇ PiPsC₇C-PIPSC₇IP»CH₇PSP₇ PrPsC₇P»C-PrP₇P₇PSPSC₇ P·P°PiP°P»CH₇PSPsPiPs PIP°P»PëPIC₇ C,P°PjC₇Cr₇P»PsP·PëPSC₇ PiC-PrC₇PsC...P»PsC₇PëPrC₇ (AUC (PiP»PsC%₀C- PiC-Pr C,,P°C₇PjP°PePsPeC-PSP₇C,PëC₇PSPsC₇ PeC₇PëPIPsC₇)) C- P₇P₇C₇C-PsPrC₇ PSP°PiC-PIPIPëPIP₇PrP₇PSPSC₇ PIPeP°P·C₇C" PSP° C,P₇, C%₀Ps C,,P°C₇PjP°PePsPeC-PSP₇C,PëC₇PSP° PrC-C₇ C,P°PjC₇Cr₇P»PsP·PëPSC₇ PiC-PrC₇PsC...P»PsC₇PëPrC₇ PjPsP₇P₇ P±C₇C,Pë C,C₇PsC...Pë C,C₇PëPIP°P»C-CëPsC₇ Cr PIP°C₇C-C"PSC,C-PI P»C-C,PSC₇PsPiPs PIC-PeC₇ PiPsC₇C-PIPSC₇IPSPs P· PjPsP»PsPrPëPjPë P·PrPsC₇PsPIPëPjPë PrPsP±C₇PsPIPsP»CH₇C₇IPjPë C₇PsP»PsPIC-C₇PsC— C₇C,P°C,C-. P'P»P°C₇PSPëP₇ PeP»C-C₇P₇PSC₇ PSP₇ P·P°P»P₇P₇PëC,CH₇ PIC-Pr P·PIB₇TM C₇IP·C₇PIP°PSPSC₇ C,P°PjC₇Cr₇P»PsP·PëPSC₇ PiC-PrC₇PsC...P»PsC₇PëPrC₇ P· P°P»CH₇C,,P°-1-PePëC₇P»PëPj PiP»C-PePsPiC₇PsC,P₇C—PSPsPj, P°P»P₇ P·PSPëP₇Cr₇C"CH₇C₇ C₇ P· PIC-PePsPj PiP°C₇C-C"PSC,P°, Cr C₇P₇P·C₇P»CH₇C,P°C,C- C₇PsPiPs P·P°PiP°P»CH₇PSP° PrC-C₇ C" PSP° 40 % C₇PëP»CH₇PSC-CëPsC₇ (AUC) Cr PiP°C₇C-C"PSC,C-PI PIC-PePsPj PIC-Pr 55 PrPs 75 C₇PsPeC-PIB PiPsC₇C-PIPSC₇IPSPs P· PrC-C"CH₇ Cr PiP°C₇C-C"PSC,C-PI PIC-PePsPjB 20B₇"32 C₇PsPePë.

P₇PëC₇PePsPIP° PSP₇PrPsC₇C,P°C,PSC-C₇C,CH

P"Cr₇C,P°C₇C,P₇C₇PëPr

P'PiP»PëPI PSPëC₇PePsPIPsC— PSP₇PrPsC₇C,P°C,PSPsC₇C,C- PSP° C,,P°C₇PjP°PePsPeC-PSP₇C,PëPeC₇ PrC₇C,P°C₇C,P₇C₇PëPrC₇ PSP₇ PIPëPIC₇P°P»Pë. P₇P»P₇ Cr C₇P₇C₇C- P»CH₇PrPëPSPë PIPëC₇IP»C₇C"CH₇C₇ PjP₇PSCëP₇ 0,1 % PrPsP·Pë PrC₇C,P°C₇C,P₇C₇PëPrC₇ 0,5 PjPi Cr C₇C-PIPSPsPIP°P₇PSC-PN₇ PePsPSC₇P₇PSC,C₇P°C₇C-C—, C,PsPjCr PeP»C-PSC-C₇PSPs P·PSP°C₇C₇C%₀PsPiPs PiC-PrPIPëC%₀P₇PSPSC₇ PePsPSC₇P₇PSC,C₇P°C₇C-C— PrC₇C,P°C₇C,P₇C₇PëPrC₇ Cr PiP»P°P·PjC- PeC₇PsPIC- Cr PiP°C₇C-C"PSC,C-PI P· PSPëC₇PePsPIPsC₇ PSP₇PrPsC₇C,P°C,PSC-C₇C,CH₇ PsC₇C-PeC₇PIP°C,Pë PSP₇ C₇P»C-PrB (PrPëPI. C₇PsP·PrC-P» B«PÿPiPsC₇C-P± P·P°C₇C,PsC₇CrPIP°PSPSC₇ C,P° PrPsP·PëB»).

PÿP°PjC₇Cr₇P»PsP·PëPS

P₇P°C₇PjP°PePsPeC-PSP₇C,PëPeP° C,P°PjC₇Cr₇P»PsP·PëPSC₇ PiC-PrC₇PsC...P»PsC₇PëPrC₇ PiPsC₇C-PIPSC₇IPIP°P»P°C₇C₇ Cr 6 PiP°C₇C-C"PSC,C-PI P· PSPëC₇PePsPIPsC₇ PSP₇PrPsC₇C,P°C,PSC-C₇C,CH₇ PIC-Pr C₇P»P°P±PePsPiPs PrPs PiPsPjC-C₇PSPsPiPs (30 ≤ CL_{cr} < 70 PjP»/C...PI/1,73 Pj²) P°P±Ps PIC-Pr PiPsPjC-C₇PSPsPiPs PrPs C,C₇IP₇PePsPiPs (10 ≤ CL_{cr} < 30 PjP»/C...PI/1,73 Pj²) C₇C,CrPiP₇PSC₇ C- Cr 6 PrPsC₇P»C-PrP₇Cr₇PIP°PSPëC... P· PSPsC₇PjP°P»CH₇PSPëPj PeP»C-C₇P₇PSC₇P₇Pj (CL_{cr} < 90 PjP»/C...PI/1,73 Pj²). Pj C,PsPN₇ C₇P°C₇ C₇IPe Cr P·P°PiP°P»CH₇PSC-PN₇ PePsPSC₇P₇PSC,C₇P°C₇C-C— C,P°PjC₇Cr₇P»PsP·PëPSC₇ PiC-PrC₇PsC...P»PsC₇PëPrC₇ Cr PiP»P°P·PjC- PeC₇PsPIC- C₇PiPsC₇C,P₇C₇C-PiP°P»P°C₇C₇ P·PjC-PSP° PI C₇P₇P·C₇P»CH₇C,P°C,C- P·PjC-PSPSPsPiPs P·PIB₇TM C₇IP·C₇PIP°PSPSC₇ P· P°P»CH₇C,,P°-1-PePëC₇P»PëPj PiP»C-PePsPiC₇PsC,P₇C—PSPsPj, PePsPSC₇P₇PSC,C₇P°C₇C-C₇ PSP₇P·PIB₇TM C₇IP·P°PSPsPiPs (P°PeC,PëPIPsPiPs) C,P°PjC₇Cr₇P»PsP·PëPSC₇ PiC-PrC₇PsC...P»PsC₇PëPrC₇, P° C,P°PePsP₇ PIP»P°C₇PSPëP₇ PeP»C-C₇P₇PSC₇ P·P°P»PëCëP°P»PëC₇C₇ PIC-PrPSPsC₇PSPs C₇C,P°P±C-P»CH₇PSPëPjPë. PÿPsPjCr PiP°C₇C-C"PSC,P°Pj P· PSPëC₇PePsPIPsC₇ PSP₇PrPsC₇C,P°C,PSC-C₇C,CH₇ PSP₇ PiPsC,C₇C-P±PSP° PePsC₇P₇PeC₇C-C₇ PrPsP·Pë C,P°PjC₇Cr₇P»PsP·PëPSC₇ PiC-PrC₇PsC...P»PsC₇PëPrC₇ PI PeP°PiC₇Cr₇P»P°C.... P₇P»P₇ PiP°C₇C-C"PSC,Pë P·

C, P_μ C₇ P_j C - P S P ° P » C H P S P S C H C I C, P ° P r C - C " C H P S P e C H P e P s P I P s C —
P S P μ P r P s C I C, P ° C, P S P s C I C, C - (C_{Lcr} < 10 P_j P » / C ... P I / 1, 73 P_j²) P S P μ
P r P s C I P » C - P r P ¶ C r P I P ° P » P e C I C I.

P_μ P_μ C ‡ C - P S P e P s P I P ° P S P μ P r P s C I C, P ° C, P S C - C I C, C H

P " C r C, P ° C I C, P_μ C₇ P e P r

P' P i P » P e P I P i P μ C ‡ C - P S P e P s P I P s C — P S P μ P r P s C I C, P ° C, P S P s C I C, C - P S P °
C „ P ° C₇ P_j P ° P e P s P e C - P S P μ C, P e P e C r P r C r C, P ° C I C, P μ C₇ P e P r C r P S P μ P I P e P I C ‡ P ° P I C I C I (P r P e P I.
C₇ P s P · P r C - P » B « P μ C₇ P s C, P e P i P s P e P ° P · P ° P S P S C I B »). P h C I P e C - P » C H P e P e
P r C r C, P ° C I C, P μ C₇ P e P r P I P e P I P s P r P e C, C H C I C I P i P μ C₇ P μ P I P ° P ¶ P S P s C e P » C I C ... P s P_j
P_j P μ C, P ° P ± P s P » C - P · P_j C r, P s C ‡ C - P e C r C " C, C H C I C I, C % P s C₇ C - P I P S C -
P r C r C, P ° C I C, P μ C₇ P e P r C r P I P i P » P ° P · P_j C - P e C₇ P s P I C - C ‡ P e C ... P i P ° C ‡ C - C " P S C, C - P I
P ± C r P r C r C, C H P i C - P r P I P e C % P μ P S P e P_j P e, P ° P i P μ C₇ C - P s P r P S P ° P i C - P I P I P e P I P μ P r P μ P S P S C I
B H " C, C₇ P e P I P ° P » C - C e P e P_j (P r P e P I. C₇ P s P · P r C - P » P e B « P Ÿ P i P s C I C - P ±
P · P ° C I C, P s C I C r P I P ° P S P S C I C, P ° P r P s P · P e B » C - B « P h C I P s P ± P » P e P I P s C I C, C -
P · P ° C I C, P s C I C r P I P ° P S P S C I B »).

P Ÿ P ° P_j C I C r P » P s P · P e P S

P α P ° C₇ P_j P ° P e P s P e C - P S P μ C, P e P e C r C, P ° P_j C I C r P » P s P · P e P S C r P i C - P r C₇ P s C ... P » P s C₇ P e P r C r
P i P s C₇ C - P I P S C H P I P ° P » P e C r 8 P i P ° C ‡ C - C " P S C, C - P I P · P i P s P_j C - C₇ P S P e P_j
P i P s C₇ C r C e P μ P S P S C I P_j C „ C r P S P e C ‡ C - C — P i P μ C ‡ C - P S P e P e (P e P » P ° C I P e C „ C - P e P ° C ‡ C - C I P · P °
P S P ° P N e P » P r P s P_j B H " P μ B H TM C H: C I C, C r P i P μ P S C - A C - B) C - C r 8 P r P s C I P » C - P r P ¶ C r P I P ° P S P e C ...
P · P S P s C₇ P_j P ° P » C H P S P s C H C „ C r P S P e C ‡ C - C " C H P i P μ C ‡ C - P S P e P e. P' C, P s P N e C ‡ P ° C I C I P e
P · P_j C - P S P ° P I P · P ° P i P ° P » C H P S C - P N e P e P s P S C ‡ P μ P S C, C₇ P ° C ‡ C - C — C, P ° P_j C I C r P » P s P · P e P S C r
P i C - P r C₇ P s C ... P » P s C₇ P e P r C r P I P i P » P ° P · P_j C - C I P i P s C I C, P μ C₇ C - P i P ° P » P ° C I C I P I
C₇ P μ P · C r P » C H C, P ° C, C - P · P_j C - P S P S P s P i P s P · P i B H TM C I P · C r P I P ° P S P S C I P · P ° P » C H C „ P ° - 1 -
P e P e C I P » P e P_j P i P » C - P e P s P i C₇ P s C, P μ C — P S P s P_j, P e P s P S C ‡ P μ P S C, C₇ P ° C ‡ C - C I
P S P μ P · P i B H TM C I P · P ° P S P s P i P s (P ° P e C, P e P I P S P s P i P s) C, P ° P_j C I C r P » P s P · P e P S C r
P i C - P r C₇ P s C ... P » P s C₇ P e P r C r P S P μ P · P ° P · P S P ° P I P ° P » P ° P · P S P ° C ‡ P S P e C ... P · P_j C - P S,
P I C - P r P · P S P ° C ‡ P ° P » P ° C I C I C, C - P » C H P e P e P i P s P_j C - C₇ P S P ° (32 %) P · P_j C - P S P °
P I P » P ° C I P S P s P i P s P e P » C - C₇ P μ P S C I C r P S P μ P · P i B H TM C I P · P ° P S P s P i P s C, P ° P_j C I C r P » P s P · P e P S C r
P i C - P r C₇ P s C ... P » P s C₇ P e P r C r. P Ÿ P s P_j C r P i P ° C ‡ C - C " P S C, P e P · P i P s P_j C - C₇ P S P e P_j
P i P s C₇ C r C e P μ P S P S C I P_j C „ C r P S P e C ‡ C - C — P i P μ C ‡ C - P S P e P e P S P μ P i P s C, C₇ P μ P ± C r C H C, C H
P e P s C₇ P μ P e C ‡ C - C — P r P s P · P e C, P ° P_j C I C r P » P s P · P e P S C r P i C - P r C₇ P s C ... P » P s C₇ P e P r C r. P " C - C I
C, P ° P_j C I C r P » P s P · P e P S C r P i C - P r C₇ P s C ... P » P s C₇ P e P r C r P S P μ P I P e P I C ‡ P ° P » P ° C I C I C r
P i P ° C ‡ C - C " P S C, C - P I P · C, C I P ¶ P e P e P_j P i P s C₇ C r C e P μ P S P S C I P_j C „ C r P S P e C ‡ C - C —
P i P μ C ‡ C - P S P e P e.

P' P μ P · P i P μ P e P ° C, P ° P e P » C - P S C - C ‡ P S C - P r P s C I P » C - P r P ¶ P μ P S P S C I.

P Ÿ P μ C₇ C ‡ P μ P I P ° P S P μ P r P s C I C, P ° C, P S C - C I C, C H

P_j 4 - C₇ C - C ‡ P S P s P_j C r P e P » C - P S C - C ‡ P S P s P_j C r P r P s C I P » C - P r P ¶ P μ P S P S C -
P · P ° C I C, P s C I C r P I P ° P S P S C I P r C r C, P ° C I C, P μ C₇ P e P r C r C r P i P s C " P r P S P ° P S P S C - P ·
C, P ° P_j C I C r P » P s P · P e P S P s P_j P r P » C I P P » C - P e C r P I P ° P S P S C I P r P s P ± C₇ P s C I P e C - C I P S P s C —
P i C - P i P μ C₇ P i P » P ° P · C - C — P i P μ C₇ P μ P r P_j C - C ... C r C₇ P s P I P s C — P · P ° P » P s P · P e C r 4844
C ‡ P s P » P s P I C - P e C - P I (P r P s C I P » C - P r P ¶ P μ P S P S C I C o m b A T) C ‡ P ° C I C, P s C, P °
P I P e P S P e P e P S P μ P S P S C I C I P μ C₇ C ‡ P μ P I P s C — P S P μ P r P s C I C, P ° C, P S P s C I C, C - (P · P ± C - C₇ P S P μ
P i P s P S C I C, C, C I) P I P i C₇ C r P i C - P e P s P_j P ± C - P S P s P I P ° P S P s C — C, P μ C₇ P ° P i C - C — P ± C r P » P °

PIPëC%PsCĤ (14/1610, 0,9 %), PSC-P¶ Cr P±CřPrCĤ-CĹPeC-PNë PiCĤCrPïC-
PjPsPSPsC,PµCĤP°PïC-C— PrCřC,P°CĹC,PµCĤPëPrPsPj (4/1623, B B 0,2 %) P°P±Ps
C,P°PjCĹCřP»PsP·PëPSPsPj (10/1611, 0,6 %).

P' PsPeCĤPµPjPsPjCr 4-CĤC-C±PSPsPjCr PeP»C-PSC-C±PSPsPjCr PïPsCĤC-PIPSCLP»CHPSPsPjCr
PrPsCĹP»C-PrP¶PµPSPSC- PïP»P°C†PµP±Ps P· C...C-PjC-C±PSPsCĤ
PïCĤPsC,,C-P»P°PeC,PëPePsCĤ PrCřC,P°CĹC,PµCĤPëPrPsPj P·P° CřC±P°CĹC,CĤ 8231
C±PsP»PsPIC-PeP° PIC-PePsPj PIC-Pr 50 PrPs 75 CĤPsPeC-PI P· PïPsPïPµCĤPµPrPSC-PjPë
PSPµPiP°C,PëPIPSPëPjPë PrP°PSPëPjPë P±C-PsPïCĹC-C— PSP° CĤP°Pe
PïPµCĤPµPrPjC-C...CřCĤPsPIPsC— P·P°P»PsP·Pë C- PIPëC...C-PrPSPëPj CĤC-PIPSPµPj PSA
PjC-P¶ 2,5 PSPI/PjP» C- 10,0 PSPI/PjP» Cř C±PsP»PsPIC-PeC-PI PIC-Pr 50 PrPs 60 CĤPsPeC-PI
P°P±Ps 3 PSPI/PjP» C- 10,0 PSPI/PjP» Cř C±PsP»PsPIC-PeC-PI PIC-PePsPj PïPsPSP°Pr 60
CĤPsPeC-PI (PrPsCĹP»C-PrP¶PµPSPSCĹ REDUCE) P±CřP»Ps PICĹC,P°PSPsPIP»PµPSPs, C%Ps
C±P°CĹC,PsC,P° PIPëPSPëPePSPµPSPSCĹ CĹPµCĤC†PµPIPsC— PSPµPrPsCĹC,P°C,PSPsCĹC,C- Cř
PïP°C†C-C"PSC,C-PI, CĹPeC- PïCĤPëPNëPjP°P»Pë PrCřC,P°CĹC,PµCĤPëPr 0,5 PjPi 1 CĤP°P· PSP°
PrPµPSCĤ (30/4105, 0,7 %), P±CřP»P° PIPëC%PsCĤ PïPsCĤC-PIPSCLPSPs P·
PïP°C†C-C"PSC,P°PjPë, CĹPeC- PïCĤPëPNëPjP°P»Pë PïP»P°C†PµP±Ps B (16/4126, 0,4 %).
P PµC,CĤPsCĹPïPµPeC,PëPIPSPëPNë P°PSP°P»C-P· C†CHPsPiPs PrPsCĹP»C-PrP¶PµPSPSCĹ
PïPsPeP°P·P°PI PIPëC%Cr C±P°CĹC,PsC,Cř CĹPµCĤC†PµPIPsC— PSPµPrPsCĹC,P°C,PSPsCĹC,C-
Cř PïP°C†C-C"PSC,C-PI, CĹPeC- PïCĤPëPNëPjP°P»Pë PrCřC,P°CĹC,PµCĤPëPr C- P°P»CHC,,P°-
P±P»PsPeP°C,PsCĤ PsPrPSPsC±P°CĹPSPs (12/1152, 1,0 %), PïPsCĤC-PIPSCLPSPs P·
CĹCřP±'C"PeC,P°PjPë, CĹPeC- PïCĤPëPNëPjP°P»Pë PrCřC,P°CĹC,PµCĤPëPr P±PµP· P°P»CHC,,P°-
P±P»PsPeP°C,PsCĤP° (18/2953, 0,6 %), PïP»P°C†PµP±Ps C- P°P»CHC,,P°-P±P»PsPeP°C,PsCĤ
(1/1399, <0,1 %) P°P±Ps PïP»P°C†PµP±Ps P±PµP· P°P»CHC,,P°-P±P»PsPeP°C,PsCĤP° (15/2727,
0,6 %). PµCĤPëC±PëPSPSPsPiPs PIP·P°C"PjPsP·PIBĤ™ CĹP·PeCř PjC-P¶
P·P°CĹC,PsCĹCřPIP°PSPSCĹPj PrCřC,P°CĹC,PµCĤPëPrCř (CĹP°PjPsCĹC,C-PNëPSPs P°P±Ps Cř
PePsPjP±C-PSP°C†C-C— P· P°P»CHC,,P°-P±P»PsPeP°C,PsCĤP°PjPë) C,P°
PIPëPSPëPePSPµPSPSCĹPj CĹPµCĤC†PµPIPsC— PSPµPrPsCĹC,P°C,PSPsCĹC,C-
PICĹC,P°PSPsPIP»PµPSPs PSPµ P±CřP»Ps (PrPëPI. CĤPsP·PrC-P» B«PhCĹPsP±P»PëPIPsCĹC,C-
P·P°CĹC,PsCĹCřPIP°PSPSCĹB»).

P P°Pe PïPµCĤPµPrPjC-C...CřCĤPsPIPsC— P·P°P»PsP·Pë C-
PSPëP·CHPePsPrPëC,,PµCĤPµPSC†C-PNëPsPIP°PSC- PïCřC...P»PëPSPë

PJ 4-CĤC-C±PSPsPjCr PrPsCĹP»C-PrP¶PµPSPSC-, PI CĹPePsPjCr PIPëPIC±P°P»Pë PrC-CĤ
PrCřC,P°CĹC,PµCĤPëPrCř PïPsCĤC-PIPSCLPSPs P· PïP»P°C†PµP±Ps, Cř 8231 C±PsP»PsPIC-PeP°
PIC-PePsPj PIC-Pr 50 PrPs 75 CĤPsPeC-PI P· PïPsPïPµCĤPµPrPSC-PjPë PSPµPiP°C,PëPIPSPëPjPë
PrP°PSPëPjPë P±C-PsPïCĹC-C— PSP° CĤP°Pe PïPµCĤPµPrPjC-C...CřCĤPsPIPsC— P·P°P»PsP·Pë
C- PIPëC...C-PrPSPëPj CĤC-PIPSPµPj PSA PjC-P¶ 2,5 PSPI/PjP» C- 10,0 PSPI/PjP» Cř
C±PsP»PsPIC-PeC-PI PIC-PePsPj PIC-Pr 50 PrPs 60 CĤPsPeC-PI P°P±Ps 3 PSPI/PjP» C- 10,0
PSPI/PjP» Cř C±PsP»PsPIC-PeC-PI PIC-PePsPj PIC-Pr 60 CĤPsPeC-PI (PrPsCĹP»C-PrP¶PµPSPSCĹ
REDUCE) 6706 CĹCřP±BĤ™C"PeC,P°Pj P±CřP»Ps PïCĤPsPIPµPrPµPSPs PïPsP»PePsPICř
P±C-PsPïCĹC-CĤ PïCĤPsCĹC,P°C,Pë (PsP±PsPIBĤ™ CĹP·PePsPICř P·P° PïPµCĤPIPëPSPSPëPj
PïCĤPsC,PsPePsP»PsPj), PrP°PSC- CĹPePsC— P±CřP»Pë PIPëPePsCĤPëCĹC,P°PSC- PrP»CĹ
P°PSP°P»C-P·Cř PrPëC,,PµCĤPµPSC†C-CĤPIP°PSPSCĹ P·P°B CëPeP°P»PsCĤ P"P»C-CĹPsPSP°. P'
PrPsCĹP»C-PrP¶PµPSPSC- P±CřP»Ps PIPëCĹPIP»PµPSPs 1517 PïP°C†C-C"PSC,C-PI P·
PrC-P°PiPSPsP·PsPj CĤP°Pe PïCĤPsCĹC,P°C,Pë. P'C-P»CHCëC-CĹC,CH PïCřC...P»PëPS
PïCĤPsCĹC,P°C,Pë (70 %), PIPëCĹPIP»PµPSPëC... P·P° PrPsPïPsPjPsPiPsCĤ P±C-PsPïCĹC-C—, PI
PsP±PsC... PïCĤCrPïP°C... P»C-PeCřPIP°PSPSCĹ PjP°P»Pë PIPëCĹPsPePëPNë CĤC-PIPµPSCĤ
PrPëC,,PµCĤPµPSC†C-CĤPIP°PSPSCĹ (5BĤ"6 P±P°P»C-PI P·P° CëPeP°P»PsCĤ P"P»C-CĹPsPSP°).

