

РiРќРЎРЎР РЈРљРiРiРi
РiР»CЦ РjРмРrРёC‡PSPsPiPs P·P°CfC,PsCfCfPiP°PSPSCЦ P»C-PeP°CЦCfCђPePsPiPs P·P°CfPsP±Cf

РђР°РiРљРЎР

(ABIXA)

РЎРёРёР»Р°Рr:
PrC-CђC‡P° CђPмC‡PsPIPёPSP°: memantine;
1 C,P°P±P»PмC,PeP° PjC-CfC,РёC,CH PjPмPjP°PSC,РёPSCf PiC-PrCђPsC...P»PsCђРёPrCf 10 PjPi;
PrPsPiPsPjC-P¶PSC- CђPмC‡PsPIPёPSPё: C‡PмP»CHP»PsP·P° PjC-PeCђPsPeCђPёCfC,P°P»C-C‡PSP°,
PSP°C,CђC-Cђ PeCђPsCfPeP°CђPjPмP»PsP·P°, PeCђPмPjPSC-Cђ PrC-PsPeCfPёPr PePsP»PsC—PrPSPёPм
P±PмP·PiPsPrPSPёPм, PjP°PiPSC-Cђ CfC,PмP°CђP°C,;
PsP±PsP»PsPSPeP°:B PiC-PiCђPsPjPмP»PsP·P°, B C,РёC,P°PSCfB PrC-PsPeCfPёPrB (P• 171),B
PjP°PeCђPsPiPsP»B 400,B P·P°P»C-P·P° PsPeCfPёPr P¶PsPIC,РёPм (P• 172).

Р·C-РёР°CђCfCђPeP° C„PsCђPjP°. РђР°РiРљРљC,PePё, PiPeCђPёC,C- PiP»C-PiPePsPiPsCH
PsP±PsP»PsPSPePsCH.
PhCfPSPsPiPSC- C„C-P·PёPePs-C...C-PjC-C‡PSC- PIP»P°CfC,РёPiPsCfC,C-: C,P°P±P»PмC,PePё, PiPeCђPёC,C-
PiP»C-PiPePsPiPsCH PsP±PsP»PsPSPePsCH, PIC-Pr P±P»C-PrPs-P¶PsPIC,PsPiPs PrPs P¶PsPIC,PsPiPs
PePsP»CHPsCђCf, PsPiP°P»CHPSPSC—C„PsCђPjPё, P· P»C-PSC-C°CH PrP»CЦ CђPsP·P»PsPjCf C,P°
PjP°CђPeCfPiP°PSPSCЦPj PIC-PrPSPsCfPSPs C—C—: B«РмB» P»C-PiPsCђCfC‡C- PiCђP°PiPsCђCfC‡P·
PsPrPSPsPiPs P±PsPeCf C- B«1B» P»C-PiPsCђCfC‡C- B«0B» PiCђP°PiPsCђCfC‡C‡ вђ” P· C-PSCёPsPiPs.
РђР°РiРљРљC,PeP° PjPsP¶Pм P±CfC,Рё CђPsP·PrC-P»PмPSP° PSP° PrPIC- PiPsP»PsPiPёPSPё.

Р»Р°CђPjP°PePsC,РмCђP°PiРмPIC,РёC‡PSP° PiCђCfPiP°.
PмCfPёC...PsP°PSP°P»PмPiC,РёPePё. P‡PSCёC- P·P°CfPsP±Pё PrP»CЦ P·P°CfC,PsCfCfPiP°PSPSCЦ Cf
CђP°P·C- PrPмPjPмPSC‡C—, РмPмPjP°PSC,РёPS.
РмPsPr РђРЎРЎ N06D X01.

Р»Р°CђPjP°PePsP»PsPiC-C‡PSC- PIP»P°CfC,РёPiPsCfC,C-.

PaP°CḡPjP°PePsFrPēPSP°PjC-PeP°.

Pj PīCḡPsCūPIP°C... CīPēPjPīC,PsPjC-PI C- PīCḡPsPīCḡPūCīCfPIP°PSPSC-
PSḡPūPēCḡPsPrPūPīPūPSPūCḡP°C,PēPIPSPsC— PrPūPjPūPSCīC-C— PIP°PḡP°PēPICf CḡPsP>Cḡ
PIC-PrC-PīCḡP°C” PīPsCḡCfCēPūPSPSCū PīP>CfC,P°PjP°C,PūCḡPīC-CāPSPsC—
PSḡPūPēCḡPsC,CḡP°PSCīPjC-CīC-C—, PsCḡPsP±P°PēPIPs P- CīCāP°CīC,Cḡ NMDA (N-PjPūC,PēP>-D-
P°CīPīP°CḡC,P°C.)-CḡPūCīPūPīC,PsCḡC-PI.
PḡPūPjP°PSC,PēPS CūPIP°CūC” CīPpsP±PsCḡ PīPsC,PūPSCīC-P°P>P°P>PūPḡPSPēPḡ,
CīPūCḡPūPrPSCḡPsC— P°C,C-PSPSPsCīC,C-, PSḡPūPePsPSPeCfCḡPūPSC,PSPēPḡ P°PSC,P°PīPsPSC-CīC,
NMDA-CḡPūCīPūPīC,PsCḡC-PI. PḡPūPjP°PSC,PēPS PjPsPrCīP>CḡC” PūC,PūPeC,Pē PīP°C,PsP>PsPīC-CāPSPs
PīC-PrPīPēC%PūPSPēC... CḡC-PīPSC-PI PīP>CfC,P°PjP°C,Cf, CūPePēPḡ PjPsPḡPū PīCḡPēP-PīPūCīC,Cē PrPs
PrPēCīC,CīPSPeCīC-C— PSPūPḡCḡPsPSC-PI.

PḡP>C-PSC-CāPSC- PrPsCīP>C-PrPḡPūPSPSCū
PḡCīPSPsPIPSḡPū PrPsCīP>C-PrPḡPūPSPSCū PjPsPSPsC,PūCḡP°PīC-C— Cf PīP°CīC-C”PSC,C-PI C-P·
CīPūCḡPūPrPSCḡPsCḡ C,P° C,CūPḡPePsCḡ CīC,P°PrC-C”Cḡ C...PīPsCḡPsP±Pē
PḡP>CḡCīPīPūPḡPjPūCḡP° (P·P° PḡPsCḡPsC,PePsCḡ CēPeP°P>PsCḡ PsCīC-PSPePē
PīCīPēC...C-PSPSPsPīPs CīC,P°C,CīCīCīC (MMSE) PsCīC-PSPeP° 3ḡḡ”14) PīPeP>CḡCīP°P>Ps
P·P°PīP°P>PsPj 252 P°PjP±CīP>P°C,PsCḡPSPēC... PīP°CīC-C”PSC,C-PI. PūPsP·PēC,PēPIPSPēPḡ
PīPīP>PēPI C,PūCḡP°PīC-C— PjPūPjP°PSC,PēPSPsPj PīPēCūPIP>CUC”C,CḡCīCū PīC-CīP>Cū 6
PjC-CīCūCīC-PI P·P°CīC,PsCīCīPīP°PSPSCū PīPsCḡC-PīPSCūPSPs P· P>C-PeCīPīP°PSPSCūPj
PīP>P°CīPūP±Ps (P°PSP°P>C-P· CīPīPsCīC,PūCḡPūPḡPūPSPēC... PīPēPīP°PrPeC-PI PrP>Cū
PsPīPēC