PJ PïCĤCrPïC- PrCřC,P°CĹC,PµCĤPëPrCř P·P°CĤPµC"CĹC,CĤPsPIP°PSPs P±C-P»CHCë

PIPëCÍPSPeCí C±P°CÍC,PsC,Cí (n = 29, 0,9 %)

PSPëP·CHPePsPrPëC,,PµCĤPµPSC†C-PMëPsPIP°PSPSPiPs CĤP°PeCí PiCĤPsCÍC,P°C,Pë (8bĤ“10 P±P°P»C-PI P·P° CëPeP°P»PsCĤ P“P»C-CÍPSPSP°) PiPsCĤC-PIPSCĤPSPs P· PiCĤCíPiPsCĤ PiP»P°C†PµP±Ps (n = 19, 0,6 %) (CĤ = 0,15). PJ PiPµCĤCëC- 2 CĤPsPePë PrPsCÍP»C-PrP¶PµPSPSCĤ PeC-P»CHPeC-CÍC,CH PiP°C†C-C”PSC,C-PI P· CĤP°PePsPj PiPµCĤPµPrPjC-C...CíCĤPsPIPsC— P·P°P»PsP·Pë P· PiPsPeP°P·PSPëPePsPj 8bĤ“10 P±P°P»C-PI P·P° CëPeP°P»PsCĤ P“P»C-CÍPSPSP° P±CíP»P° PsPrPSP°PePsPIPsCĤ PI PiCĤCíPiC- PrCíC,P°CÍC,PµCĤPëPrCí (n 17, 0,5%) C- PI PiCĤCíPiC- PiP»P°C†PµP±Ps (n = 18, 0,5 %). PµCĤPsC,CĤPiPsPj B 3bĤ“4-PiPs CĤPsPeC-PI PrPsCÍP»C-PrP¶PµPSPSCĤ P±C-P»CHCëP° PeC-P»CHPeC-CÍC,CH PIPëPiP°PrPeC-PI CĤP°PeCí PiPµCĤPµPrPjC-C...CíCĤPsPIPsC— P·P°P»PsP·Pë P· PiPsPeP°P·PSPëPePsPj 8bĤ“10 P±P°P»C-PI P·P° CëPeP°P»PsCĤ P“P»C-CÍPSPSP° P±CíP»P° PrC-P°PiPSPsCÍC,PsPIP°PSP° Cí PiCĤCíPiC- PrCíC,P°CÍC,PµCĤPëPrCí (n = 12, 0,5 %) PiPsCĤC-PIPSCĤPSPs P· PiCĤCíPiPsCĤ PiP»P°C†PµP±Ps (n = 1, < 0,1%) (CĤ = 0,0035). PkPµPjP°C” PrP°PSPëC... PiCĤPs PIPiP»PëPI PSP° CĤPëP·PëPe CĤPsP·PIPëC,PeCí CĤP°PeCí PiCĤPsCÍC,P°C,Pë Cí C±PsP»PsPIC-PeC-PI, CĤPeC- PiCĤPëPNëPjP°CĤC,CH PrCíC,P°CÍC,PµCĤPëPr PiPsPSP°Pr 4 CĤPsPePë. P’C-PrCÍPSPsC,PsPe PiP°C†C-C”PSC,C-PI P· PrC-P°PiPSPsP·PsPj CĤP°PeCí PiPµCĤPµPrPjC-C...CíCĤPsPIPsC— P·P°P»PsP·Pë P· PiPsPeP°P·PSPëPePsPj 8bĤ“10 P±P°P»C-PI P·P° CëPeP°P»PsCĤ P“P»C-CÍPSPSP° P·P±PµCĤC-PiP°PICÍCĤ PiPsCÍC,C-PMëPSPëPj PI CĤC-P·PSC- PiPµCĤC-PsPrPë PrPsCÍP»C-PrP¶PµPSPSCĤ (1bĤ“2-PëPNë CĤPsPePë, 3bĤ“4-PëPNë CĤPsPePë) PI PiCĤCíPiC- PrCíC,P°CÍC,PµCĤPëPrCí (0,5 % Cí PePsP¶PSPëPNë PiPµCĤC-PsPr C±P°CÍCí), Cí C,PsPNë C±P°CÍ CĤPe Cí PiCĤCíPiC- PiP»P°C†PµP±Ps PIC-PrCÍPSPsC,PsPe PiP°C†C-C”PSC,C-PI C-P· PSPëP·CHPePsPrPëC,,PµCĤPµPSC†C-PMëPsPIP°PSPëPj CĤP°PePsPj PiCĤPsCÍC,P°C,Pë (8bĤ“10 P±P°P»C-PI P·P° CëPeP°P»PsCĤ P“P»C-CÍPSPSP°) P±CíPI PSPëP¶C±PëPj Cí 3bĤ“4-C,PëPNë CĤPsPePë, PSC-P¶ PI 1bĤ“2-PëPNë CĤPsPePë (< 0,1 %B C- 0,5 %, PIC-PrPiPsPIC-PrPSPs) (PrPëPI. CĤPsP·PrC-P» B«PhCÍPSP±P»PëPIPsCÍC,C- P·P°CÍC,PsCÍCíPIP°PSPSCĤB»). PkPµ P±CíP»Ps PSC-CĤPePsC— CĤC-P·PSPëC†C- PI C±P°CÍC,PsC,C- PIPëPiP°PrPeC-PI CĤP°PeCí PiPµCĤPµPrPjC-C...CíCĤPsPIPsC— P·P°P»PsP·Pë P· PiPsPeP°P·PSPëPePsPj 7bĤ“10 P±P°P»C-PI P·P° CëPeP°P»PsCĤ P“P»C-CÍPSPSP° (CĤ = 0,81).

PJ 4-CĤC-C±PSPsPjCí PeP»C-PSC-C±PSPsPjCí PrPsCÍP»C-PrP¶PµPSPSC- P»C-PeCíPIP°PSPSCĤ PrPsP±CĤPsCĤPeC-CÍPSPsC— PiC-PiPµCĤPiP»P°P·C-C— PiPµCĤPµPrPjC-C...CíCĤPsPIPsC— P·P°P»PsP·PëB (CombAT), PrPµB PiPµCĤPIPëPSPSPëPj PiCĤPsC,PsPePsP»PsPj PSPµ P±CíP»Ps PiPµCĤPµPrP±P°C±PµPSPs PsP±PsPiBĤ™ CĤP·PePsPICí P±C-PsPiCÍC-CĤ C- PICÍC- PrC-P°PiPSPsP·Pë CĤP°PeCí PiCĤPsCÍC,P°C,Pë P±CíP»Pë PICÍC,P°PSPsPIP»PµPSC- PSP° P±C-PsPiCÍC-C— P·P° PiPsPeP°P·P°PSPSCĤPjPë, C±P°CÍC,PsC,P° PIPëPiP°PrPeC-PI CĤP°PeCí PiPµCĤPµPrPjC-C...CíCĤPsPIPsC— P·P°P»PsP·Pë P· PiPsPeP°P·PSPëPePsPj 8bĤ“10 P±P°P»C-PI P·P° CëPeP°P»PsCĤ P“P»C-CÍPSPSP° P±CíP»P° 0,5 % (n = 8) Cí PiCĤCíPiC- PrCíC,P°CÍC,PµCĤPëPrCí 0,7 % (n=11), Cí PiCĤCíPiC- C,P°PjCÍCíP»PsP·PëPSCí C,P° 0,3% (n=5) Cí PiCĤCíPiC- PePsPjP±C-PSPsPIP°PSPsC— C,PµCĤP°PiC-C—.

P—PiBĤ™ CĤP·PsPe PjC-P¶ P·P°CÍC,PsCÍCíPIP°PSPSCĤPj PrCíC,P°CÍC,PµCĤPëPrCí C,P° PIPëPSPëPePSPµPSPSCĤPj PSPëP·CHPePsPrPëC,,PµCĤPµPSC†C-PMëPsPIP°PSPsPiPs CĤP°PeCí PiPµCĤPµPrPjC-C...CíCĤPsPIPsC— P·P°P»PsP·Pë P·P°P»PëCëP°C”C,CHCÍCĤ PSPµP·BĤ™ CĤCÍPSPiP°PSPëPj.

P P°Pe PiCĤCáPrPSPsC— P·P°P»PsP·Pë Cí C±PsP»PsPIC-PeC-PI

P”PIP° PIPëPiP°PrPsPe-PePsPSC,CĤPsP»CHPsPIP°PSC- PµPiC-PrPµPjC-PsP»PsPiC-C±PSC- PrPsCÍP»C-PrP¶PµPSPSCĤ, PsPrPSPµ PiCĤPsPIPµPrPµPSPs Cí PŸPëPĥ B B B B B B B B B B B B B B B B (n=339 PIPëPiP°PrPeC-PI CĤP°PeCí PiCĤCíPrPSPsC— P·P°P»PsP·Pë C- n=6780 Cí PiCĤCíPiC- PePsPSC,CĤPsP»CĤ), P° C-PSCëPµ Cí P’PµP»PëPeC-PNë

$P'CH_2P\ddot{e}C, P^{\circ}PSC-C$ — B B B B B B B ($n = 398$ $PIP\ddot{e}P_iP^{\circ}PrPeC-PI$ $CH_2P^{\circ}PeCr$ $PiCH_2CrPrPSPsC-$
 $P\cdot P^{\circ}P \gg PsP\cdot P\ddot{e}C$ — $n = 3930$ Cr $PiCH_2CrPiC-$ $PePsPSC, CH_2PsP \gg CH$) Cr $P \pm P^{\circ}P\cdot P^{\circ}C \dots PrP^{\circ}PSP\ddot{e}C \dots$
 $PsC \dots PsCH_2PsPSP\ddot{e}C$ $P\cdot PrPsCH_2PsPiB\ddot{e}C$ CI , PSP_{μ} $PiPsPeP^{\circ}P\cdot P^{\circ}P \gg P\ddot{e}C$ $PSC-CI$ $PePsPiPs$
 $P\cdot P \pm C-P \gg CH_2C\epsilon P_{\mu}PSPSC$ CI $CH_2P\ddot{e}P\cdot P\ddot{e}PeCr$ $CH_2PsP\cdot PIP\ddot{e}C, PeCr$ $CH_2P^{\circ}PeCr$ $PiCH_2CrPrPSPsC-$
 $P\cdot P^{\circ}P \gg PsP\cdot P\ddot{e}C$ Cr $C \ddagger PsP \gg PsPIC-PeC-PI$ $PiCH_2P\ddot{e}C$ $P\cdot P^{\circ}C\ddot{r}C, PsC\ddot{r}CrPIP^{\circ}PSPSC-$
 $C-PSiC-P \pm C-C, PsCH_2C-PI$ $5\alpha-CH_2P_{\mu}PrCrPeC, P^{\circ}P\cdot P\ddot{e}C$. P $P_{\mu}P\cdot CrP \gg CH_2C, P^{\circ}C, P\ddot{e}C$ $PiP_{\mu}CH_2C\epsilon PsPiPs$
 $PrPsC\ddot{r}P \gg C-PrP_{\mu}P_{\mu}PSPSC$ CI PSP_{μ} $PIP\ddot{e}C$ $PIP\ddot{e}P \gg P\ddot{e}C$ $PiPsP\cdot P\ddot{e}C, P\ddot{e}PIPSPsPiPs$
 $PIP\cdot P^{\circ}C''PjPsP\cdot PiB\ddot{e}C$ CI $P\cdot PeCr$ $P\cdot CH_2P^{\circ}PePsPj$ $PiCH_2CrPrPSPsC-$ $P\cdot P^{\circ}P \gg PsP\cdot P\ddot{e}C$
 $(PIC-PrPSPsC\ddot{r}PSP\ddot{e}PN_{\ddot{e}}CH_2P\ddot{e}P\cdot P\ddot{e}PePrP \gg CI$ 3 1 $CH_2PsPeCr$ $P\cdot P^{\circ}C\ddot{r}C, PsC\ddot{r}CrPIP^{\circ}PSPSC$ CI $PrPs$
 $PIC\ddot{r}C, P^{\circ}PSPsPIP \gg P_{\mu}PSPSC$ CI $PrC-P^{\circ}PiPSPsP\cdot Cr$ $CH_2P^{\circ}PeCr$ $PiCH_2CrPrPSPsC-$ $P\cdot P^{\circ}P \gg PsP\cdot P\ddot{e}C$
 $PiPsCH_2C-PIPSC$ CI $PsPs$ $P < 1$ $CH_2PsPeCr$ $P\cdot P^{\circ}C\ddot{r}C, PsC\ddot{r}CrPIP^{\circ}PSPSC$ CI : $0,70: 95 \%$ $P''P \ddagger 0,34, 1,45$).
 Pj $PrCH_2CrPiPsPjCr$ $PrPsC\ddot{r}P \gg C-PrP_{\mu}P_{\mu}PSPSC-$ $PsC \ddagger C-PSp_{\mu}PSP\ddot{e}PN_{\ddot{e}}$ $PIC-PrPSPsC\ddot{r}PSP\ddot{e}PN_{\ddot{e}}$
 $CH_2P\ddot{e}P\cdot P\ddot{e}PeCH_2P^{\circ}PeCr$ $PiCH_2CrPrPSPsC-$ $P\cdot P^{\circ}P \gg PsP\cdot P\ddot{e}C$, $PiPsPiB\ddot{e}C$ CI $P\cdot P^{\circ}PSP\ddot{e}PN_{\ddot{e}}$ $C-P\cdot$
 $P\cdot P^{\circ}C\ddot{r}C, PsC\ddot{r}CrPIP^{\circ}PSPSC$ CI Pj $C-PSiC-P \pm C-C, PsCH_2C-PI$ $5\alpha-CH_2P_{\mu}PrCrPeC, P^{\circ}P\cdot P\ddot{e}C$
 $PiPsCH_2C-PIPSC$ CI $PsPs$ $P\cdot C, P^{\circ}PeP\ddot{e}Pj$ $PiCH_2P\ddot{e}C$ $PIC-PrC\ddot{r}CrC, PSPsC\ddot{r}C, C-$
 $P\cdot P^{\circ}C\ddot{r}C, PsC\ddot{r}CrPIP^{\circ}PSPSC$ CI $C\ddot{r}C, P^{\circ}PSPsPIP\ddot{e}PI$ $1,08: 95 \%$ $P''P \ddagger 0,62, 1,87$).

$P_{\mu}CH_2P\ddot{e}C \ddagger P\ddot{e}PSPSP\ddot{e}PN_{\ddot{e}}$ $PIP\cdot P^{\circ}C''PjPsP\cdot PiB\ddot{e}C$ CI $P\cdot PsPe$ $PjC-P_{\mu}$ $PIP\ddot{e}P_iP^{\circ}PrPeP^{\circ}PjP\ddot{e}CH_2P^{\circ}PeCr$
 $PiCH_2CrPrPSPsC-$ $P\cdot P^{\circ}P \gg PsP\cdot P\ddot{e}C$ Cr $C \ddagger PsP \gg PsPIC-PeC-PI$ C, P° $PrPsPiPiPsC, CH_2P\ddot{e}PIP^{\circ}P \gg P\ddot{e}Pj$
 $P\cdot P^{\circ}C\ddot{r}C, PsC\ddot{r}CrPIP^{\circ}PSPSC$ CI Pj $PrCrC, P^{\circ}C\ddot{r}C, P_{\mu}CH_2P\ddot{e}PrCrPSP_{\mu}$ $PIC\ddot{r}C, P^{\circ}PSPsPIP \gg P_{\mu}PSPs$.

$P_{\mu}P \gg C-PSC-C \ddagger PSC-C \dots P^{\circ}CH_2P^{\circ}PeC, P_{\mu}CH_2P\ddot{e}C\ddot{r}C, P\ddot{e}PeP\ddot{e}C$.

$P_{\mu}PsPeP^{\circ}P\cdot P^{\circ}PSPSC$ CI .

$P \gg C-PeCrPIP^{\circ}PSPSC$ CI $PiPsPjC-CH_2PSP\ddot{e}C \dots C, P^{\circ}C, CI$ $P_{\mu}P_{\mu}PeP\ddot{e}C \dots C\ddot{r}P\ddot{e}PjPiC, PsPjC-PIB$
 $PrPsP \pm CH_2PsCI$ $PeC-C\ddot{r}PSPsC-$ $PiC-PiP_{\mu}CH_2PiP \gg P^{\circ}P\cdot C-C-$ $PiP_{\mu}CH_2P_{\mu}PrPjC-C \dots CrCH_2PsPiPsC-$
 $P\cdot P^{\circ}P \gg PsP\cdot P\ddot{e}C$.

$P-PjP_{\mu}PSC\epsilon P_{\mu}PSPSC$ CI $CH_2P\ddot{e}P\cdot P\ddot{e}PeCr$ $PIP\ddot{e}PSP\ddot{e}PePSP_{\mu}PSPSC$ CI $PiPsC\ddot{r}C, CH_2PsC-$
 $P\cdot P^{\circ}C, CH_2P\ddot{e}PjPeP\ddot{e}C\ddot{r}P_{\mu}C \ddagger C-$ $C, P^{\circ}PSP_{\mu}PsP \pm C \dots C-PrPSPsC\ddot{r}C, C-$ $C \dots C-CH_2CrCH_2PiC-C \ddagger PSPsPiPs$
 $PIC, CH_2CrC \ddagger P^{\circ}PSPSC$ CI Cr $PiP^{\circ}C \ddagger C-C''PSC, C-PI$ $P\cdot$ $PiPsPjC-CH_2PSP\ddot{e}PjP\ddot{e}C, P^{\circ}C, CI$ $P_{\mu}P_{\mu}PeP\ddot{e}PjP\ddot{e}$
 $C\ddot{r}P\ddot{e}PjPiC, PsPjP^{\circ}PjP\ddot{e}PrPsP \pm CH_2PsCI$ $PeC-C\ddot{r}PSPsC-$ $PiC-PiP_{\mu}CH_2PiP \gg P^{\circ}P\cdot C-C-$
 $PiP_{\mu}CH_2P_{\mu}PrPjC-C \dots CrCH_2PsPiPsC-$ $P\cdot P^{\circ}P \gg PsP\cdot P\ddot{e}C$.

$P_{\mu}CH_2PsC, P\ddot{e}PiPsPeP^{\circ}P\cdot P^{\circ}PSPSC$ CI .

$P''CrPsPrP^{\circ}CH_2C, PSP_{\mu}$ $P\cdot P^{\circ}C\ddot{r}C, PsC\ddot{r}PsPICrCH_2C, CH_2$ $PrP \gg CI$ $P \gg C-PeCrPIP^{\circ}PSPSC$ CI
 $P_{\mu}C-PSPsPeC-$ $PrC-C, P_{\mu}PN_{\ddot{e}}$ ($PrP\ddot{e}PI$. $CH_2PsP\cdot PrC-P \gg B \ll P-P^{\circ}C\ddot{r}C, PsC\ddot{r}CrPIP^{\circ}PSPSC$ CI Cr
 $PiP_{\mu}CH_2C-PsPr$ $PIP^{\circ}PiC-C, PSPsC\ddot{r}C, C-$ $P^{\circ}P \pm Ps$ $PiPsPrCrPIP^{\circ}PSPSC$ CI $PiCH_2CrPrPrCH_2B \gg$).

$P''CrPsPrP^{\circ}CH_2C, PiCH_2PsC, P\ddot{e}PiPsPeP^{\circ}P\cdot P^{\circ}PSP\ddot{e}PN_{\ddot{e}}$ $C \dots PIPsCH_2P\ddot{e}Pj$ $P\cdot$
 $PiC-PrPIP\ddot{e}C \%$ $P_{\mu}PSPsCH_2B$ $C \ddagger CrC, P \gg P\ddot{e}PIC-C\ddot{r}C, CH_2$ $PrPs$ $PrCrC, P^{\circ}C\ddot{r}C, P_{\mu}CH_2P\ddot{e}PrCr,$
 $C-PSC\epsilon P\ddot{e}C \dots C-PSiC-P \pm C-C, PsCH_2C-PI$ $5\alpha-CH_2P_{\mu}PrCrPeC, P^{\circ}P\cdot P\ddot{e}C, C, P^{\circ}PjC\ddot{r}CrP \gg PsP\cdot P\ddot{e}PSC\ddot{r}$
 $(PIP\ddot{e}P \gg CH_2C \ddagger P^{\circ}CH_2C \ddagger P\ddot{e}C, P^{\circ}PjC\ddot{r}CrP \gg PsP\cdot P\ddot{e}PSC-PSPrCrPePsPIP^{\circ}PSP\ddot{e}PN_{\ddot{e}}$
 $P^{\circ}PSiC-PsPSP_{\mu}PICCH_2PsC, P\ddot{e}C \ddagger PSP\ddot{e}PN_{\ddot{e}}$ $PSP^{\circ}P \pm CH_2CI$ Pe), $C-PSC\epsilon P\ddot{e}C \dots PePsPjPiPsPSP_{\mu}PSC, C-PI$
 $PiCH_2P_{\mu}PiP^{\circ}CH_2P^{\circ}C, Cr$ $P^{\circ}P \pm Ps$ $PrPs$ $C\ddot{r}PsC-$ $C-$ $P^{\circ}CH_2P^{\circ}C \dots C-C\ddot{r}Cr$.

P"CrPsPrP°CTC, PiCTHPS,PePiPsPeP°P·P°PSPëPN₂ C...PIPsCTHPëPj, CIPeC- PjP°CTHC,CH
PI P°PSP°PjPSPµP·C- PsCTC,PsCÍC,P°C,PëC±PSCr P°CTC,PµCTC-P°P»CHPSCr
PiC-PiPsC,PµPSP·C-CT.

P"CrPsPrP°CTC, PiCTHPS,PePiPsPeP°P·P°PSPëPN₂ C...PIPsCTHPëPj P· C,CIP¶PePsCT
PiPµC±C-PSPePsPIPsCT PSPµPrPsCÍC,P°C,PSC-CÍC,CT.

**P'P·P°C"PjPsPrC-CI P· C-PSCëPëPjPë P»C-PeP°CTCÍCHPePëPjPë P·P°CÍPsP±P°PjPë C,P°
C-PSCëC- PIPëPrPë PIP·P°C"PjPsPrC-PN₂.**

P"PsCÍP»C-PrP¶PµPSPSC¶ PIP·P°C"PjPsPrC-C— PiCTHµPiP°CTHP°C,Cr P"CrPsPrP°CTC, P·
C-PSCëPëPjPë P»C-PeP°CTCÍCHPePëPjPë P·P°CÍPsP±P°PjPë PSPµ PiCTHPSPIPsPrPëP»PëCÍC¶.
P"°P»C- PiPsPrP°PSPs PSP°C¶IPSCr C-PSC,,PsCTHPjP°C±C-CT PiCTHPS PsPeCTHµPjC-
C-PSPiCTHµPrC-C"PSC,Pë.

P"CzC,P°CÍC,PµCTHPëPr

P"°P»C¶ PsP·PSP°PN₂PsPjP»µPSPSC¶ P· C-PSC,,PsCTHPjP°C±C-C"CT PiCTHPS P·PSPëP¶PµPSPSC¶
CTC-PIPS-C PI PiCTHPSCÍC,P°C,-CÍPiPµC±PëC,,C-C±PSPsPiPs P°PSC,PëPiPµPSCr (PSA) PI
CÍPëCTHPSPIP°C,C±C- PeCTHPSIC- PiC-Pr C±P°CÍ P»C-PeCÍPIP°PSPSC¶
PrCrC,P°CÍC,PµCTHPëPrPsPj C,P° P· CTHµPePsPjPµPSPPrP°C±C-C¶PjPë PIC-PrPSPsCÍPSPs
PIPëC¶IP»µPSPSC¶ CTC°PeCr PiPµCTHµPrPjC-C...CrCTHPSPIPsC— P·P°P»PsP·Pë PrPëPI.
CTHPS·PrC-P» B«PhCÍPsP±P»PëPIPsCÍC,C- P·P°CÍC,PsCÍCrPIP°PSPSC¶B».

*P'PiP»PëPI C-PSCëPëC... P»C-PeP°CTCÍCHPePëC... P·P°CÍPsP±C-PI PSP°
C,,P°CTHPjP°PePsPeC-PSPµC,PëPeCz PrCzC,P°CÍC,PµCTHPëPrCz*

*P—P°CÍC,PsCÍCzPIP°PSPSC¶ CTC°P·PsPj P· C-PSPiC-P±C-C,PsCTHP°PjPë CYP3A4 C,P°/P°P±Ps P-
PiP»C-PePsPiCTHPS,µC—PSCz*

P"CrC,P°CÍC,PµCTHPëPr PiPµCTHµPIP°P¶PSPs PIPëPIPsPrPëC,CHCÍC¶ CëP»C¶C...PsPj
PjPµC,P°P±PsP»C-P·PjCr. P"PsCÍP»C-PrP¶PµPSPSC¶ *in vitro* PiPsPeP°P·CrCTHC,CH, C%Ps
PeP°C,P°P»C-P·P°C,PsCTHP°PjPë PjPµC,P°P±PsP»C-P·PjCr C" CYP3A4 C- CYP3A5.
PhC,,C-C±C-PN₂PSC- PrPsCÍP»C-PrP¶PµPSPSC¶ PIP·P°C"PjPsPrC-C— P· P°PeC,PëPIPSPëPjPë
C-PSPiC-P±C-C,PsCTHP°PjPë CYP3A4 PSPµ PiCTHPSPIPsPrPëP»PëCÍC¶. PµCTHPS,µ PI
PrPsCÍP»C-PrP¶PµPSPSC- PiPsPiCrP»C¶C±C-PN₂PSPsC— C,,P°CTHPjP°PePsPeC-PSPµC,PëPePë
PePsPSC±PµPSC,CTHP°C±C-C— PrCrC,P°CÍC,PµCTHPëPrCr PI CÍPëCTHPSPIP°C,C±C- PeCTHPSIC-
P±CrP»Pë PI CÍPµCTHµPrPSCHPsPjCr PI 1,6B"1,8 CTC°P·Pë PIPëC%PëPjPë Cr
PSPµPIPµP»PëPePsC— PeC-P»CHPePsCÍC,C- C,PëC... PiP°C±C-C"PSC,C-PI, CIPeC-
PsPrPSPsC±P°CÍPSPs P»C-PeCÍPIP°P»PëCÍC¶ PIPµCTHP°PiP°PjC-P»PsPj P°P±Ps
PrPëP»C,C-P°P·PµPjPj (PiPsPjC-CTHPS-C-PSPiC-P±C-C,PsCTHPë CYP3A4 C,P°
C-PSPiC-P±C-C,PsCTHPë P-PiP»C-PePsPiCTHPS,µC—PSCr), PSC-P¶ Cr C-PSCëPëC...
PiP°C±C-C"PSC,C-PI.

PµCTHPë PrPsPIPiPsCÍC,CTHPSPePsPIPsPjCr P·P°CÍC,PsCÍCrPIP°PSPSC- PePsPjP±C-PSP°C±C-C—
PrCrC,P°CÍC,PµCTHPëPrCr P· P»C-PeP°CTCÍCHPePëPjPë P·P°CÍPsP±P°PjPë, CIPeC- C"
CÍPëP»CHPSPsPrC-CTC±PëPjPë C-PSPiC-P±C-C,PsCTHP°PjPë C,,PµCTHPjPµPSC,Cr CYP3A4
(PSP°PiCTHPëPeP»P°Pr: CTHPëC,PsPSP°PIC-CTH, C-PSPrPëPSP°PIC-CTH, PSPµC,,P°P·PsPrPsPS,
C-C,CTHP°PePsPSP°P·PsP», PePµC,PsPePsPSP°P·PsP», CIPeC- PIPIPsPrPëP»Pë
PiPµCTHPSCTHP°P»CHPSPs), PePsPSC±PµPSC,CTHP°C±C-C¶ PrCrC,P°CÍC,PµCTHPëPrCr PI

$\text{C}\Gamma\text{P}\ddot{\text{e}}\text{C}\tilde{\text{H}}\text{P}\text{sP}\text{IP}^\circ\text{C}, \text{C}\dagger\text{C}-\text{P}\text{e}\text{C}\tilde{\text{H}}\text{P}\text{sP}\text{IC}-\text{PjP}\text{sP}\P\text{P}\mu\text{P}\ddot{\text{I}}\text{C}-\text{PrP}\text{IP}\ddot{\text{e}}\text{C}\%_{\text{O}}\text{C}\Gamma\text{P}\text{IP}^\circ\text{C}, \text{P}\ddot{\text{e}}\text{C}\Gamma\text{C}\Pi.$
 $\text{P}\mu\text{P}\text{sPrP}^\circ\text{P}\gg\text{C}\tilde{\text{H}}\text{C}\epsilon\text{P}\mu\text{C}-\text{PSPiC}-\text{P}\pm\text{C}\Gamma\text{P}\text{IP}^\circ\text{PSPSC}\Pi\text{ } 5\alpha\text{-C}\tilde{\text{H}}\text{P}\mu\text{PrC}\Gamma\text{P}\text{eC}, \text{P}^\circ\text{P}\cdot\text{P}\ddot{\text{e}}\text{P}\ddot{\text{I}}\text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}$
 $\text{P}\ddot{\text{I}}\text{C}-\text{PrC}\Gamma\text{P}\ddot{\text{e}}\text{P}\gg\text{P}\mu\text{PSC}-\text{PN}\ddot{\text{e}}\text{PrC}-\text{C}-\text{PrC}\Gamma\text{C}, \text{P}^\circ\text{C}\Gamma\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{PrC}\Gamma\text{C}''\text{PjP}^\circ\text{P}\gg\text{PsPN}\ddot{\text{e}}\text{PjP}\text{sP}\text{IC}-\text{C}\tilde{\text{H}}\text{PSP}\ddot{\text{e}}\text{Pj}.$
 $\text{P}\tilde{\text{H}}\text{P}\gg\text{P}\mu\text{PjP}\text{sP}\P\text{P}\gg\text{P}\ddot{\text{e}}\text{P}\text{IP}\mu\text{P}\cdot\text{PjP}\mu\text{PSC}\epsilon\text{P}\mu\text{PSPSC}\Pi\text{C}\ddagger\text{P}^\circ\text{C}\Gamma\text{C}, \text{PsC}, \text{P}\ddot{\text{e}}\text{P}\text{IP}\text{IP}\mu\text{PrP}\mu\text{PSPSC}\Pi\text{PrP}\text{sP}\cdot$
 $\text{PrC}\Gamma\text{C}, \text{P}^\circ\text{C}\Gamma\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{PrC}\Gamma\text{C}\text{C}\Gamma\text{C}\tilde{\text{H}}\text{P}^\circ\text{P}\cdot\text{C}-\text{C}\tilde{\text{H}}\text{P}\text{sP}\cdot\text{P}\text{IP}\ddot{\text{e}}\text{C}, \text{P}\text{eC}\Gamma\text{P}\ddot{\text{I}}\text{PsP}\pm\text{C}-\text{C}\ddagger\text{PSP}\ddot{\text{e}}\text{C}\dots\text{P}\mu\text{C}, \text{P}\mu\text{P}\text{eC}, \text{C}-\text{PI}.$
 $\text{P}\ddot{\text{Y}}\text{P}\gg\text{C}-\text{PrP}\text{IC}-\text{PrP}\cdot\text{PSP}^\circ\text{C}\ddagger\text{P}\ddot{\text{e}}\text{C}, \text{P}\ddot{\text{e}}, \text{C}\%_{\text{O}}\text{PsC}\Gamma\text{C}\tilde{\text{H}}\text{P}^\circ\text{P}\cdot\text{C}-\text{P}\ddot{\text{I}}\text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{P}\text{IPSC}-\text{C}\ddagger\text{P}\mu\text{PSPSC}\Pi$
 $\text{P}^\circ\text{P}\text{eC}, \text{P}\ddot{\text{e}}\text{P}\text{IPSPsC}\Gamma\text{C}, \text{C}-\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{PjP}\mu\text{PSC}, \text{C}\Gamma\text{PrP}\text{sP}\text{IPiP}\ddot{\text{e}}\text{PN}\ddot{\text{e}}\text{P}\ddot{\text{I}}\text{P}\mu\text{C}\tilde{\text{H}}\text{C}-\text{PsPr}$
 $\text{PSP}^\circ\text{P}\ddot{\text{I}}\text{C}-\text{P}\text{IP}\text{IP}\ddot{\text{e}}\text{P}\text{IP}\mu\text{PrP}\mu\text{PSPSC}\Pi\text{PjP}\text{sP}\P\text{P}\mu\text{C}\Gamma\text{C}, \text{P}^\circ\text{C}, \text{P}\ddot{\text{e}}\text{C}\%_{\text{O}}\text{P}\mu\text{PrP}\text{sP}\text{IC}\epsilon\text{P}\ddot{\text{e}}\text{PjC}-\text{C}\Gamma\text{C}\Gamma\text{P}\ddot{\text{I}}\text{C}\Gamma\text{C}, \text{PSC}\Pi$
 $\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{P}^\circ\text{P}\ddot{\text{I}}\text{C}-\text{C}\Pi\text{PjP}\text{sP}\P\text{P}\mu\text{C}\Gamma\text{C}, \text{P}^\circ\text{P}\text{eP}\text{sPjC}\Gamma\text{P}\ddot{\text{e}}\text{P}\text{IP}^\circ\text{PrP}\text{eC}\Gamma\text{C}, \text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{P}\text{IP}^\circ\text{C}, \text{P}\ddot{\text{e}}\text{P}\ddot{\text{I}}\text{PsPSP}^\circ\text{Pr } 6$
 $\text{PjC}-\text{C}\Gamma\text{C}\Pi\text{C}\dagger\text{C}-\text{PIPrP}\text{sC}, \text{PsP}\text{IPs}, \text{C}\Pi\text{P}\text{eP}\pm\text{C}\Gamma\text{PrP}\mu\text{PrP}\text{sC}\Gamma\text{C}\Pi\text{P}\text{IPSC}\Gamma\text{C}, \text{PsPSPsP}\text{IPsC}-\text{B}$
 $\text{C}\tilde{\text{H}}\text{C}-\text{P}\text{IPSPsP}\text{IP}^\circ\text{P}\P\text{PSPsC}-\text{P}\text{eP}\text{sPSC}\dagger\text{P}\mu\text{PSC}, \text{C}\tilde{\text{H}}\text{P}^\circ\text{C}\dagger\text{C}-\text{C}-.$

$\text{P}-\text{P}^\circ\text{C}\Gamma\text{C}, \text{PsC}\Gamma\text{C}\Gamma\text{P}\text{IP}^\circ\text{PSPSC}\Pi\text{ } 12\text{PiC}\dots\text{PsP}\gg\text{P}\mu\text{C}\Gamma\text{C}, \text{P}\ddot{\text{e}}\text{C}\tilde{\text{H}}\text{P}^\circ\text{PjC}-\text{PSC}\Gamma\text{C}\ddagger\text{P}\mu\text{C}\tilde{\text{H}}\text{P}\mu\text{P}\cdot\text{P}\ddot{\text{I}}\text{PsPrP}\ddot{\text{e}}\text{PSC}\Gamma$
 $\text{P}\ddot{\text{I}}\text{C}-\text{C}\Gamma\text{P}\gg\text{C}\Pi\text{P}\cdot\text{P}^\circ\text{C}\Gamma\text{C}, \text{PsC}\Gamma\text{C}\Gamma\text{P}\text{IP}^\circ\text{PSPSC}\Pi\text{PsPrPSPsC}\tilde{\text{H}}\text{P}^\circ\text{P}\cdot\text{PsP}\text{IPsC}-\text{PrP}\text{sP}\cdot\text{P}\ddot{\text{e}}\text{ } 5\text{PjPi}$
 $\text{PrC}\Gamma\text{C}, \text{P}^\circ\text{C}\Gamma\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{PrC}\Gamma\text{C}\text{PSP}\mu\text{P}\text{IPiP}\gg\text{P}\ddot{\text{e}}\text{P}\text{IP}^\circ\text{P}\gg\text{PsPSP}^\circ\text{C}, \text{P}^\circ\text{C}\tilde{\text{H}}\text{PjP}^\circ\text{P}\text{eP}\text{sP}\text{eC}-\text{PSP}\mu\text{C}, \text{P}\ddot{\text{e}}\text{P}\text{eC}\Gamma$
 $\text{PrC}\Gamma\text{C}, \text{P}^\circ\text{C}\Gamma\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{PrC}\Gamma\text{C}.$

P'P\ddot{I}P\ggP\ddot{e}PIPrC\acute{z}C, P^\circ C\Gamma C, P\mu C\tilde{H}P\ddot{e}PrC\acute{z}PSP^\circ C, P^\circ C\tilde{H}PjP^\circ P\text{eP}\text{sP}\text{eC}-PSP\mu C, P\ddot{e}P\text{eC}\acute{z}C-PSC\epsilon P\ddot{e}C\dots
P\gg C-P\text{eP}^\circ C\tilde{H}C\Gamma C\tilde{H}P\text{eP}\ddot{e}C\dots P\cdot P^\circ C\Gamma P\text{sP}\pm C-PI

$\text{PJ PSP}\mu\text{P}\text{IP}\mu\text{P}\gg\text{P}\ddot{\text{e}}\text{P}\text{eP}\text{sPjC}\Gamma\text{PrP}\text{sC}\Gamma\text{P}\gg\text{C}-\text{PrP}\P\text{P}\mu\text{PSPSC}-\text{(N=24)C}, \text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{P}\text{IP}^\circ\text{P}\gg\text{C}-\text{C}\Gamma\text{C}, \text{C}\tilde{\text{H}}\text{PIPrP}\text{IP}^\circ$
 $\text{C}, \text{P}\ddot{\text{e}}\text{P}\P\text{PSC}-\text{P}\cdot\text{P}^\circ\text{C}\Gamma\text{C}\ddagger\text{P}^\circ\text{C}\Gamma\text{C}, \text{C}\tilde{\text{H}}\text{P}\cdot\text{PrP}\text{sC}\tilde{\text{H}}\text{P}\text{sP}\text{IP}\ddot{\text{e}}\text{C}\dots\text{C}\ddagger\text{PsP}\gg\text{PsP}\text{IC}-\text{P}\text{eC}-\text{PIPrC}\Gamma\text{C}, \text{P}^\circ\text{C}\Gamma\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{Pr}$
 $\text{(0,5PSP}^\circ\text{PrP}\text{sP}\pm\text{C}\Gamma\text{P})\text{PSP}\mu\text{P}\text{IPiP}\gg\text{P}\ddot{\text{e}}\text{P}\text{IP}^\circ\text{PIPSP}^\circ\text{C}, \text{P}^\circ\text{C}\tilde{\text{H}}\text{PjP}^\circ\text{P}\text{eP}\text{sP}\text{eC}-\text{PSP}\mu\text{C}, \text{P}\ddot{\text{e}}\text{P}\text{eC}\Gamma$
 $\text{C}, \text{P}^\circ\text{PjC}\Gamma\text{C}\Gamma\text{P}\gg\text{PsP}\cdot\text{P}\ddot{\text{e}}\text{PSC}\Gamma\text{P}^\circ\text{P}\pm\text{PsC}, \text{P}\mu\text{C}\tilde{\text{H}}\text{P}^\circ\text{P}\cdot\text{PsP}\cdot\text{P}\ddot{\text{e}}\text{PSC}\Gamma\text{P}.$
 $\text{PJ C}\dagger\text{C}\tilde{\text{H}}\text{PsPjC}\Gamma\text{B}$
 $\text{PrP}\text{sC}\Gamma\text{P}\gg\text{C}-\text{PrP}\P\text{P}\mu\text{PSPSC}-\text{C}, \text{P}^\circ\text{P}\text{eP}\text{sP}\P\text{P}\mu\text{PSP}\mu\text{P}\pm\text{C}\Gamma\text{P}\gg\text{PsP}\text{IP}\ddot{\text{e}}\text{C}\Pi\text{IP}\gg\text{P}\mu\text{PSPsPsP}\cdot\text{PSP}^\circ\text{P}\text{e}$
 $\text{C}, \text{P}^\circ\text{C}\tilde{\text{H}}\text{PjP}^\circ\text{P}\text{eP}\text{sPrP}\ddot{\text{e}}\text{PSP}^\circ\text{PjC}-\text{C}\ddagger\text{PSPsC}-\text{P}\text{IP}\cdot\text{P}^\circ\text{C}''\text{PjP}\text{sPrC}-\text{C}-.$

$\text{P}''\text{C}\Gamma\text{C}, \text{P}^\circ\text{C}\Gamma\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{PrPSP}\mu\text{P}\text{IPiP}\gg\text{P}\ddot{\text{e}}\text{P}\text{IP}^\circ\text{C}''\text{PSP}^\circ\text{C}, \text{P}^\circ\text{C}\tilde{\text{H}}\text{PjP}^\circ\text{P}\text{eP}\text{sP}\text{eC}-\text{PSP}\mu\text{C}, \text{P}\ddot{\text{e}}\text{P}\text{eC}\Gamma$
 $\text{P}\text{IP}^\circ\text{C}\tilde{\text{H}}\text{C}, \text{P}^\circ\text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{PSC}\Gamma\text{P}^\circ\text{P}\pm\text{PsPrP}\ddot{\text{e}}\text{P}\text{IPsP}\text{eC}\Gamma\text{P}\ddot{\text{e}}\text{PSC}\Gamma\text{P}.$
 $\text{P}\ddot{\text{I}}\text{P}\mu\text{P}\text{IP}\text{eP}^\circ\text{P}\cdot\text{C}\Gamma\text{C}''\text{PSP}^\circ\text{C}, \text{P}\mu, \text{C}\%_{\text{O}}\text{Ps}$
 $\text{PrC}\Gamma\text{C}, \text{P}^\circ\text{C}\Gamma\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{PrPSP}\mu\text{C}-\text{PSPiC}-\text{P}\pm\text{C}\Gamma\text{C}''/\text{PSP}\mu\text{C}-\text{PSP}\Gamma\text{C}\Gamma\text{P}\text{eC}\Gamma\text{C}''\text{P}^\circ\text{P}\text{eC}, \text{P}\ddot{\text{e}}\text{P}\text{IPSC}-\text{C}\Gamma\text{C}, \text{C}\tilde{\text{H}}$
 $\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{PjP}\mu\text{PSC}, \text{C}\Gamma\text{CYP2C9P}^\circ\text{P}\pm\text{PsP}\cdot\text{P}\ddot{\text{I}}\text{P}\gg\text{C}-\text{P}\text{eP}\text{sP}\ddot{\text{I}}\text{C}\tilde{\text{H}}\text{P}\text{sC}, \text{P}\mu\text{C}-\text{PSC}\Gamma\text{P}.$
 $\text{P}\ddot{\text{I}}\text{P}\mu\text{C}\tilde{\text{H}}\text{P}\mu\text{PSPsC}\Gamma\text{PSP}\ddot{\text{e}}\text{P}\text{eP}^\circ\text{P}^\circ\text{P}\text{PSC}-\text{PrP}\text{sC}\Gamma\text{P}\gg\text{C}-\text{PrP}\P\text{P}\mu\text{PSC}\tilde{\text{H}}\text{P}\text{IP}\cdot\text{P}^\circ\text{C}''\text{PjP}\text{sPrC}-\text{C}-\text{in vitro}$
 $\text{P}\text{IP}\text{eP}^\circ\text{P}\cdot\text{C}\Gamma\text{C}\tilde{\text{H}}\text{C}, \text{C}\tilde{\text{H}}\text{PSP}^\circ\text{C}, \text{P}\mu, \text{C}\%_{\text{O}}\text{PsPrC}\Gamma\text{C}, \text{P}^\circ\text{C}\Gamma\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{PrPSP}\mu\text{C}-\text{PSPiC}-\text{P}\pm\text{C}\Gamma\text{C}''$
 $\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{PjP}\mu\text{PSC}, \text{P}\ddot{\text{e}}\text{CYP1A2}, \text{CYP2D6}, \text{CYP2C9}, \text{CYP2C19P}^\circ\text{P}\pm\text{PsCYP3A4}.$

P\ddot{y}P^\circ PjC\Gamma C\acute{z}P\gg PsP\cdot P\ddot{e}PS

$\text{P}\tilde{\text{H}}\text{PrPSPsC}\ddagger\text{P}^\circ\text{C}\Gamma\text{PSP}\mu\text{P}\cdot\text{P}^\circ\text{C}\Gamma\text{C}, \text{PsC}\Gamma\text{C}\Gamma\text{P}\text{IP}^\circ\text{PSPSC}\Pi\text{C}, \text{P}^\circ\text{PjC}\Gamma\text{C}\Gamma\text{P}\gg\text{PsP}\cdot\text{P}\ddot{\text{e}}\text{PSC}\Gamma$
 $\text{P}\ddot{\text{I}}\text{C}-\text{PrC}\tilde{\text{H}}\text{PsC}\dots\text{P}\gg\text{PsC}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{PrC}\Gamma\text{P}\cdot\text{P}\gg\text{C}-\text{P}\text{eP}^\circ\text{C}\tilde{\text{H}}\text{C}\Gamma\text{C}\tilde{\text{H}}\text{P}\text{eP}\ddot{\text{e}}\text{PjP}\ddot{\text{e}}\text{P}\cdot\text{P}^\circ\text{C}\Gamma\text{PsP}\pm\text{P}^\circ\text{PjP}\ddot{\text{e}}, \text{C}\Pi\text{P}\text{eC}-$
 $\text{PjP}\text{sP}\P\text{C}\Gamma\text{C}, \text{C}\tilde{\text{H}}\text{P}\cdot\text{PSP}\ddot{\text{e}}\text{P}\P\text{C}\Gamma\text{P}\text{IP}^\circ\text{C}, \text{P}\ddot{\text{e}}\text{P}^\circ\text{C}\tilde{\text{H}}\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{C}-\text{P}^\circ\text{P}\gg\text{C}\tilde{\text{H}}\text{PSP}\ddot{\text{e}}\text{PN}\ddot{\text{e}}\text{C}, \text{P}\ddot{\text{e}}\text{C}\Gamma\text{P}\text{e}, \text{C}\Gamma\text{C}, \text{PsPjC}\Gamma$
 $\text{C}\ddagger\text{P}\ddot{\text{e}}\text{C}\Gamma\text{P}\gg\text{C}-\text{C}\tilde{\text{H}}\text{P}^\circ\text{P}\cdot\text{PsPjP}\cdot\text{C}-\text{P}\cdot\text{PSP}\mu\text{P}\pm\text{PsP}\gg\text{C}\tilde{\text{H}}\text{P}\text{IP}^\circ\text{P}\gg\text{C}\tilde{\text{H}}\text{PSP}\ddot{\text{e}}\text{PjP}\ddot{\text{e}}\text{P}\gg\text{C}-\text{P}\text{eP}^\circ\text{C}\tilde{\text{H}}\text{C}\Gamma\text{C}\tilde{\text{H}}\text{P}\text{eP}\ddot{\text{e}}\text{PjP}\ddot{\text{e}}$
 $\text{P}\cdot\text{P}^\circ\text{C}\Gamma\text{PsP}\pm\text{P}^\circ\text{PjP}\ddot{\text{e}}, \text{C}-\text{PSPiC}-\text{P}\pm\text{C}-\text{C}, \text{PsC}\tilde{\text{H}}\text{P}^\circ\text{PjP}\ddot{\text{e}}\text{ } 5\text{-C}, \text{PsC}\Gamma\text{C}, \text{PsPrC}-\text{P}\mu\text{C}\Gamma\text{C}, \text{P}\mu\text{C}\tilde{\text{H}}\text{P}^\circ\text{P}\cdot\text{P}\ddot{\text{e}}\text{C}, \text{P}^\circ$
 $\text{C}-\text{PSC}\epsilon\text{P}\ddot{\text{e}}\text{PjP}\ddot{\text{e}}\text{P}^\circ\text{P}\gg\text{C}\tilde{\text{H}}\text{C}, \text{P}^\circ\text{-1-P}^\circ\text{PrC}\tilde{\text{H}}\text{P}\mu\text{PSPsP}\pm\text{P}\gg\text{PsP}\text{eP}^\circ\text{C}, \text{PsC}\tilde{\text{H}}\text{P}^\circ\text{PjP}\ddot{\text{e}}\text{PjP}\text{sP}\P\text{P}\mu$
 $\text{C}, \text{P}\mu\text{PsC}\tilde{\text{H}}\text{P}\mu\text{C}, \text{P}\ddot{\text{e}}\text{C}\ddagger\text{PSPsB}\text{P}\ddot{\text{I}}\text{C}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{P}\cdot\text{P}\text{IPsPrP}\ddot{\text{e}}\text{C}, \text{P}\ddot{\text{e}}\text{PrP}\text{sP}\ddot{\text{I}}\text{PsC}\Gamma\text{P}\ddot{\text{e}}\text{P}\gg\text{P}\mu\text{PSPsC}-$
 $\text{P}\ddot{\text{I}}\text{C}-\text{P}\ddot{\text{I}}\text{PsC}, \text{P}\mu\text{PSP}\cdot\text{P}\ddot{\text{e}}\text{P}\text{IPSPsC}-\text{PrC}-\text{C}-.$
 $\text{P}''\text{C}\Gamma\text{PsPrP}^\circ\text{C}\tilde{\text{H}}\text{C}, \text{PSP}\mu\text{C}\Gamma\text{P}\gg\text{C}-\text{PrP}\cdot\text{P}^\circ\text{C}\Gamma\text{C}, \text{PsC}\Gamma\text{PsP}\text{IC}\Gamma\text{P}\text{IP}^\circ\text{C}, \text{P}\ddot{\text{e}}\text{PIPeP}\text{sPjP}\pm\text{C}-\text{PSP}^\circ\text{C}\dagger\text{C}-\text{C}-\text{P}\cdot\text{C}-\text{PSC}\epsilon\text{P}\ddot{\text{e}}\text{PjP}\ddot{\text{e}}\text{P}^\circ\text{P}\gg\text{C}\tilde{\text{H}}\text{C}, \text{P}^\circ\text{-1-}$
 $\text{P}^\circ\text{PrC}\tilde{\text{H}}\text{P}\mu\text{PSPsP}\pm\text{P}\gg\text{PsP}\text{eP}^\circ\text{C}, \text{PsC}\tilde{\text{H}}\text{P}^\circ\text{PjP}\ddot{\text{e}}.$

$\text{P}\tilde{\text{H}}\text{PrPSPsC}\ddagger\text{P}^\circ\text{C}\Gamma\text{PSP}\mu\text{P}\cdot\text{P}^\circ\text{C}\Gamma\text{C}, \text{PsC}\Gamma\text{C}\Gamma\text{P}\text{IP}^\circ\text{PSPSC}\Pi\text{C}, \text{P}^\circ\text{PjC}\Gamma\text{C}\Gamma\text{P}\gg\text{PsP}\cdot\text{P}\ddot{\text{e}}\text{PSC}\Gamma$
 $\text{P}\ddot{\text{I}}\text{C}-\text{PrC}\tilde{\text{H}}\text{PsC}\dots\text{P}\gg\text{PsC}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{PrC}\Gamma\text{C}, \text{P}^\circ\text{PeP}\mu\text{C}, \text{PsPeP}\text{sPSP}^\circ\text{P}\cdot\text{PsP}\gg\text{C}\Gamma\text{(C}\Gamma\text{P}\ddot{\text{e}}\text{P}\gg\text{C}\tilde{\text{H}}\text{PSPsP}\text{IPs}$
 $\text{C}-\text{PSPiC}-\text{P}\pm\text{C}-\text{C}, \text{PsC}\tilde{\text{H}}\text{P}^\circ\text{CYP3A4})\text{B}\text{B}\text{P}\cdot\text{P}\pm\text{C}-\text{P}\gg\text{C}\tilde{\text{H}}\text{C}\epsilon\text{C}\Gamma\text{C}''\text{P}\ddot{\text{Y}}_{\text{max}}\text{C}, \text{P}^\circ\text{AUC}$
 $\text{C}, \text{P}^\circ\text{PjC}\Gamma\text{C}\Gamma\text{P}\gg\text{PsP}\cdot\text{P}\ddot{\text{e}}\text{PSC}\Gamma\text{P}\ddot{\text{I}}\text{C}-\text{PrC}\tilde{\text{H}}\text{PsC}\dots\text{P}\gg\text{PsC}\tilde{\text{H}}\text{P}\ddot{\text{e}}\text{PrC}\Gamma\text{C}\text{C}\Gamma\text{ } 2,2\text{C}, \text{P}^\circ\text{ } 2,8\text{C}\tilde{\text{H}}\text{P}^\circ\text{P}\cdot\text{P}^\circ$

PIC-PrPⁱPsPIC-PrPSPs.

P^YC^rPjC-C^rPSP_μ P·P°C^rC,PsC^rC^rPIP°PSPSC_μ C,P°PjC^rC^rP»PsP·PëPSC^r
PiC-PrC_μPsC...P»PsC_μPëPrC^r C,P° PⁱP°C_μPsPeC^rP_μC,PëPSC^r (C^rPëP»CH_μPSPsPiPs
C-PSPiC-P±C-C,PsC_μP° CYP2D6) P·P±C-P»CH_μC€C^rC” P^Y_{max} C,P° AUC C,P°PjC^rC^rP»PsP·PëPSC^r
PiC-PrC_μPsC...P»PsC_μPëPrC^r C^r 1,3 C,P° 1,6 C_μP°P·P° PIC-PrPⁱPsPIC-PrPSPs.
P_μPSP°P»PsPiC-C_μPSP_μ P·P±C-P»CH_μC€P_μPSPSC_μ PsC_μC-PeC^rC”C,CH_μC^rC_μ C^r
PⁱP°C^rC-C”PSC,C-PI P·C- C^rP»P°P±PePëPj PjP_μC,P°P±PsP»C-P·PjPsPj CYP2D6
PⁱPsC_μC-PIPSC_μPSPs P· C,P°PePëPjPë P· C-PSC,P_μPSC^rPëPIPSPëPj PjP_μC,P°P±PsP»C-P·PjPsPj
PiC_μPë C^rC^rPjC-C^rPSPsPjC^r P·P°C^rC,PsC^rC^rPIP°PSPSC- P· C^rPëP»CH_μPSPëPjPë
C-PSPiC-P±C-C,PsC_μP°PjPë CYP3A4.

P·C,,P_μPeC, C^rC^rPjC-C^rPSPsPiPs P·P°C^rC,PsC^rC^rPIP°PSPSC_μ PsP±PsC...
C-PSPiC-P±C-C,PsC_μC-PI CYP3A4 C,P° C-PSPiC-P±C-C,PsC_μC-PI CYP2D6 P·
C,P°PjC^rC^rP»PsP·PëPSPsPj PeP»C-PSC-C_μPSPs PSP_μ PrPsC^rP»C-PrP^μC^rPIP°PIC^rC_μ, PsPrPSP°Pe
PⁱPsC,P_μPSC^rC-PM_μPSPs PjPsP^μP_μ C^rC^rC,C,C”PIPs P·P±C-P»CH_μC€C^rPIP°C,PëC^rC_μ
PePsPSC^rP_μPSC,C_μP°C^rC-C_μ C,P°PjC^rC^rP»PsP·PëPSC^r (PrPëPI. C_μPsP·PrC-P»
B«PhC^rPsP±P»PëPIPsC^rC,C- P·P°C^rC,PsC^rC^rPIP°PSPSC_μB»).

PhPrPSPsC_μP°C^rPSP_μ P·P°C^rC,PsC^rC^rPIP°PSPSC_μ C,P°PjC^rC^rP»PsP·PëPSC^r
PiC-PrC_μPsC...P»PsC_μPëPrC^r (0,4 PjPi) C- C_μPëPjP_μC,PëPrPëPSC^r (400 PjPi PePsP^μPSC- 6
PiPsPrPëPS PiC_μPsC,C_μPiPsPj 6 PrPSC-PI) PiC_μPëP·PIPsPrPëP»Ps PrPs P·PSPëP^μP_μPSPSC_μ
PeP»C-C_μP_μPSC^rC^r (26 %) C- P·P±C-P»CH_μC€P_μPSPSC_μ AUC (PⁱP»PsC%_μC- PiC-Pr
C,,P°C_μPjP°PePsPeC-PSP_μC,PëC_μPSPsC_μ PeC_μPëPIPsC_μ) (44 %) C,P°PjC^rC^rP»PsP·PëPSC^r
PiC-PrC_μPsC...P»PsC_μPëPrC^r. P”C^rPsPrP°C_μC, C^rP»C-Pr P· PsP±P_μC_μP^μP^μPSC-C^rC,C_μ
P·P°C^rC,PsC^rPsPIC^rPIP°C,Pë PI PePsPjP±C-PSP°C^rC-C— P· C_μPëPjP_μC,PëPrPëPSPsPj.

P’PëC_μP_μC_μPⁱPSP_μ PrPsC^rP»C-PrP^μP_μPSPSC_μ PIP·P°C”PjPsPrC-C— C,P°PjC^rC^rP»PsP·PëPSC^r
PiC-PrC_μPsC...P»PsC_μPëPrC^r C- PIP°C_μC,,P°C_μPëPSC^r PSP_μ PiC_μPsPIPsPrPëP»PsC^rC_μ.
P P_μP·C^rP»CH_μC,P°C,Pë PsP±PjP_μP^μP_μPSPëC... PrPsC^rP»C-PrP^μP_μPSC_μ *in vitro* C,P° *in vivo* C”
PSP_μPrPsC^rC,P°C,PSC-PjPë. P— PsP±P_μC_μP_μP^μPSC-C^rC,C_μ C^rP»C-Pr PiC_μPsPIPsPrPëC,Pë
PsPrPSPsC_μP°C^rPSP_μ P»C-PeC^rPIP°PSPSC_μ PIP°C_μC,,P°C_μPëPSPsPj C-
C,P°PjC^rC^rP»PsP·PëPSC^r PiC-PrC_μPsC...P»PsC_μPëPrPsPj.

P_μC-C_μPePsC— PIP·P°C”PjPsPrC-C— PSP_μ C^rPiPsC^rC,P_μC_μC-PiP°P»PsC^rC_μ, PePsP»Pë
C,P°PjC^rC^rP»PsP·PëPSC^r PiC-PrC_μPsC...P»PsC_μPëPr PIPiPsPrPëP»Pë PsPrPSPsC_μP°C^rPSPs P·
P°C,P_μPSPsP»PsP»PsPj P°P±Ps P_μPSP°P»P°PiC_μPëP»PsPj, P°P±Ps PSC-C,,P_μPrPëPiC-PSPsPj,
P°P±Ps C,P_μPsC,,C-P»C-PSPsPj. PhPrPSPsC_μP°C^rPSP_μ P·P°C^rC,PsC^rC^rPIP°PSPSC_μ
C,,C^rC_μPsC^rP_μPjC-PrC^r PiC_μPëP·PIPsPrPëC,CH PrPs P·PSPëP^μP_μPSPSC_μ C_μC-PIPSC-PI
C,P°PjC^rC^rP»PsP·PëPSC^r PI C^rPëC_μPsPIP°C,C^rC- PeC_μPsPIC-, P°P»P_μ, PsC^rPeC-P»CH_μPePë
C^rC- C_μC-PIPSC- P·P°P»PëC€P°C_μC,CH_μC^rC_μ C^r PjP_μP^μP°C... PSPsC_μPjP°P»CH_μPSPsPiPs
PrC-P°PiP°P·PsPSC^r, PePsC_μP_μPeC^rC-C_μ PrPsP·Pë PSP_μ PiPsC,C_μC-P±PSP°.

P’ C^rPjPsPIP°C... *in vitro* PSC- PrC-P°P·P_μPiP°Pj, PSC- PiC_μPsPiC_μP°PSPsP»PsP», PSC-
C,C_μPëC...P»PsC_μPjP_μC,C-P°P·PëPr, PSC- C...P»PsC_μPjP°PrPëPSPsPS, PSC-
P°PjC-C,C_μPëPiC,PëP»C-PS, PSC- PrPëPeP»PsC,,P_μPSP°Pe, PSC- PiP»C-P±P_μPSPeP»P°PjC-Pr,
PSC- C^rPëPjPIP°C^rC,P°C,PëPS PSP_μ P·PjC-PSC_μC_μC,CH PIC-P»CH_μPSC^r C,,C_μP°PeC^rC-C_μ
C,P°PjC^rC^rP»PsP·PëPSC^r C^r PiP»P°P·PjC- PeC_μPsPIC- P»CH_μPrPëPSPë. P^YP°PjC^rC^rP»PsP·PëPS
C,P°PePsP^μ PSP_μ P·PjC-PSC_μC” PIC-P»CH_μPSC- C,,C_μP°PeC^rC-C— PrC-P°P·P_μPiP°PjC^r,
PiC_μPsPiC_μP°PSPsP»PsP»C^r, C,C_μPëC...P»PsC_μPjP_μC,C-P°P·PëPrC^r C-
C...P»PsC_μPjP°PrPëPSPsPSC^r.

PsC†C-PSC†PIP°P»Pë C†PëP·PëPe C†PsP·PIPëC,PeC† P†PsP±C-C†PSPëC... C†PµP°PeC†C-PNë P·
P±PsPeC† C†PµC†PµPIP-S-C†C†PrPëPSPSPsC— C†PëC†C,PµPjPë P†C†Pë
P·P°C†C,PsC†C†PIP°PSPSC- PrC†C,P°C†C,PµC†PëPrC† (P†PsC†C-PIPSC†PSPs P·
PePsPSC,C†PsP»C†PSPsC† P†C†C†P†PsC†). P†Pµ P±C†P»Ps PIC†C,P°PSPsPIP»PµPSPs
C†C,C-PNëPePsPiPs C†C,P°C,PëC†C,PëC†PSPs P·PSP°C†C†C%oPsPiPs P·P±C-P»C†C†PµPSPSC†
C†PëP·PëPeC†B C†PµC†PµPIP-S— PSPµPrPsC†C,P°C,PSPsC†C,C- (RR 1,05; 95 % P†P† 0,71,
1,57), P†PsC†C,C†PsPiPs C-PSC,P°C†PeC,C† PjC-PsPeP°C†PrP° B B B B B B B B B B (RR 1,00;
95 % P†P† 0,77, 1,30) P°P±Ps C-PSC†C†P»C†C,C† (RR 1,20; 95 % P†P† 0,88, 1,64).

P P°Pe P†PµC†PµPrPjC-C...C†C†PsPIP-S— P·P°P»PsP·Pë C,P° P†C†C...P»PëPSPë PIPëC†PsPePsPiPs
C†C,C†P†PµPSC† P†C†P°PrP°C†C-C— P·P° P°P»C-C†PsPSPsPj (PSPëP·C†PePs-
PrPëC,,PµC†PµPSC†C-PNëPsPIP°PSC-)

Pj C...PsPrC- 4-C†C-C†PSPsPiPs PeP»C-PSC-C†PSPsPiPs PrPsC†P»C-PrPµPµPSPSC† P·P°
C†C†P°C†C,C† > 8000 C†PsP»PsPIC-PeC-PI PIC-PePsPj PIC-Pr 50 PrPs 75 C†PsPeC-PI P·
P†PsPiPµC†PµPrPSC†PsC† PSPµPiP°C,PëPIPSPsC† P±C-PsPiC†C-C†C† PSP° C†P°Pe
P†PµC†PµPrPjC-C...C†C†PsPIP-S— P·P°P»PsP·Pë C,P° C,,PsPSPsPIPëPj C†C-PIPSPµPj PSA PjC-Pµ
2,5 PSp†PjP» C,P° 10,0 PSp†PjP» (PrPsC†P»C-PrPµPµPSPSC† REDUCE) C† 1517
C†PsP»PsPIC-PeC-PI P±C†P»Ps PrC-P°PiPSPsC†C,PsPIP°PSPs C†P°Pe
P†PµC†PµPrPjC-C...C†C†PsPIP-S— P·P°P»PsP·Pë. P°C-PrP·PSP°C†P°P»P°C†C† PIPëC%oP°
C†P°C†C,PsC,P° PIPëPSPëPePSPµPSPSC† C†P°PeC† P†PµC†PµPrPjC-C...C†C†PsPIP-S—
P·P°P»PsP·Pë P· PrPëC,,PµC†PµPSC†C-C†PIP°PSPSC†Pj 8B†“10 P±P°P»C-PI P·P° C†PeP°P»PsC†
P°P»C-C†PsPSP° C† P†C†C†P†C- PrC†C,P°C†C,PµC†PëPrC† (n=29, 0,9 %) P†PsC†C-PIPSC†PSPs
P· P†C†C†P†PsC† P†P»P°C†PµP±Ps (n=19, 0,6 %). PµC†PëC†PëPSPSPsPiPs
PIP·P°C†PjPsP·P†B†™ C†P·PeC† PjC-Pµ P·P°C†C,PsC†C†PIP°PSPSC†Pj PrC†C,P°C†C,PµC†PëPrC†
C,P° PIPëPSPëPePSPµPSPSC†Pj PSPëP·C†PePsPrPëC,,PµC†PµPSC†C-PNëPsPIP°PSPsPiPs
C†P°PeC† P†PµC†PµPrPjC-C...C†C†PsPIP-S— P·P°P»PsP·Pë PIC†C,P°PSPsPIP»PµPSPs PSPµ
P±C†P»Ps. PµP»C-PSC-C†PSP° P·PSP°C†C†C%oC-C†C,C† C†PëC,,C†PsPIP-SPiPs
PrPëC†P±P°P»P°PSC†C† PSPµ PIC†C,P°PSPsPIP»PµPSP°.B

P†PsP»PsPIC-PePë, C†PeC- P·P°C†C,PsC†PsPIC†C†C,C† P°C†PsPrP°C†C,, PjP°C†C,C†
C†PµPiC†P»C†C†PSPs P†C†PsC...PsPrPëC,Pë PsP±C†C,PµPµPµPSPSC† C%oPsPrPs
PIPëP·PSP°C†PµPSPSC† C†PëP·PëPeC† C†PsP·PIPëC,PeC† C†P°PeC†
P†PµC†PµPrPjC-C...C†C†PsPIP-S— P·P°P»PsP·Pë, PIPëP»C†C†P°C†C†Pë C,PµC†C, PSP°
P†C†PsC†C,P°C,-C†P†PµC†PëC,,C-C†PSPëPNë P°PSC,PëPiPµPS.

Pj PrPsPrP°C,PePsPiPsPjC† P†PsC†P»C-PrPsPIPSPsPjC† 2-C†C-C†PSPsPjC†
C†P†PsC†C,PµC†PµPµPµPSPSC- PsC†PëPiC-PSP°P»C†PSPëC... P†P°C†C-C†PSC,C-PI P†C†Pë
P·P°C†C,PsC†C†PIP°PSPSC- B PrC†C,P°C†C,PµC†PëPrC† PI C†PePsC†C,C- C...C-PjC-C†PSPsC—
P†C†PsC,,C-P»P°PeC,PëPePë (PrPsC†P»C-PrPµPµPSPSC† REDUCE) P±C†P»P°
PIC†C,P°PSPsPIP»PµPSP° PSPëP·C†PeP° C†P°C†C,PsC,P° PSPsPIPëC... PIPëP†P°PrPeC-PI
C†P°PeC† P†PµC†PµPrPjC-C...C†C†PsPIP-S— P·P°P»PsP·Pë (P†C†C†P†P°
PrC†C,P°C†C,PµC†PëPrC† [n=14, 1,2 %]) C,P° P†C†C†P†P° P†P»P°C†PµP±Ps [n=7, 0,7 %]) P·
PIC-PrC†C†C,PSC-C†C,C† PSPsPIPëC... C-PrPµPSC,PëC,,C-PePsPIP°PSPëC... PIPëP†P°PrPeC-PI
C†P°PeC† P†PµC†PµPrPjC-C...C†C†PsPIP-S— P·P°P»PsP·Pë P·
PrPëC,,PµC†PµPSC†C-C†PIP°PSPSC†Pj 8B†“10 P±P°P»C-PI P·P° C†PeP°P»PsC†
P°P»C-C†PsPSP°.

P°P†PiPsC,C†PëPIP°P»Pµ P†PsC†P»C-PrPsPIPSPµ (PrPs 18 C†PsPeC-PI)
C†P†PsC†C,PµC†PµPµPµPSPSC† P†P°C†C-C†PSC,C-PI P· PeP»C-PSC-C†PSPsPiPs
PrPsC†P»C-PrPµPµPSPSC† C-P· P·P°C†C,PsC†C†PIP°PSPSC†Pj C-PSC†PsPiPs
C-PSPiC-P±C-C,PsC†P° 5α-C†PµPrC†PeC,P°P·Pë (C,,C-PSP°C†C,PµC†PëPrC†) PI C†PePsC†C,C-

C...C-PjC-C†PSPsC— P†CḡPsC,,C-P»P°PeC,PëPePë PSPµ P†PsPeP°P·P°P»Ps
C†C,P°C,PëC†C,PëC†PSPs P·PSP°C†CrC%PsC— CḡC-P·PSPëC†C- PjC-P¶ P†CḡCrP†P°PjPë
C,,C-PSP°C†C,PµCḡPëPrCr C,P° P†P»P°C†PµP±Ps Cr P†PsPeP°P·PSPëPeP°C...
P·P°P†P°P»CHPSPsPiPs P†PëP¶PëP†P°PSPSC¶ (HR1,02, 95 % P”P† 0,97Бḡ“1,08) P°P±Ps
P†PëP¶PëP†P°PSPSC¶ P†C-C†P»C¶ PrC-P°PiPSPsC†C,C†P†P°PSPSC¶ CḡP°PeCr
P†PµCḡPµPrPjC-C...CrCḡPsP†PsC— P·P°P»PsP·Pë (HR1,01, 95 % P”P† 0,85Бḡ“1,20).B B B

P’P†P»PëP† PSP° P†CḡPsC†C,P°C,-C†P†PµC†PëC,,C-C†PSPëPNë P°PSC,PëP†PµPS (PSA)

PḡPsPSC†PµPSC,CḡP°C†C-C¶ P†CḡPsC†C,P°C,-C†P†PµC†PëC,,C-C†PSPsPiPs P°PSC,PëP†PµPSC†
P†PµCḡPµPrPjC-C...CrCḡPsP†PsC— P·P°P»PsP·Pë (PSA) B C” P†P°P¶P»PëP†PëPj
PePsPjP†PsPSPµPSC,PsPj C†PeCḡPëPSC-PSP†PsP†PsP†Ps P†CḡPsC†PµC†C† PrP»C¶
P†PëC¶P†P»PµPSPSC¶ CḡP°PeCr P†PµCḡPµPrPjC-C...CrCḡPsP†PsC— P·P°P»PsP·Pë.
P”CrC,P°C†C,PµCḡPëPr P·PrP°C,PSPëPNë P·PSPëP¶C†P†P°C,Pë CḡC-PIPµPSCḡ
C†PëCḡPsP†P°C,PePsP†PsP†Ps P†CḡPsC†C,P°C,-C†P†PµC†PëC,,C-C†PSPsPiPs P°PSC,PëP†PµPSC† Cr
C...P†PsCḡPëC... P†CḡPëP±P»PëP·PSPs PSP° 50 % C†PµCḡPµP· 6 PjC-C†C¶C†C-PI
P»C-PëC†P†P°PSPSC¶.

PµP°C†C-C”PSC,Pë, C¶PeC- P†CḡPëPNëPjP°CḡC,CH P”CrPsPrP°CḡC,, P†PsP†PëPSPSC- PjP°C,Pë
PSPsP†PëPNë P†PëC...C-PrPSPëPNë CḡC-PIPµPSCḡ PSA, P†C†C,P°PSPsP†P»PµPSPëPNë C†PµCḡPµP·
6 PjC-C†C¶C†C-PI P†C-C†P»C¶ P»C-PëC†P†P°PSPSC¶ C†PëPj P†CḡPµP†P°CḡP°C,PsPj.
PḡP°PrP°P»C- C†PµPNë CḡC-PIPµPSCḡ CḡPµPePsPjPµPSP†CrC”C,CHC†C¶
P†PµCḡPµPIC-CḡC¶C,Pë CḡPµP†CrP»C¶CḡPSPs. P”CrPrCH-C¶PePµ P†C-PrC,PIPµCḡPrP¶PµPSPµ
P·P±C-P»CHC€PµPSPSC¶ CḡC-PIPSC¶ PSA PIC-Pr PSP°PNëPSPëP¶C†PsPiPs CḡC-PIPSC¶ P†C-Pr
C†P°C† P·P°C†C,PsC†C†P†P°PSPSC¶ P”CrPsPrP°CḡC,C† PjPsP¶Pµ P±CrC,Pë
C†PIC-PrC†PµPSPSC¶Pj PSP°C¶P†PSPsC†C,C- CḡP°PeCr P†PµCḡPµPrPjC-C...CrCḡPsP†PsC—
P·P°P»PsP·Pë P°P±Ps PSPµPrPsC,CḡPëPjP°PSPSC¶ CḡPµP¶PëPjCr P»C-PëC†P†P°PSPSC¶
P”CrPsPrP°CḡC,PsPj C- P†PsC,CḡPµP±CrC” CḡPµC,PµP»CHPSPsPiPs P†PëPIC†PµPSPSC¶,
PSP°PIC-C,CH C¶PeC%Ps P†PsPeP°P·PSPëPePë PSA P·PSP°C...PsPrC¶C,CHC†C¶ Cr
PjPµP¶P°C... PSPsCḡPjPë Cr C†PsP»PsPIC-PëC-PI, C¶PeC- PSPµ P»C-PëC†P†P°P»PëC†CH
C-PSP†C-P±C-C,PsCḡP°PjPë 5α-CḡPµPrCrPeC,P°P·Pë. PµCḡPë C-PSC,PµCḡP†CḡPµC,P°C†C-C—
P†PsPeP°P·PSPëPeC-PI PSA Cr C...P†PsCḡPëC..., C¶PeC- P»C-PëC†CḡC,CHC†C¶
P”CrPsPrP°CḡC,PsPj, C†P»C-Pr P·P†P°P¶P°C,Pë PSP° P†PsP†PµCḡPµPrPSC- P†PsPeP°P·PSPëPePë
PSA PrP»C¶ P†PsCḡC-PIPSC¶PSPSC¶.

P—P°C†C,PsC†C†P†P°PSPSC¶ P”CrPsPrP°CḡC,C† PSPµ P†P†P»PëP†P°C” PSP°
P†PëPePsCḡPëC†C,P°PSPSC¶ CḡC-PIPSC¶ P†CḡPsC†C,P°C,-C†P†PµC†PëC,,C-C†PSPsPiPs
P°PSC,PëP†PµPSC† PrP»C¶ PrC-P°PiPSPsC†C,PëPePë CḡP°PeCr
P†PµCḡPµPrPjC-C...CrCḡPsP†PsC— P·P°P»PsP·Pë P†C-C†P»C¶ PIC†C,P°PSPsP†P»PµPSPSC¶
PNëPsPiPs PSPsP†PsPiPs P†PëC...C-PrPSPsPiPs CḡC-PIPSC¶.

P—P°PiP°P»CHPSPëPNë CḡC-PIPµPSCḡ C†PëCḡPsP†P°C,PePsP†PsP†Ps P†CḡPsC†C,P°C,-
C†P†PµC†PëC,,C-C†PSPsPiPs P°PSC,PëP†PµPSC† P†PsP†PµCḡC,P°C”C,CHC†C¶ PrPs
P†PëC...C-PrPSPsPiPs CḡC-PIPSC¶ P†CḡPsC,C¶PiPsPj 6 PjC-C†C¶C†C-PI P†C-C†P»C¶
P†CḡPëP†PëPSPµPSPSC¶ P»C-PëC†P†P°PSPSC¶.

PŸP†C-PIPIC-PrPSPsC€PµPSPSC¶ P¶ PIC-P»CHPSPsPiPs P†CḡPsC†C,P°C,-
C†P†PµC†PëC,,C-C†PSPsPiPs P°PSC,PëP†PµPSC† C- P·P°PiP°P»CHPSPsPiPs PNëPsPiPs
CḡC-PIPSC¶ P·P°P»PëC€P°C”C,CHC†C¶ C†C,P°P»PëPj PSP°PIC-C,CH P†C-Pr C†P°C†
P»C-PëC†P†P°PSPSC¶ P”CrPsPrP°CḡC,PsPj. P†PeC%Ps PrP»C¶ P†PëP·PSP°C†PµPSPSC¶
CḡP°PeCr P†PµCḡPµPrPjC-C...CrCḡPsP†PsC— P·P°P»PsP·Pë P»C-PëP°Cḡ P†PëCḡC-C€PëC,CH
P†PëPePsCḡPëC†C,P°C,Pë PIC-PrC†PsC,PePsP†PëPNë P†PsPeP°P·PSPëPe PIC-P»CHPSPsPiPs

ΠΐCΉPsCΓC, P°C, -CΓΐΠΐC†PēC,, C-C†PSPsPiPs P°PSC, PēPiPμPSCr CΓ C... PIPsCΉPsPiPs, CΐPePēPN₂
P»C-PeCΓC"C, CHCΓCΐ P"CΓPsPrP°CΉC, PsPj, PSC-CΐPePsPiPs PePsCΉPμPiCΓPIP°PSPSCΐ
ΠΐPsPeP°P·PSPēPeP° PIC-P»CHPSPsPiPs ΠΐCΉPsCΓC, P°C, -CΓΐΠΐC†PēC,, C-C†PSPsPiPs
P°PSC, PēPiPμPSCr ΠΐCΉPsPiPsPrPēC, Pē PSPμ ΠΐPsC, CΉC-P±PSPs.

PμPμCΉPμPr ΠΐPsC†P°C, PePsPj PeCΓCΉCΓCΓ P»C-PeCΓPIP°PSPSCΐ P"CΓPsPrP°CΉC, PsPj C, P°
ΠΐPμCΉC-PsPrPēC†PSPs ΠΐC-Pr C†P°CΓ P»C-PeCΓPIP°PSPSCΐ ΠΐP°C†C-C"PSC, P°Pj P·
PrPsP±CΉPsCΐPeC-CΓPSPsCΉ ΠΐC-ΠΐPμCΉΠΐP»P°P·C-C"CΉ ΠΐPμCΉPμPrPjC-C... CΓCΉPsPiPsC—
P·P°P»PsP·Pē CΓP»C-Pr CΉPsP±PēC, Pē ΠΐP°P»CHC†PμPIPμ CΉPμPeC, P°P»CHPSPμ
PsP±CΓC, PμP¶PμPSPSCΐ, P° C, P°PePsP¶ P·P°CΓC, PsCΓPsPICΓPIP°C, Pē C-PSC€C- PjPμC, PsPrPē
PIPēCΐPIP»PμPSPSCΐ CΉP°PeCΓ ΠΐPμCΉPμPrPjC-C... CΓCΉPsPiPsC— P·P°P»PsP·Pē.

PκPēCΉPePsPIP° PSPμPrPsCΓC, P°C, PSC-CΓC, CH

P>C-PeCΓPIP°PSPSCΐ C... PIPsCΉPēC... P· C, CΐP¶PePsCΉ PSPēCΉPePsPiPsCΉ
PSPμPrPsCΓC, P°C, PSC-CΓC, CΉB (PeP»C-CΉPμPSCΓ PeCΉPμP°C, PēPSC-PSCr < 10 PjP»/C...PI)
ΠΐPsC, CΉC-P±PSPs ΠΐCΉPsPiPsPrPēC, Pē P· PsP±PμCΉPμP¶PSC-CΓC, CΉ, PsCΓPeC-P»CHPePē
C,, P°CΉPjP°PePsPeC-PSPμC, PēPeP° PrCΓC, P°CΓC, PμCΉPēPrCΓ CΓ C, P°PePēC... C... PIPsCΉPēC...
PSPμ PIPēPIC†P°P»P°CΓCΉ.

PϋCΉC, PμCΉC-P°P»CHPSP° ΠΐC-ΠΐPsC, PμPSP·C-Cΐ

PμPsPrC-P±PSPs PrPs C-PSC€PēC... P°P»CHC,, P°1-P°PrCΉPμPSPsP±P»PsPeP°C, PsCΉC-PI,
PsCΉC, PsCΓC, P°C, PēC†PSP° ΠΐC-ΠΐPsC, PμPSP·C-Cΐ PjPsP¶Pμ PIPēPSPēPePSCrC, Pē CΓ
ΠΐP°C†C-C"PSC, C-PI, CΐPeC- P»C-PeCΓPIP°P»PēCΓCΐ C, P°PjCΓCΓP»PsP·PēPSPsPj, C- CΓ
CΉC-PrPeC-CΓPSPēC... PIPēΠΐP°PrPeP°C... PjPsP¶Pμ ΠΐCΉPēP·PIPμCΓC, Pē PrPs CΓPēPSPePsPiPμ.

PμCΉPē ΠΐPμCΉC€PēC... PsP·PSP°PeP°C... PsCΉC, PsCΓC, P°C, PēC†PSPsC— ΠΐC-ΠΐPsC, PμPSP·C-C—
(P·P°ΠΐP°PjPsCΉPsC†PμPSPSCΐ, CΓP»P°P±PeC-CΓC, CΉ) ΠΐP°C†C-C"PSC, C-PI, CΐPeC-
CΉPsP·ΠΐPsC†P°P»Pē P»C-PeCΓPIP°PSPSCΐ P"CΓPsPrP°CΉC, PsPj, CΓP»C-Pr ΠΐPμCΉPμPIPμCΓC, Pē
CΓ ΠΐPsP»PsP¶PμPSPSCΐ CΓPēPrCΐC†Pē P°P±Ps P»PμP¶P°C†Pē, ΠΐPsPePē CΓPēPjΠΐC, PsPjPē PSPμ
PjPēPSCrC, CΉ.

PŸP»C-Pr PrPsC, CΉPēPjCΓPIP°C, PēCΓCΉ PsP±PμCΉPμP¶PSPsCΓC, C- ΠΐCΉPē
CΓCΓPjC-CΓPSPsPjCΓ P·P°CΓC, PsCΓCΓPIP°PSPSC- P°P»CHC,, P°-
P°PrCΉPμPSPsP±P»PsPeP°C, PsCΉC-PI, PIPeP»CΉC†P°CΉC†Pē C, P°PjCΓCΓP»PsP·PēPS, C, P°
C-PSPiC-P±C-C, PsCΉC-PI 5-C,, PsCΓC,, PsPrC-PμCΓC, PμCΉP°P·Pē. PϋP»CHC,, P°-
P°PrCΉPμPSPsP±P»PsPeP°C, PsCΉPē C, P° C-PSPiC-P±C-C, PsCΉPē 5-
C,, PsCΓC,, PsPrC-PμCΓC, PμCΉP°P·Pē C" PIP°P·PsPrPēP»P°C, P°C, PsCΉP°PjPē C, P° PjPsP¶CΓC, CΉ
P·PSPēP¶CΓPIP°C, Pē P°CΉC, PμCΉC-P°P»CHPSPēPN₂ C, PēCΓPe. PŸCΓPjC-CΓPSPμ
P·P°CΓC, PsCΓCΓPIP°PSPSCΐ C†PēC... PrPIPsC... PeP»P°CΓC-PI P»C-PeP°CΉCΓCΉPePēC...
P·P°CΓPsP±C-PI PjPsP¶Pμ ΠΐPsC, PμPSC†C-PN₂PSPs CΓΠΐCΉPēC†PēPSPēC, Pē
CΓPēPjΠΐC, PsPjP°C, PēC†PSCr P°CΉC, PμCΉC-P°P»CHPSCr ΠΐC-ΠΐPsC, PμPSP·C-CΉ (PrPēPI.
CΉPsP·PrC-P» B<P·P·P°C"PjPsPrC-Cΐ P· C-PSC€PēPjPē P»C-PeP°CΉCΓCΉPePēPjPē
P·P°CΓPsP±P°PjPē C, P° C-PSC€C- PIPēPrPē PIP·P°C"PjPsPrC-PN₂B»).

P†PSC, CΉP°PsPiPμCΉP°C†C-PN₂PSPēPN₂ CΓPēPSPrCΉPsPj P°C, PsPSC-C†PSPsC—
CΉP°PN₂PrCΓP¶PePē

PμC-Pr C†P°CΓ PsPiPμCΉP°C†C-C— P· ΠΐCΉPēPIPsPrCΓ PeP°C, P°CΉP°PeC, Pē C, P°
ΠΐP»P°CΓPePsPjPē CΓ PrPμCΐPePēC... ΠΐP°C†C-C"PSC, C-PI, CΐPeC- CΉP°PSC-C€Pμ
PsC, CΉPēPjCΓPIP°P»Pē C, P°PjCΓCΓP»PsP·PēPS, P±CΓPI PIC-PrP·PSP°C†PμPSPēPN₂

C-PSC, C₇H₆P°PsPīPμC₇H₆P°C†C-PM₂PSPēPM₂ C†PēPSPrC₇H₆PsPj P°C, PsPSC-C†PSPsC—
C₇H₆P°PM₂PrC†P¶PēPē (P†PŸP₇P, PIP°C₇H₆-P°PSC, C†PēPSPrC₇H₆PsPjC† PIC†P·CH₂PePsC—
P·C-PSPēC†C-). P†PSC, C₇H₆P°PsPīPμC₇H₆P°C†C-PM₂PSPēPM₂ C†PēPSPrC₇H₆PsPj P°C, PsPSC-C†PSPsC—
C₇H₆P°PM₂PrC†P¶PēPē PjPsP¶Pμ PīC₇H₆PēP·PIPμC†C, Pē PrPs P·P±C-P»CH₂CēPμPSPSC¶
C₇H₆PēP·PēPeC† PsC†PSPēC... C†C†PēP»P°PrPSPμPSC₇ PīC-Pr C†P°C† P°P±Ps PīC-C†P»C¶
PsPīPμC₇H₆P°C†C-C—. PŸPsPjC† P»C-PeC†PIP°PSPSC¶ P°C†PsPrP°C₇H₆, PsPj PSPμ
C₇H₆PePsPjPμPSPrC†C°C, CH₂C†C¶ PīP°C†C-C°PSC, P°Pj, C¶PePēPj P·P°PīP»P°PSPsPIP°PSPs
PsPīPμC₇H₆P°C†C-C₇ P· PīC₇H₆PēIPsPrC† PeP°C, P°C₇H₆P°PeC, Pē.

PμC-Pr C†P°C† PīPμC₇H₆PμPrPsPīPμC₇H₆P°C†C-PM₂PSPsPiPs PsP±C†C, PμP¶PμPSPSC¶
C...C-C₇H₆C†C₇Pi-PsC,, C, P°P»CH₂PjPsP»PsPi C- PM₂PsPiPs P±C₇H₆PēPiP°PrP° PjP°C₇H₆, CH₂
P·B₇TM C¶C†C†PIP°C, Pē, C†Pē PīC₇H₆PēP·PSP°C†P°P»Pē PīP°C†C-C°PSC, PsPIC- C₇H₆P°PSC-CēPμ
P°P±Ps C, PμPīPμC₇ P°C†PsPrP°C₇H₆,. PīPμ PrP°C†C, CH₂ P·PjPsPiC† PīPμC₇H₆PμPrP±P°C†PēC, Pē
PjPsP¶P»PēPIC† PīPsC¶PIC† C-PSC, C₇H₆P°PsPīPμC₇H₆P°C†C-PM₂PSPsPiPs C†PēPSPrC₇H₆PsPjC†
P°C, PsPSC-C†PSPsC— C₇H₆P°PM₂PrC†P¶PēPē PīC-Pr C†P°C† PsPīPμC₇H₆P°C†C-C—.

PhC, C₇H₆PēPjP°PSPs PīPsPsPrPēPSPsPeC- PīPsPIC-PrPsPjP»PμPSPSC¶ PīC₇H₆Ps
PīPsP·PēC, PēPIPSPēPM₂ PμC,, PμPeC, PIC-PrPjC-PSPē C, P°PjC†C†P»PsP·PēPSC† P·P° 1B₇"2
C, PēP¶PSC- PrPs PsPīPμC₇H₆P°C†C-C— P· PīC₇H₆PēIPsPrC† PeP°C, P°C₇H₆P°PeC, Pē C, P°
PīP»P°C†PePsPjPē, PsPrPSP°Pe PīPμC₇H₆PμPIP°PiPē C, P° C†C, C₇H₆PsPePē PīC₇H₆PēPīPēPSPμPSPSC¶
P»C-PeC†PIP°PSPSC¶ PrPs PsPīPμC₇H₆P°C†C-C— P· PīC₇H₆PēIPsPrC† PeP°C, P°C₇H₆P°PeC, Pē C, P°
PīP»P°C†PePsPjPē PSPμ PIC†C, P°PSPsPIP»PμPSC-.

PkPμPiPμC₇H₆PjPμC, PēC†PSC- PeP°PīC†C†P»Pē

P°C†C, P°C†C, PμC₇H₆PēPr P°P±C†PsC₇H₆P±C†C°C, CH₂C†C¶ C†PμC₇H₆Pμ· CēPeC-C₇H₆C†, C, PsPjC†
P¶C-PSPēPē C, P° PrC-C, Pē PjP°C₇H₆, CH₂ C†PSPēPeP°C, Pē PePsPSC, P°PeC, C† P·
PSPμPiPμC₇H₆PjPμC, PēC†PSPēPjPē PeP°PīC†C†P»P°PjPē. PīPeC%Ps C₇H₆-PrPēPSP° P·
PeP°PīC†C†P»Pē PīPsC, C₇H₆P°PīPēP»P° PSP° CēPeC-C₇H₆C†, C—C— C†P»C-Pr PSPμPiP°PM₂PSPs
P·PjPēC, Pē PIPsPrPsC₇ P· PjPēP»PsPj.

P†PSPiC-P±C-C, PsC₇H₆Pē PŸYP3A4 C, P° CYP2D6

PŸC†PjC-C†PSPμ P·P°C†C, PsC†C†PIP°PSPSC¶ C, P°PjC†C†P»PsP·PēPSC†
PīC-PrC₇H₆PsC...P»PsC₇H₆PēPrC† P· C†PēP»CH₂PSPēPjPē C-PSPiC-P±C-C, PsC₇H₆P°PjPē PŸYP3A4
(PSP°PīC₇H₆PēPeP»P°Pr PePμC, PsPePsPSP°P·PsP»PsPj) P°P±Ps B₇" C† PjPμPSC€C-PM₂ PjC-C₇H₆-
B₇" P· C†PēP»CH₂PSPēPjPē C-PSPiC-P±C-C, PsC₇H₆P°PjPē CYP2D6 (PSP°PīC₇H₆PēPeP»P°Pr
PīP°C₇H₆PsPeC†PμC, PēPSPsPj) PjPsP¶Pμ P·P±C-P»CH₂CēC†PIP°C, Pē PePsPSC†PμPSC, C₇H₆P°C†C-C₇
C, P°PjC†C†P»PsP·PēPSC† (PrPēPI. C₇H₆PsP·PrC-P» B«P·P·P°C°PjPsPrC-C¶ P· C-PSCēPēPjPē
P»C-PeP°C₇H₆C†CH₂PePēPjPē P·P°C†PsP±P°PjPē C, P° C-PSC€C- PIPēPrPē PIP·P°C°PjPsPrC-PM₂B»).
PŸPsPjC† PSPμ C₇H₆PePsPjPμPSPrC†C°C, CH₂C†C¶ P·P°C†C, PsC†PsPIC†PIP°C, Pē
C, P°PjC†C†P»PsP·PēPS PīP°C†C-C°PSC, P°Pj, C¶PeC- P»C-PeC†C₇H₆, CH₂C†C¶
C†PēP»CH₂PSPēPjPē C-PSPiC-P±C-C, PsC₇H₆P°PjPē PŸYP3A4, C, P° C₇H₆PePsPjPμPSPrPsPIP°PSPs
P· PsP±PμC₇H₆PμP¶PSC-C†C, C₇ P·P°C†C, PsC†PsPIC†PIP°C, Pē C, P°PjC†C†P»PsP·PēPS
PīP°C†C-C°PSC, P°Pj, C¶PeC- P»C-PeC†C₇H₆, CH₂C†C¶ PīPsPjC-C₇H₆PSPēPjPē
C-PSPiC-P±C-C, PsC₇H₆P°PjPē PŸYP3A4 (PSP°PīC₇H₆PēPeP»P°Pr PμC₇H₆PēC, C₇H₆PsPjC-C†PēPSPsPj),
C†PēP»CH₂PSPēPjPē P°P±Ps PīPsPjC-C₇H₆PSPēPjPē C-PSPiC-P±C-C, PsC₇H₆P°PjPē CYP2D6,
PePsPjP±C-PSP°C†C-C°C₇ PsP±PsC... PŸYP3A4 C, P° CYP2D6 C-PSPiC-P±C-C, PsC₇H₆-PI P°P±Ps
PīP°C†C-C°PSC, P°Pj P·C- C†P»P°P±PePēPj PjPμC, P°P±PsP»C-P·PjPsPj CYP2D6.

PμPμC†C-PSPePsPIP° PSPμPrPsC†C, P°C, PSC-C†C, CH₂

P'P'IP»PëPI P'PµC‡C-PSPePsPIPsC— PSPµPrPsCf'c,P°c,PSPsCf'c,C- PSP°
C,,P°Cf'cPjP°PePsPeC-PSPµC,PëPeCf'cB PrCf'c,P°Cf'c,PµCf'cPëPrCf'c PSPµ PIPëPIC‡P°PICf'cCf'c.
PšPµCf'cPµP· P°PeC,PëPIPSpëPNë PjPµC,P°P±PsP»C-P·Pj PrCf'c,P°Cf'c,PµCf'cPëPrCf'c C,P° 3Bf'c"5-
C,Pë C,PëP¶PSPµPIPëPNë P'PµCf'cC-PsPr PNëPsPiPs PSP°P'c-PIPIPëPIPµPrPµPSPSCf'c
P»C-PeCf'cPIP°PSPSCf'c PrCf'c,P°Cf'c,PµCf'cPëPrPsPj C...PIPsCf'cPëC... P· P»PµPiPePsCf'c P°P±Ps
P'PsPjC-Cf'cPSPsCf'c P'PµC‡C-PSPePsPIPsCf'c PSPµPrPsCf'c,P°c,PSC-Cf'c,Cf'c Cf'cP»C-Pr
P'cCf'cPsPIPsPrPëC,Pë P· PsP±PµCf'cPµP¶PSC-Cf'c,Cf'c (PrPëPI. Cf'cPsP·PrC-P»Pë B«P'ŸPiPsCf'cC-P±
P·P°Cf'c,PsCf'cCf'cPIP°PSPSCf'c C,P° PrPsP·PëB», B«PµCf'cPsC,PëP'PsPeP°P·P°PSPSCf'cB»,
B«PαP°Cf'cPjP°PePsP»PsPiC-C‡PSC- PIP»P°Cf'c,PëPIPsCf'c,C-B»).

P"PsPiPsPjC-P¶PSC- Cf'cPµC‡PsPIPëPSPë

P"Cr'PsPrP°Cf'c, PjC-Cf'c,PëC,Cf'c P±P°Cf'cPIPSpëPe FD&C Yellow 6 (E 110), C%oPs PjPsP¶Pµ
Cf'cPiCf'cPëC‡PëPSPëC,Pë P°P»PµCf'cPiC-C‡PSC- Cf'cPµP°PeC‡C-C—.

P P°Pe PiCf'cCf'cPrPSPsC— P·P°P»PsP·Pë Cr C‡PsP»PsPIC-PeC-PI

PµPsPIC-PrPsPjP»Cf'cP»PsCf'cCf'c P'cCf'cPs Cf'cC-PrPeC-Cf'cPSC- PIPëPiP°PrPePë Cf'cP°PeCf'c
PiCf'cCf'cPrPSPsC— P·P°P»PsP·Pë Cr C‡PsP»PsPIC-PeC-PI P'c-Pr C‡P°Cf'c PeP»C-PSC-C‡PSPëC...
PrPsCf'cP»C-PrP¶PµPSCf'c C,P° Cr P'PsCf'c,PjP°Cf'cPePµC,PëPSPiPsPIPëPNë P'PµCf'cC-PsPr. PµCf'cPë
C‡Cf'cPsPjCf'c PµP'c-PrPµPjC-PsP»PsPiC-C‡PSC- PrPsCf'cP»C-PrP¶PµPSPSCf'c PIPeP°P·Cf'cCf'c,Cf'c
PSP° PIC-PrCf'cCf'c,PSC-Cf'c,Cf'c P'c-PrPIPëC%oPµPSPSCf'c Cf'cPëP·PëPeCf'c Cf'cPsP·PIPëC,PeCf'c
Cf'cP°PeCf'c PiCf'cCf'cPrPSPsC— P·P°P»PsP·Pë Cr C‡PsP»PsPIC-PeC-PI P'cCf'cPë
P·P°Cf'c,PsCf'cCf'cPIP°PSPSC- C-PSPiC-P±C-C,PsCf'cC-PI 5α-Cf'cPµPrCf'cPeC,P°P·Pë. P»C-PeP°Cf'cC-
P'PsPIPëPSPSC- P'PsPiPµCf'cPµPrPëC,Pë Cf'cPIPsC—C... P'P°C‡C-C"PSC,C-PI P'cCf'cPs
PSPµPsP±C...C-PrPSC-Cf'c,Cf'c PSPµPiP°PNëPSPs P'PsPIC-PrPsPjP»Cf'cC,Pë P'cCf'cPs P±Cf'cPrCf'c-
Cf'cPeC- P·PjC-PSPë Cr C,PeP°PSPëPSC- PiCf'cCf'cPrPSPsC— P·P°P»PsP·Pë, PSP°P'cCf'cPëPeP»P°Pr
P'cCf'cPs PIPëPrC-P»PµPSPSCf'c C-P· Cf'cPsCf'cPeP° P°P±Ps P'cCf'cPëP'cCf'c...P»C-Cf'c,Cf'c.

***P—P°Cf'c,PsCf'cCf'cPIP°PSPSCf'c C‡ P'PµCf'cC-PsPr PIP°PiC-C,PSPsCf'c,C- P°P±Ps
PiPsPrC‡PIP°PSPSCf'c PiCf'cC‡PrPrCf'c.***

P"Cr'PsPrP°Cf'c, P'cCf'cPsC,PëP'PsPeP°P·P°PSPëPNë PrP»Cf'c P»C-PeCf'cPIP°PSPSCf'c P¶C-PSPsPe.
P"PsCf'cP»C-PrP¶PµPSPSCf'c P· PIPëPIC‡PµPSPSCf'c PIPiP»PëPICf'c P"Cr'PsPrP°Cf'c,Cf'c PSP°
PIP°PiC-C,PSC-Cf'c,Cf'c, P»P°PeC,P°C‡C-Cf'c C,P° C,,PµCf'c,PëP»Cf'cPSC-Cf'c,Cf'c PSPµ
P'cCf'cPsPIPsPrPëP»Pë. P"°P»C- PSP°PrP°C"C,Cf'cCf'cCf'c C-PSC,PsCf'cPjP°C‡C-Cf'c P'cCf'cPs
P·P°Cf'c,PsCf'cCf'cPIP°PSPSCf'c PePsP¶PSPsPiPs PePsPjPiPsPSPµPSC,P° PsPeCf'cPµPjPs.

PαPµCf'cC,PëP»Cf'cPSC-Cf'c,Cf'c.

P"Cr'c,P°Cf'c,PµCf'cPëPr PIPiP»PëPIP°C" PSP° C...P°Cf'cP°PeC,PµCf'cPëCf'c,PëPePë
PµCf'cPeCf'cP»Cf'cCf'c,Cf'c (P·PjPµPSC€PµPSPSCf'c PeC-P»Cf'cPePsCf'c,C-
Cf'cPiPµCf'cPjP°C,PsP·PsC—PrC-PI, PsP±Bf'c™C"PjCf'c PµCf'cPeCf'cP»Cf'cCf'c,Cf'c C,P°
Cf'cCf'cC...PsPjPsCf'c,C- Cf'cPiPµCf'cPjP°C,PsP·PsC—PrC-PI). PκPµ PIPëPeP»Cf'cC‡PµPSPëPNë
Cf'cPëP·PëPe P·PSPëP¶PµPSPSCf'c C‡PsP»PsPIC-C‡PsC— C,,PµCf'cC,PëP»Cf'cPSPsCf'c,C-.

PκC‡C-PSPeP° PIPiP»PëPICf'c C,P°PjCf'cCf'cP»PsP·PëPSCf'c PiC-PrCf'cPsC...P»PsCf'cPëPrCf'c PSP°
PeC-P»Cf'cPeC-Cf'c,Cf'c P°P±Ps C,,Cf'cPSPeC‡C-Cf'c Cf'cPiPµCf'cPjP°C,PsP·PsC—PrC-PI PSPµ
P'cCf'cPsPIPsPrPëP»P°Cf'cCf'c.

P°P°PiC-C,PSC-CÍC,CH.

PİPe C- C-PSC€C- C-PSPiC-P±C-C,PsCȚPë 5α-CȚPμPrCřPeC,P°P·Pë, PrCřC,P°CÍC,PμCȚPëPr
PİPμCȚPμC€PePsPrP¶P°C” PİPμCȚPμC,PIPsCȚPμPSPSCȚ C,PμCÍC,PsCÍC,PμCȚPsPSCř Cř
PrPëPiC-PrCȚPsC,PμCÍC,PsCÍC,PμCȚPsPS, C°Ps PjPsP¶Pμ PiP°P»CHPjCřPIP°C,Pë
CȚPsP·PIPëC,PsPe P·PsPIPSC-C€PSC-C... CÍC,P°C,PμPIPëC... PsCȚPiP°PSC-PI Cř PİP»PsPrP°
C±PsP»PsPIC-C±PsC— CÍC,P°C,C-. PkPμP·PSP°C±PSCř PeC-P»CHPeC-CÍC,CH
PrCřC,P°CÍC,PμCȚPëPrCř P±CřP»Ps PIPëCȚPIIP»PμPSPs PI PμCȚPeCřP»CȚC,C- PİCȚPë
PİCȚPsPIPμPrPμPSPSC- PrPsCÍP»C-PrP¶PμPSPSCȚ. PkPμPIC-PrPsPjPs, C±Pë PIPİP»PëPIP°C”
PrCřC,P°CÍC,PμCȚPëPr, C°Ps PİPsC,CȚP°PİPëPI PrPs PsCȚPiP°PSC-P·PjCř P¶C-PSPePë P·
CÍC-PjBȚ™ CȚPj C±PsP»PsPIC-PeP°, CȚPePëPNë P»C-PeCřC”C,CHCÍCȚ P”CřPsPrP°CȚC,PsPj,
PSP° PİP»C-Pr C±PsP»PsPIC-C±PsC— CÍC,P°C,C-.

PİPe C- Cř CȚP°P·C- P·P°CÍC,PsCÍCřPIP°PSPSCȚ C-PSC€PëC... C-PSPiC-P±C-C,PsCȚC-PI 5α-
CȚPμPrCřPeC,P°P·Pë, CȚPμPePsPjPμPSPřCřC”C,CHCÍCȚ PePsCȚPëCÍC,CřPIP°C,PëCÍCH
PİCȚPμP·PμCȚPIP°C,PëPIPsPj PiC-Pr C±P°CÍ CÍC,P°C,PμPIPsPiPs P°PeC,Cř, CȚPeC°Ps
P¶C-PSPeP° PIP°PiC-C,PSP°, P° C±PsP»PsPIC-Pe P»C-PeCřC”C,CHCÍCȚ P”CřPsPrP°CȚC,PsPj, P·
PjPμC,PsCȚ P·P°PİPsP±C-PİP°PSPSCȚ PİPsC,CȚP°PİP»CȚPSPSCȚ CÍC-PjBȚ™ CȚ PrPs
PsCȚPiP°PSC-P·PjCř P¶C-PSPePë.

PkPμPjP°C” PrPsPeP°P·C-PI C,PsPiPs, C°Ps PİPIPμPrPμPSPSCȚ C,P°PjCÍCřP»PsP·PëPSCř
PiC-PrCȚPsC...P»PsCȚPëPrCř PIP°PiC-C,PSPëPj CÍP°PjPeP°Pj C°CřCȚC-PI C,P° PeCȚPsP»C-PI
Cř PrPsP·P°C..., C°Ps PİPμCȚPμPIPëC°CřCȚC,CH C,PμCȚP°PİPμPIC,PëC±PSC-,
PSPμPiP°C,PëPIPsPs PIPİP»PëPIP°C” PSP° PİP»C-Pr.

P”PsPrCřPIP°PSPSCȚ PiCȚCřPrPrCȚ.

PkPμPIC-PrPsPjPs, C±Pë PİCȚPsPSPëPeP°CȚC,CH PrCřC,P°CÍC,PμCȚPëPr C,P°
C,P°PjCÍCřP»PsP·PëPS Cř PiCȚCřPrPSPμ PjPsP»PsPePs P¶C-PSPePë.

***P—PᵣP°C,PSC-CÍC,CH PIPİP»PëPIP°C,Pë
PSP° C€PIPëPᵣPεC-CÍC,CH
CȚPμP°PeC†C-C— PİCȚPë
PePμCȚC¿PIP°PSPSC-
P°PIC,PsC,CȚP°PSCÍPİPsCȚC,PsPj P°P±Ps
C-PSC€PëPjPë PjPμC...P°PSC-P·PjP°PjPë.***

P”PsCÍP»C-PrP¶PμPSPSCȚ P· PIPëPIC±PμPSPSCȚ PIPİP»PëPICř P”CřPsPrP°CȚC,Cř PSP°
P·PrP°C,PSC-CÍC,CH PePμCȚCřPIP°C,Pë P°PIC,PsC,CȚP°PSCÍPİPsCȚC,PsPj P°P±Ps
C-PSC€PëPjPë PjPμC...P°PSC-P·PjP°PjPë PSPμ PİCȚPsPIPsPrPëP»PëCÍCȚ. PkPrPSP°Pe
PİP°C†C-C”PSC,C-PI PİPsC,CȚC-P±PSPs PİCȚPsC-PSC,,PsCȚPjCřPIP°C,Pë PİCȚPs
PjPsP¶P»PëPIPμ PIPëPSPëPePSPμPSPSCȚ CÍPëPjPİC,PsPjC-PI, PİPsPIBȚ™ CȚP·P°PSPëC... P·
PsCȚC,PsCÍC,P°C,PëC±PSPsCȚ P°CȚC,PμCȚC-P°P»CHPSPsCȚ PiC-PİPsC,PμPSP·C-C”CȚ,

$P \cdot PsPeC_{\overline{H}}P_{\overline{u}}P_jP^\circ$ $\overline{P}iC_{\overline{H}}Ps$ $P \cdot P^\circ \overline{P}iP^\circ P_jPsC_{\overline{H}}PsC_{\overline{+}}P_{\overline{u}}PSPSC_{\overline{u}}$, $\overline{P}iC_{\overline{H}}P_{\overline{e}} P \cdot P^\circ C_{\overline{f}}C$, $PsC_{\overline{f}}C_{\overline{f}}PIP^\circ PSPSC-$
 $P''C_{\overline{f}}PsPrP^\circ C_{\overline{H}}C$, $C_{\overline{f}}$.

$\overline{P}Y\overline{P}iPsC_{\overline{f}}C-P_{\pm} P \cdot P^\circ C_{\overline{f}}C$, $PsC_{\overline{f}}C_{\overline{f}}PIP^\circ PSPSC_{\overline{u}}$ C , $P^\circ PrPsP \cdot P_{\overline{e}}$.

$P''PsC_{\overline{H}}PsC_{\overline{f}}P_{\overline{u}}C-$ ($PIPeP_{\overline{u}}C_{\overline{H}}C_{\overline{+}}P^\circ C_{\overline{H}}C_{\overline{+}}P_{\overline{e}} C \dots PIPsC_{\overline{H}}P_{\overline{e}}C \dots P_{\overline{u}}C-C$, $PsC_{\overline{H}}PsPiPs PIC-PeC_{\overline{f}}$)

$P P_{\overline{u}}PePsP_jP_{\overline{u}}PSPPrPsPIP^\circ PSP^\circ PrPsP \cdot P^\circ P''C_{\overline{f}}PsPrP^\circ C_{\overline{H}}C$, $C_{\overline{f}}$ $\overline{b}_{\overline{H}}''$ $1 PeP^\circ \overline{P}iC_{\overline{f}}C_{\overline{f}}P_{\overline{u}}P^\circ$ ($0,5 P_jPi/0,4 P_jPi$) $PSP^\circ PrPsP_{\pm}C_{\overline{f}}$. $P_{\overline{u}}C_{\overline{H}}P_{\overline{u}}\overline{P}iP^\circ C_{\overline{H}}P^\circ C$, $\overline{P}iC_{\overline{H}}P_{\overline{e}}PN_{\overline{e}}P_jP^\circ C_{\overline{H}}C$, CH $\overline{P}iP_{\overline{u}}C_{\overline{H}}PsC_{\overline{H}}P^\circ P_{\overline{u}}CHPsPs$
 $C_{\overline{+}}P_{\overline{u}}C_{\overline{H}}P_{\overline{u}}P \cdot 30 C \dots PIP_{\overline{e}}P_{\overline{u}}P_{\overline{e}}PS \overline{P}iC-C_{\overline{f}}P_{\overline{u}}C_{\overline{u}}$ $C_{\overline{f}}\overline{P}iPsP_{\overline{u}}\overline{P}iPIP^\circ PSPSC_{\overline{u}}$ $C-P_{\overline{u}}C-$ $PI PsPrP_{\overline{e}}PS C-$
 C , $PsPN_{\overline{e}} P_{\overline{u}}P_{\overline{u}} C_{\overline{+}}P^\circ C_{\overline{f}}$. $P_{\overline{u}}P^\circ \overline{P}iC_{\overline{f}}C_{\overline{f}}P_{\overline{u}}C_{\overline{f}} C_{\overline{f}}P_{\overline{u}}C-P_r PePsPIC$, $P^\circ C$, $P_{\overline{e}} C_{\overline{+}}C-P_{\overline{u}}PsC_{\overline{H}}$, $PSP_{\overline{u}}$
 $PIC-PrPeC_{\overline{H}}P_{\overline{e}}PIP^\circ C$, $P_{\overline{e}} C$, $P^\circ PSP_{\overline{u}}$ $C_{\overline{H}}PsP \cdot P_{\overline{u}}PsPIC_{\overline{f}}PIP^\circ C$, $P_{\overline{e}}$, $PsC_{\overline{f}}PeC-P_{\overline{u}}CHPeP_{\overline{e}} \overline{P}iC_{\overline{H}}P_{\overline{e}}$
 $PePsPSC$, $P^\circ PeC$, $C-$ $C-P$. $PIP_jC-C_{\overline{f}}C$, $PsP_j PeP^\circ \overline{P}iC_{\overline{f}}C_{\overline{f}}P_{\overline{u}}P_{\overline{e}} P_jPsP_{\overline{u}}P_{\overline{u}}P_{\overline{e}}PIP_{\overline{u}}$
 $\overline{P}iPsPrC_{\overline{H}}P^\circ P \cdot PSP_{\overline{u}}PSPSC_{\overline{u}}$ $C_{\overline{f}}P_{\overline{u}}P_{\overline{e}}P \cdot PsPIP_sC-$ $PsP_{\pm}PsP_{\overline{u}}PsPSPeP_{\overline{e}} C_{\overline{H}}PsC$, $P^\circ C$, $P^\circ PiP_{\overline{u}}PsC$, $PeP_{\overline{e}}$.

$P''C_{\overline{f}}PsPrP^\circ C_{\overline{H}}C$, $P_jPsP_{\overline{u}}PSP^\circ P \cdot P^\circ C_{\overline{f}}C$, $PsC_{\overline{f}}PsPIC_{\overline{f}}PIP^\circ C$, $P_{\overline{e}} PrP_{\overline{u}}C_{\overline{u}}$ $P \cdot P^\circ P_jC-C_{\overline{u}}P_{\overline{u}}PSPSC_{\overline{u}}$
 $PePsP_jP_{\pm}C-PSPsPIP^\circ PSPsC-$ C , $P_{\overline{u}}C_{\overline{H}}P^\circ \overline{P}iC-C-$ $PrC_{\overline{f}}C$, $P^\circ C_{\overline{f}}C$, $P_{\overline{u}}C_{\overline{H}}P_{\overline{e}}PrPsP_j C$, P°
 C , $P^\circ P_jC_{\overline{f}}C_{\overline{f}}P_{\overline{u}}PsP \cdot P_{\overline{e}}PSC_{\overline{f}}$ $PiC-PrC_{\overline{H}}PsC \dots P_{\overline{u}}PsC_{\overline{H}}P_{\overline{e}}PrPsP_j P \cdot P_jP_{\overline{u}}C$, $PsC_{\overline{H}}$ $\overline{P}iPsP_{\overline{u}}P_{\overline{u}}PiC_{\overline{e}}P_{\overline{u}}PSPSC_{\overline{u}}$
 $P_{\overline{u}}C-PeC_{\overline{f}}PIP^\circ PSPSC_{\overline{u}}$.

$P-P^\circ P_jC-PSP^\circ P''C_{\overline{f}}PsPrP^\circ C_{\overline{H}}C$, $PsP_jB PrC_{\overline{f}}C$, $P^\circ C_{\overline{f}}C$, $P_{\overline{u}}C_{\overline{H}}P_{\overline{e}}PrC_{\overline{f}} P^\circ P_{\pm}Ps$
 C , $P^\circ P_jC_{\overline{f}}C_{\overline{f}}P_{\overline{u}}PsP \cdot P_{\overline{e}}PSC_{\overline{f}}$ $PiC-PrC_{\overline{H}}PsC \dots P_{\overline{u}}PsC_{\overline{H}}P_{\overline{e}}PrC_{\overline{f}} \overline{P}iC_{\overline{H}}P_{\overline{e}} P_jPsPSPsC$, $P_{\overline{u}}C_{\overline{H}}P^\circ \overline{P}iC-C-$
 $P_jPsP_{\overline{u}}P_{\overline{u}}P_{\overline{e}}PIP^\circ$, $C_{\overline{u}}PeC_{\overline{u}}Ps C_{\overline{+}}P_{\overline{u}} PeP_{\overline{u}}C-PSC-C_{\overline{+}}PSPs PsP_{\pm}T'C_{\overline{H}}C_{\overline{f}}PSC$, $PsPIP^\circ PSPs$.

$\overline{P}K_{\overline{e}}C_{\overline{H}}PePsPIP^\circ PSP_{\overline{u}}PrPsC_{\overline{f}}C$, $P^\circ C$, $PSC-C_{\overline{f}}C$, CH

$P_{\overline{u}}P^\circ C_{\overline{H}}P_jP^\circ PePsPeC-PSP_{\overline{u}}C$, $P_{\overline{e}}PeP^\circ P''C_{\overline{f}}PsPrP^\circ C_{\overline{H}}C$, $C_{\overline{f}}$ $C_{\overline{f}}$ $C \dots PIPsC_{\overline{H}}P_{\overline{e}}C \dots P \cdot PSP_{\overline{e}}C_{\overline{H}}PePsPIP_sC_{\overline{H}}$
 $PSP_{\overline{u}}PrPsC_{\overline{f}}C$, $P^\circ C$, $PSC-C_{\overline{f}}C$, CH $PSP_{\overline{u}}$ $PIP_{\overline{e}}PIC_{\overline{+}}P^\circ P_{\overline{u}}P^\circ C_{\overline{f}}CH$. $P-P_jC-PSC_{\overline{H}}PIP^\circ C$, $P_{\overline{e}} PrPsP \cdot C_{\overline{f}}$
 $\overline{P}iC_{\overline{H}}P_{\overline{u}}\overline{P}iP^\circ C_{\overline{H}}P^\circ C$, $C_{\overline{f}}P_{\overline{u}}C_{\overline{u}}$ $P_{\overline{u}}C-PeC_{\overline{f}}PIP^\circ PSPSC_{\overline{u}}$ C , $P^\circ PeP_{\overline{e}}C \dots C \dots PIPsC_{\overline{H}}P_{\overline{e}}C \dots PSP_{\overline{u}}$
 $\overline{P}iPsC$, $C_{\overline{H}}C-P_{\pm}PSPs$ ($PrP_{\overline{e}}PI$. $C_{\overline{H}}PsP \cdot PrC-P_{\overline{u}}$ $B \ll PhC_{\overline{f}}PsP_{\pm}P_{\overline{u}}P_{\overline{e}}PIP_sC_{\overline{f}}C$, $C-$
 $P \cdot P^\circ C_{\overline{f}}C$, $PsC_{\overline{f}}C_{\overline{f}}PIP^\circ PSPSC_{\overline{u}}B \gg C$, $P^\circ B \ll P_{\overline{u}}P^\circ C_{\overline{H}}P_jP^\circ PePsPeC-PSP_{\overline{u}}C$, $P_{\overline{e}}PeP^\circ B \gg$).

$\overline{P}_{\overline{u}}P_{\overline{u}}C_{\overline{+}}C-PSPePsPIP^\circ PSP_{\overline{u}}PrPsC_{\overline{f}}C$, $P^\circ C$, $PSC-C_{\overline{f}}C$, CH

$P_{\overline{u}}P^\circ C_{\overline{H}}P_jP^\circ PePsPeC-PSP_{\overline{u}}C$, $P_{\overline{e}}PeP^\circ P''C_{\overline{f}}PsPrP^\circ C_{\overline{H}}C$, $C_{\overline{f}}$ $C_{\overline{f}}$ $C \dots PIPsC_{\overline{H}}P_{\overline{e}}C \dots P \cdot$
 $\overline{P}iP_{\overline{u}}C_{\overline{+}}C-PSPePsPIP_sC_{\overline{H}}$ $PSP_{\overline{u}}PrPsC_{\overline{f}}C$, $P^\circ C$, $PSC-C_{\overline{f}}C$, CH $PSP_{\overline{u}}$ $PIP_{\overline{e}}PIC_{\overline{+}}P^\circ P_{\overline{u}}P^\circ C_{\overline{f}}CH$, C , $PsP_jC_{\overline{f}}$
 $\overline{P}iC_{\overline{H}}P_{\overline{u}}\overline{P}iP^\circ C_{\overline{H}}P^\circ C$, $C_{\overline{f}}P_{\overline{u}}C_{\overline{u}}$ $P_{\overline{u}}C-PeC_{\overline{f}}PIP^\circ PSPSC_{\overline{u}}$ C , $P^\circ PeP_{\overline{e}}C \dots C \dots PIPsC_{\overline{H}}P_{\overline{e}}C \dots PSP_{\overline{u}}$
 $\overline{P}iPsC$, $C_{\overline{H}}C-P_{\pm}PSPs$ ($PrP_{\overline{e}}PI$. $C_{\overline{H}}PsP \cdot PrC-P_{\overline{u}}$ $B \ll PhC_{\overline{f}}PsP_{\pm}P_{\overline{u}}P_{\overline{e}}PIP_sC_{\overline{f}}C$, $C-$
 $P \cdot P^\circ C_{\overline{f}}C$, $PsC_{\overline{f}}C_{\overline{f}}PIP^\circ PSPSC_{\overline{u}}B \gg C$, $P^\circ B \ll P_{\overline{u}}P^\circ C_{\overline{H}}P_jP^\circ PePsPeC-PSP_{\overline{u}}C$, $P_{\overline{e}}PeP^\circ B \gg$). $\overline{P}iPIP_sC_{\overline{H}}P_{\overline{e}}P_j P \cdot$
 C , $C_{\overline{u}}P_{\overline{u}}PePsC_{\overline{H}}$ $\overline{P}iP_{\overline{u}}C_{\overline{+}}C-PSPePsPIP_sC_{\overline{H}}$ $PSP_{\overline{u}}PrPsC_{\overline{f}}C$, $P^\circ C$, $PSC-C_{\overline{f}}C$, CH $\overline{P}iC_{\overline{H}}P_{\overline{u}}\overline{P}iP^\circ C_{\overline{H}}P^\circ C$,
 $\overline{P}iC_{\overline{H}}PsC$, $P_{\overline{e}}\overline{P}iPsPeP^\circ P \cdot P^\circ PSP_{\overline{e}}PN_{\overline{e}}$ ($PrP_{\overline{e}}PI$. $C_{\overline{H}}PsP \cdot PrC-P_{\overline{u}}$ $B \ll P_{\overline{u}}C_{\overline{H}}PsC$, $P_{\overline{e}}\overline{P}iPsPeP^\circ P \cdot P^\circ PSPSC_{\overline{u}}B \gg$).

$P''C-C$, $P_{\overline{e}}$. B

$P-P^\circ C_{\overline{f}}C$, $PsC_{\overline{f}}C_{\overline{f}}PIP^\circ PSPSC_{\overline{u}}$ $\overline{P}iC_{\overline{H}}PsC$, $P_{\overline{e}}\overline{P}iPsPeP^\circ P \cdot P^\circ PSPs$.

$\overline{P}_{\overline{u}}P_{\overline{u}}C_{\overline{H}}P_{\overline{u}}PrPsP \cdot C_{\overline{f}}PIP^\circ PSPSC_{\overline{u}}$.

P"°PSPëC... PïCḡPs PIPëPï°PrPePë PïPµCḡPµPrPsP·CḡPIP°PSPSCḡ P"ḡPsPrP°CḡC,PsPj
PSPµPjP°C". P"°P»C- PSP°PrP°C"C,CHCḡCḡ C-PSC,,PsCḡPjP°CḡC-Cḡ PïCḡPs
P·P°CḡC,PsCḡCḡPIP°PSPSCḡ PePsP¶PSPsPiPs PePsPjPïPsPSPµPSC,P° PsPeCḡPµPjPs.

P"ḡCḡC,P°CḡC,PµCḡPëPr

P—P° PrP°PSPëPjPë PeP»C-PSC-CḡPSPëC... PrPsCḡP»C-PrP¶PµPSCḡ, Cḡ
PrPsP±CḡPsPIPsP»CHCḡC-PI CḡP°P·PsPIC- PrPsP·Pë PrCḡC,P°CḡC,PµCḡPëPrCḡ PrPs 40
PjPi/PrPsP±Cḡ (CḡB 80 CḡP°P·C-PI PIPëC%oC- P·P° C,PµCḡP°PïPµPIC,PëCḡPSC-)
PïCḡPsC,CḡPiPsPj 7 PrPSC-PI PSPµ PIPëPeP»PëPeP°P»Pë P·P°PSPµPïPsPePsC"PSCḡ C%oPsPrPs
P±PµP·PïPµPePë C—C... P·P°CḡC,PsCḡCḡPIP°PSPSCḡ. PµC-Pr CḡP°Cḡ PeP»C-PSC-CḡPSPëC...
PrPsCḡP»C-PrP¶PµPSCḡ P·P°CḡC,PsCḡPsPICḡPIP°P»Pë PrPsP·Pë PrCḡC,P°CḡC,PµCḡPëPrCḡ PïPs
5 PjPi/PrPsP±Cḡ PïCḡPsC,CḡPiPsPj 6 PjC-CḡCḡCḡC-PI P±PµP· PïPsCḡPIPë
PrPsPrP°C,PePsPIPëC... PïPsP±C-CḡPSPëC... CḡPµP°PeCḡC-PNḡ PïPsCḡC-PIPSCḡPSPs C-P·
P·P°CḡC,PsCḡCḡPIP°PSPSCḡPj PrCḡC,P°CḡC,PµCḡPëPrCḡ Cḡ PrPsP·C- 0,5 PjPi/PrPsP±Cḡ.

PŸPïPµCḡPëC,,C-CḡPSPsPiPs P°PSC,PëPrPsC,Cḡ PSPµPjP°C", C,PsPjCḡ Cḡ CḡP°P·C-
PjPsP¶P»PëPIPsPiPs PïPµCḡPµPrPsP·CḡPIP°PSPSCḡ PïCḡPsPIPsPrCḡC,CH
CḡPëPjPiC,PsPjP°C,PëCḡPSCḡ C,P° PïC-PrC,CḡPëPjCḡPIP°P»CHPSCḡ C,PµCḡP°PïC-Cḡ.

PjP°PjCḡCḡP»PsP·PëPS

PḡP°PrC...PsPrPëP»Pë PïPsPIC-PrPsPjP»PµPSPSCḡ C%oPsPrPs PïPsCḡC,CḡPsPiPs
PïPµCḡPµPrPsP·CḡPIP°PSPSCḡ C,P°PjCḡCḡP»PsP·PëPSCḡ PïC-PrCḡPsC...P»PsCḡPëPrPsPj Cḡ
PrPsP·C- 5 PjPi, PIB CḡPµP·CḡP»CHC,P°C,C-B CḡPePsPiPsB
CḡPïPsCḡC,PµCḡC-PiP°P»P°CḡCHB PïPsCḡC,CḡP°B P°CḡC,PµCḡC-P°P»CHPSP°B
PiC-PïPsC,PµPSP·C-CḡB (CḡPëCḡC,PsP»C-CḡPSPëPNḡ B P°CḡC,PµCḡC-P°P»CHPSPëPNḡB
C,PëCḡPe 70 PjPj/Hg), P±P»CHPIP°PSPSCḡ C,P° PrC-P°CḡPµCḡ, CḡPeC- P»C-PeCḡPIP°P»Pë
C-PSC,,CḡP·C-C"ḡ CḡC-PrPëPSPë, PïC-CḡP»Cḡ CḡPsPiPs PïP°CḡC-C"PSC, PIC-PrCḡCḡPI
PïPsP»PµPiC¶PµPSPSCḡ Cḡ C,PsPNḡ CḡP°PjPëPNḡ PrPµPSCḡ. Pj CḡP°P·C- PïPsCḡC,CḡPsC—
P°CḡC,PµCḡC-P°P»CHPSPsC— PiC-PïPsC,PµPSP·C-C—, CḡPeP° PIPëPSPëPeP°C" PïC-CḡP»Cḡ
PïPµCḡPµPrPsP·CḡPIP°PSPSCḡ C,P°PjCḡCḡP»PsP·PëPSCḡ PïC-PrCḡPsC...P»PsCḡPëPrPsPj,
CḡP»C-Pr P·P°P±PµP·PïPµCḡPëC,Pë PïC-PrC,CḡPëPjPeCḡ PrC-CḡP»CHPSPsCḡC,C-
CḡPµCḡCḡPµPIPs-CḡCḡPrPëPSPSPsC— CḡPëCḡC,PµPjPë. PµP°CḡC-C"PSC,Cḡ PI C,P°PePsPjCḡ
CḡC,P°PSC- CḡP»C-Pr PïCḡPëPNḡPSCḡC,Pë PïPsCḡPëP·PsPSC,P°P»CHPSPµ
PïPsP»PsP¶PµPSPSCḡ, C%oPsP± PIC-PrPSPsPIPëC,Pë P°CḡC,PµCḡC-P°P»CHPSPëPNḡ C,PëCḡPe
C- PSPsCḡPjP°P»C-P·CḡPIP°C,Pë CḡP°CḡC,PsC,Cḡ CḡPµCḡCḡPµPIPëC... CḡPePsCḡPsCḡPµPSCḡ.
PïPeC%oPs CḡPµ PSPµ PrPsPïPsPjPsP¶Pµ, CḡP»C-Pr PïCḡPëP·PSP°CḡPëC,Pë
PïP»P°P·PjPsP·P°PjC-PSPSPëPePë, P° Cḡ CḡP°P·C- PSPµPsP±C...C-PrPSPsCḡC,C- ḡḡ"
CḡCḡPrPëPSPsP·PICḡP¶CḡPIP°P»CHPSC- P·P°CḡPsP±Pë. PµPsC,CḡC-P±PSPs CḡC,PµP¶PëC,Pë
P·P° C,,CḡPSPeCḡC-C"ḡ PSPëCḡPsPe C- PïCḡPsPIPsPrPëC,Pë
P·P°PiP°P»CHPSPsPïC-PrC,CḡPëPjCḡPIP°P»CHPSCḡ C,PµCḡP°PïC-Cḡ. P"ḡC-P°P»C-P· PjPsP¶Pµ
PIPëCḡPIPëC,PëCḡCḡ PSPµPµC,,PµPeC,PëPIPSPëPj, PsCḡPeC-P»CHPePë
C,P°PjCḡCḡP»PsP·PëPSCḡ PïC-PrCḡPsC...P»PsCḡPëPr PjP°PNḡP¶Pµ PïPsPIPSC-CḡC,Cḡ
P·PiBḡ™ CḡP·CḡC"C,CHCḡCḡ P· P±C-P»PeP°PjPë PïP»P°P·PjPë.

PµCḡPë PïPµCḡPµPrPsP·CḡPIP°PSPSC-, C%oPsP± P·P°PïPsP±C-PiC,Pë P°P±CḡPsCḡP±CḡC-C—,
Cḡ PïP°CḡC-C"PSC,P° PSPµPsP±C...C-PrPSPs PIPëPeP»PëPeP°C,Pë P±P»CHPIP°PSPSCḡ.
PïPeC%oPs PïCḡPëPNḡPSCḡC,Ps PIPµP»PëPeC- PrPsP·Pë PïCḡPµPïP°CḡP°C,Cḡ,
PSPµPsP±C...C-PrPSPs PïCḡPsPIPµCḡC,Pë PïCḡPsPjPëPIP°PSPSCḡ C¶P»CḡPSPeP°, PrP°C,Pë
P°PeC,PëPIPsPIP°PSPµ PICḡPiC-P»P»Cḡ C,P° PïCḡPsPSPsCḡPSPµ, PSP°PïCḡPëPeP»P°Pr
CḡCḡP»CHC,,P°C, PSP°C,CḡC-Cḡ.

РљР»С-РSC-С±РSC- РrPsCfP»С-РrP¶PµPSPSC¶ P·P°CfC,PsCfCfPIP°PSPSC¶ P”CfPsPrP°C¶C,Cf PSPµ PiC¶PsPIPsPrPëP»PëCfC¶, PsPrPSP°Pë P±CfP»Ps PiC¶PsPrPµPjPsPSCfC,C¶PsPIP°PSPs P±C-РsPµPePIC-PIP°P»PµPSC,PSC-CfC,CH P”CfPsPrP°C¶C,Cf PiC¶Pë CfCfPjC-CfPSPsPjCf P·P°CfC,PsCfCfPIP°PSPSC- РrCfC,P°CfC,PµC¶PëPrCf C,P° C,P°PjCfCfP»PsP·PëPSCf. P†PSC,,PsC¶PjP°C†C-С¶ PiC¶Ps PsPrPSPsC±P°CfPSPµ P·P°CfC,PsCfCfPIP°PSPSC¶ PsPrPµC¶P¶P°PSPs P· РrPsCfP»С-РrP¶PµPSPSC¶ CombAT (PePsPjP±C-PSP°C†C-С¶ P¶PIPsPrP°C¶C,Cf C,P° C,P°PjCfCfP»PsP·PëPSCf), PI С¶PePsPjCf PiPsC¶C-PIPSC¶PIP°P»Pë PePsPjP±C-PSP°C†C-С¶ PrCfC,P°CfC,PµC¶PëPrCf PI РrPsP·C- 0,5 PjPi C,P° C,P°PjCfCfP»PsP·PëPSCf 0,4 PjPi PsPrPëPS C¶P°P· PSP° РrPsP±Cf PiC¶PsC,С¶PiPsPj 4 C¶PsPeC-PI P°P±Ps PjPsPSPsC,PµC¶P°PiC-С¶ C†PëPjPë PiC¶PuPiP°C¶P°C,P°PjPë.

P†PSC,,PsCḡPjP°C†C-Cḡ PiCḡPs PīPsP±C-C†PSC- CḡPμP°PeC†C-C— PePsP¶PSPsPiPs
PePsPjPīPsPSPμPSC,P° PsPeCḡPμPjPs (PrCfC,P°CfC,PμCḡPēPr C,P° C,P°PjCfCfP»PsP·PēPS)
C,P°PePsP¶ PSP°PIPμPrPμPSPs PSPēP¶C†Pμ. PkPμ PiCḡPs PICfC- PīPsP±C-C†PSC-
CḡPμP°PeC†C-C—, C%Ps PIC-PrPjC-C†P°P»Pē PiCḡPē P·P°CfC,PsCfCfPIP°PSPSC-
PePsP¶PSPsPiPs PePsPjPīPsPSPμPSC,P° PsPeCḡPμPjPs, PīPsPIC-PrPsPjP»C‡P»Pē PiCḡPē
P·P°CfC,PsCfCfPIP°PSPSC- P°CfPsPrP°CḡC,Cf, C- C,PsPjCf C-PSC,,PsCḡPjP°C†C-Cḡ PiCḡPs
PīPsP±C-C†PSC- CḡPμP°PeC†C-C— PiCḡPē P·P°CfC,PsCfCfPIP°PSPSC- PsPeCḡPμPjPēC...
PePsPjPīPsPSPμPSC,C-PI P°CfPsPrP°CḡC,Cf C,P°PePsP¶ PIPEP»CḡC†PμPSPs PI PrP°PSCf
C-PSCfC,CḡCfPeC†C-Cḡ.

P—P° PrP°PSPëPjPë 4-C₇H₅-C- $\dagger$ PSPsPiPs PrPsC₇P»C-PrP¶PµPSPSC₇ CombAT, PIC-PrC₇PsC,PsPe
PiPsP±C-C $\dagger$ PSPëC... C₇H₅PµP°PeC $\dagger$ C-PN₂, PIPëP·PSP°C $\dagger$ PµPSPëC... PrPsC₇P»C-PrPSPëPeP°PjPë
PiC₇PsC,C₇PiPsPj PiPµC₇CëPsPiPs, PrC₇C₇PiPsPiPs, C,C₇PµC,C₇PsPiPs C,P°
C $\dagger$ PµC,PIPµC₇C,PsPiPs C₇PsPeC-PI P»C-PeC₇PIP°PSPSC₇, P·PjC-PSC₇PIP°PIC₇C₇:
PIC-PrPiPsPIC-PrPSPs 22 %, 6 %, 4 % C,P° 2 % PiC₇Pë PePsPjP±C-PSPsPIP°PSC-PN₂
C,PµC₇P°PiC-C— PrC₇C,P°C₇C,PµC₇PëPr C,P°PjC₇C₇P»PsP·PëPS; 15 %, 6 %, 3 % C,P° 2 %B
PiC₇Pë PjPsPSPsC,PµC₇P°PiC-C— PrC₇C,P°C₇C,PµC₇PëPrPsPj; 13 %, 5 %, 2 %, C,P° 2 % PiC₇Pë
PjPsPSPsC,PµC₇P°PiC-C— C,P°PjC₇C₇P»PsP·PëPSPsPj. P°PëC%PëPN₂ PIC-PrC₇PsC,PsPe
PiPsP±C-C $\dagger$ PSPëC... C₇H₅PµP°PeC $\dagger$ C-PN₂ C₇ PiC₇C₇PiC-, C₇PeP° PsC,C₇PëPjC₇PIP°P»P°
PePsPjP±C-PSPsPIP°PSC₇ C,PµC₇P°PiC-C₇, PiC₇PsC,C₇PiPsPj PiPµC₇CëPsPiPs C₇PsPeC₇
P»C-PeC₇PIP°PSPSC₇ P·C₇PjPsPIP»PµPSPëPN₂ PIPëC%PëPjPë PiPsPeP°P·PSPëPeP°PjPë
C₇H₅PµPiC₇PsPrC₇PeC,PëPIPSPëC... PiPsC₇C₇CëPµPSC₇, P° C₇P°PjPµ PiPsC₇C₇CëPµPSPSC₇
PµC₇PeC₇P»C₇C $\dagger$ C-C—, C₇PeC- C₇PiPsC₇C,PµC₇C-PIP°P»PëC₇C₇ C₇ PiC₇C₇PiC-.

$\text{P}_{\mu}\text{PsP}_{\pm}\text{C}-\text{C}\ddagger\text{PSC}-\text{C}\ddagger\text{P}_{\mu}\text{P}^{\circ}\text{PeC}\ddagger\text{C}-\text{C}-$, $\text{C}\%_{\text{Ps}}\text{PIP}\ddot{\text{e}}\text{PSP}\ddot{\text{e}}\text{PeP}^{\circ}\text{P}\gg\text{P}\ddot{\text{e}}\text{P}\cdot\text{C}\ddagger\text{P}^{\circ}\text{C}\ddagger\text{C},\text{PsC},\text{PsC}\ddot{\text{H}}\text{B}\geq 1\%$
 $\text{P}\ddot{\text{e}}\text{C}\ddot{\text{H}}\text{PsC},\text{C}\ddagger\text{P}\text{PsPj}\text{P}\ddot{\text{e}}\text{P}_{\mu}\text{C}\ddot{\text{H}}\text{C}\ddot{\text{e}}\text{PsP}\text{Ps}\text{C}\ddot{\text{H}}\text{PsPeC}\ddagger\text{P}\cdot\text{P}^{\circ}\text{C}\ddagger\text{C},\text{PsC}\ddagger\text{C}\ddagger\text{PIP}^{\circ}\text{PSPSC}\ddagger\ddagger, \text{P}\cdot\text{P}^{\circ}\text{PrP}^{\circ}\text{PSP}\ddot{\text{e}}\text{PjP}\ddot{\text{e}}\text{PrPsC}\ddagger\text{P}\gg\text{C}-\text{PrPSP}\ddot{\text{e}}\text{C}\ddagger\text{C}\ddot{\text{H}}\text{PeP}\ddot{\text{e}}\text{C}\dots\text{P}^{\circ}\text{PSP}^{\circ}\text{P}\gg\text{C}-\text{P}\cdot\text{C}-\text{PI}\text{CombAT}, \text{REDUCE}, \text{P}^{\circ}\text{C}, \text{P}^{\circ}\text{PePsP}\P$
 $\text{PeP}\gg\text{C}-\text{PSC}-\text{C}\ddagger\text{PSP}\ddot{\text{e}}\text{C}\dots\text{PrPsC}\ddagger\text{P}\gg\text{C}-\text{PrP}\P\text{P}_{\mu}\text{PSC}\ddot{\text{H}}\text{PjPsPSPsC}, \text{P}_{\mu}\text{C}\ddot{\text{H}}\text{P}^{\circ}\text{P}\ddot{\text{e}}\text{C}-\text{PN}_{\text{e}}$
 $\text{PePsPjP}\ddot{\text{e}}\text{PsPSP}_{\mu}\text{PSC}, \text{P}^{\circ}\text{PjP}\ddot{\text{e}}\text{P}^{\circ}\text{C}\ddagger\text{PsPrP}^{\circ}\text{C}\ddot{\text{H}}\text{C}, \text{C}\ddagger, \text{PSP}^{\circ}\text{PIP}_{\mu}\text{PrP}_{\mu}\text{PSC}-\text{PI}\text{C}, \text{P}^{\circ}\text{P}\pm\text{P}\gg\text{P}\ddot{\text{e}}\text{C}\ddagger\text{C}-.$

$P \vdash PSC, PsC \vdash PjP^{\circ}C \vdash C - C \vdash C\%PsPrPs \vdash P \vdash PsP \pm C - C \vdash PSP \ddot{e}C \dots C \vdash P_{\mu}P^{\circ}PeC \vdash C - PN_{\mathbb{Z}}$
 $C, P^{\circ}PjC \vdash C \vdash P \gg PsP \cdot P \ddot{e}PSC \vdash P \pm P^{\circ}P \cdot CrC^{\circ}C, CHC \vdash C \vdash PSP^{\circ}PrP^{\circ}PSP \ddot{e}C \dots, C\%PsPrPsC \vdash C, CrP \vdash PSC -$
 $PSP^{\circ}PIC - PrP \vdash PsPIC - PrPSP \ddot{e}C \dots PjP_{\mu}PrP \ddot{e}C \vdash PSP \ddot{e}C \dots C \vdash P_{\mu}C \vdash C \vdash C \vdash C \vdash P^{\circ}C \dots P \ddot{S}P^{\circ}C \vdash C, PsC, P^{\circ}$
 $C - C \dots PIP \ddot{e}PSP \ddot{e}PePSP_{\mu}PSPSC \vdash PjPsP \vdash P_{\mu}P \cdot C \vdash PsC \vdash C, P^{\circ}C, P \ddot{e}PiC \vdash P \ddot{e}PsPrPSPsC \vdash P^{\circ}C \vdash PSPsPjCr$
 $P \cdot P^{\circ}C \vdash C, PsC \vdash C \vdash PIP^{\circ}PSPSC - PrCrC, P^{\circ}C \vdash C, P_{\mu}C \vdash P \ddot{e}PrCrC, P^{\circ}C, P^{\circ}PjC \vdash C \vdash P \gg PsP \cdot P \ddot{e}PSC \vdash$

P\$P°CfC,PsC,P° PIPëPSPëPePSPµPSPSCl PİPsP±C-C±PSPëC... CĤPµP°PeC†C-PNē,
PIPëCłIPIP»PµPSPëC... Cř PeP»C-PSC-C±PSPëC... PrPsCłP»C-PrP¶PµPSPSClC...: C±P°CfC,Ps (≥
1/100 B B-B B B B B B B B < 1/10), PSPµC±P°CfC,PsB (≥ 1/1000 B- B < 1/100), CĤC-PrPePs (≥
1/10000 B- B < 1/1000), PrCřP¶Pµ CĤC-PrPePs (< 1/10000). PµPsP±C-C±PSC- CĤPµP°PeC†C-C—,
PeP»P°CłPëC„C-PePsPIP°PSC- P·P° PeP»P°CłP°PjPë CłPëCłC,PµPj PsCĤPİP°PSC-PI,
PİCĤPµPrCłC,P°PIP»PµPSPs Cř PİPsCĤCłIPrPeCř P·PSPëP¶PµPSPSCl C—C... CłC,CřPİPµPSCl
C,CłP¶PePsCłC,C-.

PµP»P°Cł CłPëCłC,PµPjPë PsCĤPİP°PSC-PI	PµPsP±C-C±PSC- CĤPµP°PeC†C-C—	P°CřC,P°CfC,PµCĤPëPr C,P°PjCłCřP»PsP·PëPS	P°CřC,P°CfC,PµCĤPëPr	PÿP°PjCłCřP»PsP·PëPS
P— P±PsPeCř PSPµCĤPIPSPİPsC— CłPëCłC,PµPjPë	PkPµPİCĤPëC,PsPjPSC-CłC,CH P—P°PİP°PjPsCĤPsC†PµPSPSCl P°PsP»PsPIPSPëPNē P±C-P»CH	- P\$P°CfC,Ps -	- -	P C-PrPePs P\$P°CfC,Ps PkPµC±P°CfC,Ps
P— P±PsPeCř CłPµCĤC†Cł	B PÿPµCĤC†PµPIP° PSPµPrPsCłC,P°C,PSC-CłC,CH (P·P±C-CĤPSPµ PİPsPSClC,C,Cł¹) PµPsCłPëP»PµPSPµ CłPµCĤC†PµP±PëC,C,Cł	PkPµC±P°CfC,Ps -	PkPµC±P°CfC,Ps ^d -	- PkPµC±P°CfC,Ps
P— P±PsPeCř CłCřPrPëPSPSPsC— CłPëCłC,PµPjPë	PµPsCłC,CřCĤP°P»CHPSP° PİC-PİPsC,PµPSP·C-Cł	-	-	PkPµC±P°CfC,Ps
P— P±PsPeCř PrPëC...P°P»CHPSPsC— CłPëCłC,PµPjPë, PsCĤPİP°PSC-PI PİCĤCřPrPSPsC— PeP»C-C,PëPë C-B CłPµCĤPµPrPsCłC,C-PSPSCl	P PëPSC-C,	-	-	PkPµC±P°CfC,Ps
P— P±PsPeCř CëP»CřPSPePsPIP- PePëCëPePsPIPSPİPs C,CĤP°PeC,Cř	P—P°PİPsCĤ P°C-P°CĤPµCł PkCřPrPsC,P° P°P»CĤPIP°PSPSCl	- - - -	- - - -	PkPµC±P°CfC,Ps PkPµC±P°CfC,Ps PkPµC±P°CfC,Ps PkPµC±P°CfC,Ps
P— P±PsPeCř CëPeC-CĤPë C,P° PİC-PrCëPeC-CĤPSPëC... C,PeP°PSPëPS	PĥPSPİC-PsPSPµPİCĤPsC,PëC†PSPëPNē- PSP°P±CĤCłPe PÿPëPSPrCĤPsPj PÿC,C-PIPµPSCłP° BĤ° P°P¶PsPSCłPsPSP° PµCĤPsPİPëPİCłPSPeP° P°PëCłPëPİ PÿPİPµCĤP±C-P¶	- - - - - -	- - - - -	P C-PrPePs P°CřP¶Pµ CĤC-PrPePs PkPµC±P°CfC,Ps PkPµC±P°CfC,Ps PkPµC±P°CfC,Ps
P— P±PsPeCř CĤPµPİCĤPsPrCřPeC,PëPIPSPsC— CłPëCłC,PµPjPë C- PİCĤCřPrPSPëC... P·P°P»PsP·	PµCĤC-P°PİC-P·Pj P†PjPİPsC,PµPSC†C-Cł³ PµPsCĤCřCëPµPSPSCl³ (P·PSPëP¶PµPSPSCl) P»C-P±C-PrPs PµPsCĤCřCëPµPSPSCl³~ PµCłPeCřP»CłC†C-C— P—P°C...PİPsCĤCĤPIP°PSPSCl PİCĤCřPrPSPëC... P·P°P»PsP·²	- P\$P°CfC,Ps P\$P°CfC,Ps P\$P°CfC,Ps P\$P°CfC,Ps P\$P°CfC,Ps	P\$P°CfC,Ps ^b P\$P°CfC,Ps ^b P\$P°CfC,Ps ^b -	- - P\$P°CfC,Ps -
P—P°PİP°P»CHPSC- CĤPsP·P»P°PrPë	PĥCłC,PµPSC-Cł	-	-	PkPµC±P°CfC,Ps

aB B B B B B B B B B B B P°CřC,P°CfC,PµCĤPëPr C,P°PjCłCřP»PsP·PëPS:B Cř PrPsCłP»C-PrP¶PµPSPSC-
CombAT C±P°CfC,PsC,P° C†PëC... PİPsP±C-C±PSPëC... CĤPµP°PeC†C-PNē
P·PSPëP¶CřC°C,CHCłCł P· PePsP¶PSPëPj PSP°CfC,CřPİPSPëPj CĤPsPePsPj P· 1-PİPs PrPs 4-PİPs.

bB B B B B B B B B B B P— PrPsCłP»C-PrP¶PµPSCĤ PjPsPSPsC,PµCĤP°PİC-C—
PrPsP±CĤPsCłPeC-CłPSPsC— PİC-PİPµCĤPİP»P°P·C-C— PİPµCĤPµPrPjC-C...CřCĤPsPIPSC—
P·P°P»PsP·Pë PrCřC,P°CfC,PµCĤPëPrPsPj.

cB B B B B B B B B B B B P— PsCłPSPsPIPSPsC— C-PSC„PsCĤPjP°C†C-C— P· P±PµP·PİPµPePë
C,P°PjCłCřP»PsP·PëPSCř PrP»Cł PeCĤP°C—PS P·Pÿ.

dB B B B B B B B B B B B P°PsCłP»C-PrP¶PµPSPSCl REDUCE.

¹B B B B B B B B B B B P—P±C-CĤPSPµ PİPsPSClC,C,Cł B«CłPµCĤC†PµPIP°
PSPµPrPsCłC,P°C,PSC-CłC,CHB» PİPeP»CĤC±P°C° P·P°CłC,C-PNēPSCř CłPµCĤC†PµPİCř

PSPμPrPsCfC,P°C,PSC-CfC,CH, CΓPμCḡC†PμPICr PSPμPrPsCfC,P°C,PSC-CfC,CH,
PSPμPrPsCfC,P°C,PSC-CfC,CH P»C-PIPsPiPs C€P»CfPSPsC†PeP°, PiPsCfC,CḡCf
CΓPμCḡC†PμPICr PSPμPrPsCfC,P°C,PSC-CfC,CH, PeP°CḡPrC-PsPiPμPSPSPëPNë C€PsPe,
PiPsCfC,CḡCf PSPμPrPsCfC,P°C,PSC-CfC,CH P»C-PIPsPiPs C€P»CfPSPsC†PeP°,
PSPμPrPsCfC,P°C,PSC-CfC,CH PiCḡP°PIPsPiPs C€P»CfPSPsC†PeP°, PiPsCfC,CḡCf
PSPμPrPsCfC,P°C,PSC-CfC,CH PiCḡP°PIPsPiPs C€P»CfPSPsC†PeP°,
PSPμPrPsCfC,P°C,PSC-CfC,CH C€P»CfPSPsC†PeP°,
PeP°CḡPrC-PsPiC†P»CHPjPsPSP°P»CHPSCr PSPμPrPsCfC,P°C,PSC-CfC,CH,
B P·P°CfC,C-PNëPSCr PeP°CḡPrC-PsPiP°C,C-CH.

2B B B B B B B B B B B B B B B B P'PeP»CHC†P°CḡC†Pë PiC-PiPμCḡPμCfC,PμP·C-CH C,P°
P·P±C-P»CHC€PμPSPSCḡ PiCḡCfPrPSPëC... P·P°P»PsP·.

3B B B B B B B B B B B B B B B B PiPsP±C-C†PSC- CḡPμP°PeC†C-C—, PiPsPiBḡ™ CḡP·P°PSC- P·
PiPsCḡCfC€PμPSPSCḡPj CΓPμPeCfC†P°P»CHPSPsC— C„CfPSPeC†C-C—,
P°CfPsC†C-CHCHC,CHCfCḡ P· P»C-PeCfPIP°PSPSCḡPj PrCfC,P°CfC,PμCḡPëPrPsPj
(PIPeP»CHC†P°CḡC†Pë PjPsPSPsC,PμCḡP°PiC-CH C,P° PePsPjP±C-PSP°C†C-CH P·
C,P°PjCfC†P»PsP·PëPSPsPj). PiC- PiPsP±C-C†PSC- CḡPμP°PeC†C-C— PjPsP¶CfC,CH
C,CḡPëPIP°C,Pë PiC-CfP»Cḡ PiCḡPëPiPëPSPμPSPSCḡ P»C-PeCfPIP°PSPSCḡ. P'PiP»PëPI
PrCfC,P°CfC,PμCḡPëPrCf PSP° C—C... C,CḡPëPIP°P»C-CfC,CH PSPμPIC-PrPsPjP°.

^B B B B B B B B P'PeP»CHC†P°CḡC†Pë P·PjPμPSC€PμPSPSCḡ PsP±Bḡ™ C”PjCf CΓPiPμCḡPjPë.

P”P°PSC- PiPsCfC,PjP°CḡPePμC,PëPSPiPsPIPëC... PrPsCfP»C-PrP¶PμPSCḡ

PJ PiPsCfC,PjP°CḡPePμC,PëPSPiPsPIPsPjCf CΓPiPsCfC,PμCḡPμP¶PμPSPSC- PiPsP±C-C†PSC-
CḡPμP°PeC†C-C— C„C-PeCfC†PIP°P»PëCfCḡ P·C- CΓPiPsPSC,P°PSPSPëC...
PiPsPIC-PrPsPjP»PμPSCḡ, C,PsPjCf C,PsC†PSP° C†P°CfC,PsC,P° C,P°PePëC... CḡPμP°PeC†C-PNë
PSPμPIC-PrPsPjP°.

PḡPsPSPsC,PμCḡP°PiC-Cḡ PrCfC,P°CfC,PμCḡPëPrPsPj

P— P±PsPeCž C-PjCžPSPSPsC— CΓPëCfC,PμPjPë

PŞP°CfC,PsC,P° PSPμPIC-PrPsPjP°: P°P»PμCḡPiC-C†PSC- CḡPμP°PeC†C-C—,
PIPeP»CHC†P°CḡC†Pë PiPëCfPëPi, CΓPIPμCḡP±C-P¶, PeCḡPsPiPëPiBḡ™ CḡPSPeCf,
P»PsPeP°P»C-P·PsPIP°PSPëPNë PSP°P±CḡCḡPe C,P° P°PSPiC-PsPSPμPICḡPsC,PëC†PSPëPNë
PSP°P±CḡCḡPe.

P— P±PsPeCž PiCfPëC...C-PePë

PŞP°CfC,PsC,P° PSPμPIC-PrPsPjP°: PrPμPiCḡPμCfC-Cḡ.

P— P±PsPeCž C€PeC-CḡPë C,P° PiC-PrC€PeC-CḡPSPsC— PeP»C-C,PePsPIPëPSPë

P C-PrPePs: P°P»PsPiPμC†C-Cḡ (PiPsP»PsPIPSPëPj C†PëPSPsPj, PIC,CḡP°C,P° PIPsP»PsCfCfCḡ
PSP° C,C-P»C-), PiC-PiPμCḡC,CḡPëC...PsP·.

P— P±PsPeCž CḡPμPiCḡPsPrCžPeC,PëPIPSPsC— CΓPëCfC,PμPjPë C,P° PiCḡCžPrPSPëC...
P·P°P»PsP·.

$\text{P}\ddot{\text{S}}\text{P}^{\circ}\text{C}\dot{\text{I}}\text{C}, \text{PsC}, \text{P}^{\circ} \text{PSP}_{\mu}\text{PIC}-\text{PrPsPjP}^{\circ}: \text{C}, \text{P}_{\mu}\text{C}\dot{\text{I}}\text{C}, \text{P}\ddot{\text{e}}\text{PeC}\dot{\text{r}}\text{P}\gg\text{C}\dot{\text{I}}\text{C}\ddot{\text{H}}\text{PSP}\ddot{\text{e}}\text{PN}_{\text{e}} \text{P}\pm\text{C}-\text{P}\gg\text{CHb} \text{C}, \text{P}^{\circ} \text{PSP}^{\circ}\text{P}\pm\text{C}\ddot{\text{H}}\text{C}\dot{\text{I}}\text{Pe}.$

$$\underline{\text{P}_{\text{Hb}}\text{P}_{\text{S}}\text{P}_{\text{S}}\text{C}, \text{P}_{\text{H}}\text{C}_{\text{H}}\text{P}^{\circ}\text{P}_{\text{I}}\text{C}-\text{Cl} \text{ C}, \text{P}^{\circ}\text{P}_{\text{j}}\text{C}_{\text{I}}'\text{C}_{\text{r}}\text{P}}\text{P}_{\text{S}}\text{P}\cdot\text{P}_{\text{e}}\text{P}_{\text{S}}\text{P}_{\text{j}}}$$

P—P° PrP°PSPëPjPë PiPsCÍ'C,PjP°CĤjPePμC,PëPSPiPsPIPsPiPs CÍ'PiPsCÍ'C,PμCĤjPμP¶PμPSPSC¶
PiC—Pr C‡P°CÍ' PsPiPμCĤjP°C†C—C— P· PiCĤjPëPIPsPrCr PeP°C,P°CĤjP°PeC,Pë C,P°
PiP»P°CrPePsPjPë Cr PrPμC¶PePëC... PiP°C†C—C"PSC,C—PI, C¶PeC— CĤjP°PSC—CεPμ
PsC,CĤjPëPjCÍ'PIP°P»Pë P°P»CHC,,P°1—P°PrCĤjPμPSPsP±P»PsPeP°C,PsCĤjPë,
PIPeP»CHC‡P°CHC‡Pë C,P°PjCÍ'CrP»PsP·PëPS, P±CrPI PIC—PrP·PSP°C‡PμPSPëPN₂
C—PSC,CĤjP°PsPiPμCĤjP°C†C—PN₂PSPëPN₂ CÍ'PëPSPrCĤjPsPj P°C,PsPSC—C‡PSPsC—
CĤjP°PM₆PrCrP¶PePë (P†PŸP₃P , PIP°CĤjC—P°PSC, CÍ'PëPSPrCĤjPsPjCÍ' PICrP·CHPePsC—
P·C—PSPëC†C—) (PrPëPI. B«PhCÍ'PsP±P»PëPIPsCÍ'C,C— P·P°CÍ'C,PsCÍ'CrPIP°PSPSC¶B»).

$\text{P}_{\mu}\text{C}-\text{Pr}\text{C}\ddagger\text{P}^{\circ}\text{C}\acute{\text{r}}\text{P}\ddot{\text{r}}\text{C}-\text{C}\acute{\text{r}}\text{P}\gg\text{C}\ddot{\text{r}}\text{C}\ddot{\text{r}}\text{P}_{\mu}\text{C}^{\circ}\text{C}\acute{\text{r}}\text{C},\text{C}\ddot{\text{r}}\text{P}^{\circ}\text{C}\ddagger\text{C}-\text{P}\text{N}_{\text{e}}\text{PSPsPiPs}\text{P}\cdot\text{P}^{\circ}\text{C}\acute{\text{r}}\text{C},\text{PsC}\acute{\text{r}}\text{C}\acute{\text{r}}\text{PIP}^{\circ}\text{PSPSC}\ddot{\text{r}}\text{P}\pm\text{C}\acute{\text{r}}\text{P}\gg\text{P}\ddot{\text{e}}\text{PrPsPrP}^{\circ}\text{C},\text{PePsPIC}-\text{P}\ddot{\text{r}}\text{PsPIC}-\text{PrPsPjP}\gg\text{P}_{\mu}\text{PSPSC}\ddot{\text{r}}\text{P}\ddot{\text{r}}\text{C}\ddot{\text{r}}\text{Ps}\text{PIP}\ddot{\text{e}}\text{P}\ddot{\text{r}}\text{P}^{\circ}\text{PrPeP}\ddot{\text{e}}\text{C},,\text{C}-\text{P}\pm\text{C}\ddot{\text{r}}\text{P}\ddot{\text{e}}\text{P}\gg\text{C}\ddot{\text{r}}\text{C}\ddagger\text{C}-\text{C}-\text{P}\ddot{\text{r}}\text{P}_{\mu}\text{C}\ddot{\text{r}}\text{P}_{\mu}\text{PrC}\acute{\text{r}}\text{P}_{\mu}\text{C}\ddot{\text{r}}\text{PrC}\text{H},\text{P}^{\circ}\text{C}\ddot{\text{r}}\text{P}\ddot{\text{e}}\text{C},\text{PjC}-\text{C}-,\text{C},\text{P}^{\circ}\text{C}...\text{C}-\text{PeP}^{\circ}\text{C}\ddot{\text{r}}\text{PrC}-\text{C}-,\text{PrP}\ddot{\text{e}}\text{C}\acute{\text{r}}\text{P}\ddot{\text{r}}\text{PSPsP}_{\mu},\text{B}\text{PSPsC}\acute{\text{r}}\text{PsPIP}\ddot{\text{e}}\text{C}...\text{PeC}\ddot{\text{r}}\text{PsPIP}\text{PsC},\text{P}_{\mu}\text{C}\ddagger,\text{P}\ddot{\text{r}}\text{PsC}\ddot{\text{r}}\text{C}\text{€P}_{\mu}\text{PSC}\text{H}\text{P}\cdot\text{PsC}\ddot{\text{r}}\text{C}\acute{\text{r}},\text{P}\text{I}\text{C},,\text{C}\ddagger.\text{C}\acute{\text{r}}\text{PIP}\ddot{\text{e}}\text{P}\ddot{\text{r}}\text{P}\gg\text{C}\ddot{\text{r}}\text{PrC}-\text{P}\cdot\text{PSP}\ddot{\text{e}}\text{P}\P\text{P}_{\mu}\text{PSPSC}\ddot{\text{r}}\text{P}\text{N}_{\text{e}}\text{PsPiPs}\text{PiPsC}\acute{\text{r}}\text{C},\text{C}\ddot{\text{r}}\text{PsC},\text{P}\ddot{\text{e}},\text{PjC}\acute{\text{r}}\text{P}\gg\text{C}\text{H}\text{C},\text{P}\ddot{\text{e}}\text{C},,\text{PsC}\ddot{\text{r}}\text{PjPSPsC}-\text{P}_{\mu}\text{C}\ddot{\text{r}}\text{P}\ddot{\text{e}}\text{C},\text{P}_{\mu}\text{PjP}\ddot{\text{e}},\text{P}_{\mu}\text{PeC}\acute{\text{r}}\text{C},,\text{PsP}\gg\text{C}-\text{P}^{\circ}\text{C},\text{P}\ddot{\text{e}}\text{PIPSPsPiPs}\text{PrP}_{\mu}\text{C}\ddot{\text{r}}\text{PjP}^{\circ}\text{C},\text{P}\ddot{\text{e}}\text{C},\text{C}\acute{\text{r}}\text{C},\text{P}^{\circ}\text{C}\acute{\text{r}}\text{C}\acute{\text{r}}\text{C}...\text{PsC}\acute{\text{r}}\text{C},\text{C}-\text{C}\acute{\text{r}}\text{P}\gg\text{P}\ddot{\text{e}}\text{P}\cdot\text{PsPIP}\text{PsC}-\text{PsP}\pm\text{PsP}\gg\text{PsPSPeP}\ddot{\text{e}}\text{C}\ddot{\text{r}}\text{PsC},\text{PsPIP}\text{PsC}-\text{P}\ddot{\text{r}}\text{PsC}\ddot{\text{r}}\text{PsP}\P\text{PSP}\ddot{\text{e}}\text{PSP}\ddot{\text{e}},\text{P}^{\circ}\text{C}\acute{\text{r}}\text{PsC}\ddagger\text{C}-\text{P}\text{N}_{\text{e}}\text{PsPIP}^{\circ}\text{PSC}-\text{C}-\text{P}\cdot\text{P}\cdot\text{P}^{\circ}\text{C}\acute{\text{r}}\text{C},\text{PsC}\acute{\text{r}}\text{C}\acute{\text{r}}\text{PIP}^{\circ}\text{PSPSC}\ddot{\text{r}}\text{Pj}\text{C},\text{P}^{\circ}\text{PjC}\acute{\text{r}}\text{C}\acute{\text{r}}\text{P}\gg\text{PsP}\cdot\text{P}\ddot{\text{e}}\text{PSC}\acute{\text{r}}.$

P†PSC€C- P†P°PSC-

Pj C...PsPrC- PeP»C-PSC-C†PSPsPiPs PrPsCfP»C-PrP¶PµPSPSC¶ (PrPsCfP»C-PrP¶PµPSPSC¶ REDUCE) Cr C†PsP»PsPIC-PeC-PI, C¶PeC- P»C-PeCrPIP»P»PëCfC¶ PrCrC,P°CfC,PµC¶PëPrPsPj, PIC-PrP·PSP°C†P°P»P°CfC¶ PIPëC%oP° C†P°CfC,PsC,P° PIPëPSPëPePSPµPSPSC¶ C¶P°PeCr PîPµC¶PµPrPjC-C...CrC¶PsPIPsC— P·P°P»PsP·PëB (P·P° C€PeP°P»PsC¶ P°P»C-CfPsPSP° Б¶“ 8Б¶“10) PîPsC¶C-PIPsC¶PSPs P· PiC¶CrPîPsC¶ PîP»P°C†PµP±Ps (PrPëPI. C¶PsP·PrC-P»Pë B«PhCfPsP±P»PëPIPsCfC,C- P·P°CfC,PsCfCrPIP°PSPSC¶B» C,P° B«PαP°C¶PjP°PePsP»PsPiC-C†PSC- PIP»P°CfC,PëPIPsCfC,C-B»). PµC¶PëC†PëPSPSPsPiPs P·PIБ¶™ C¶P·PeCr PjC-P¶ P·P°CfC,PsCfCrPIP°PSPSC¶Pj PrCrC,P°CfC,PµC¶PëPrCr C,P° PIPëPSPëPePSPµPSPSC¶Pj C¶P°PeCr PîPµC¶PµPrPjC-C...CrC¶PsPIPsC— P·P°P»PsP·Pë PIPëCfPsPePsC— PiC¶P°PrP°C†C-C— P·P° P°P»C-CfPsPSPsPj PICfC,P°PSPsPIP»PµPSPs PSPµ P±CrP»Ps.

P—P° PrP°PSPëPjPë PeP»C—PSC—C‡PSPëC... PrPsCÍP»C—PrP¶PµPSCH C,P°
PiPsCÍC,PjP°C‡PePµC,PëPSPiPsPIPëC... CÍPiPsCÍC,PµC‡PµP¶PµPSCH P±CÍP»Pë
PiPsPIC—PrPsPjP»PµPSPSC‡ PiC‡Ps PIPëPiP°PrPePë C‡P°PeCÍ PiC‡CÍPrPSPsC— P·P°P»PsP·Pë CÍ
C‡PsP»PsPIC—PeC—PI (PrPëPI. C‡PsP·PrC—P» B«PhCÍPsP±P»PëPIPsCÍC,C—
P·P°CÍC,PsCÍCÍPIP°PSPSC‡B»).

P_γP_μC_βP_jC-PS P_iC_βP_öP_ΓP[°]C,PSP_sC_ΓC,C-.

2 C₇H₅PsPεPë.

PJPjPsPIPë P·P±PμCḡC-PiP°PSPSCİİ.

P—P±PμCḡC-PiP°C,Pë PiCḡPë C,PμPjPiPμCḡP°C,CrCḡC- PSPμ PIPëC‰Pμ 25 B°PŸ.

P—P±PμCḡC-PiP°C,Pë Cr PSPμPrPsCḡC,CrPiPSPsPjCr PrP»Cİİ PrC-C,PμPNë PjC-CḡC†C-.

PJPiP°PePsPIPeP°.

PkPμPiCḡPsP·PsCḡC- C„P»P°PePsPSPë P· PiPsP»C-PμC,PëP»PμPSCr PIPëCḡPsPePsC—
C‰C-P»CHbPSPsCḡC,C- P±C-P»PsPiPs PePsP»CHbPsCḡCr P·
PiPsP»C-PiCḡPsPiC-P»PμPSPsPIPëPjPë PeCḡPëC€PeP°PjPë, C‰Ps PjP°CḡC,CHb P·P°C...PëCḡC,
PIC-Pr PIC-PrPeCḡPëC,C,Cİİ PrC-C,CHbPjPë, PiPs 30 P°P±Ps PiPs 90 PeP°PiCḡCrP» Cr
C„P»P°PePsPSC-; PiPs 1 C„P»P°PePsPSCr PI PeP°CḡC,PsPSPSC-PNë PePsCḡPsP±C†C-.

PḡP°C,PμPiPsCḡC-Cİİ PIC-PrPiCḡCḡPeCr. P—P° CḡPμC†PμPiC,PsPj.

P’PëCḡPsP±PSPëPe.

PḡP°C,P°P»PμPSC, P”P¶PμCḡPjPμPSC- PëPsCḡPSPrPsCḡC„ P”PjP±PIḡ, PkC-PjPμC†C†PëPSP°/

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**PḡC-CḡC†PμP·PSP°C...PsPrP¶PμPSPSCİİ PIPëCḡPsP±PSPëPeP° C,P° PNëPsPiPs
P°PrCḡPμCḡP° PjC-CḡC†Cİİ PiCḡPsPIP°PrP¶PμPSPSCİİ PrC-CİİP»CHbPSPsCḡC,C-.**

PëC,P°PNëPSP±P°PNëC€C,CḡP°CḡCḡPμ 1 C- 2, PëPsCḡPSPrPsCḡC„, P’P°PrPμPS-
P’CrCḡC,PμPjP±PμCḡPi, 73614, PkC-PjPμC†C†PëPSP°/

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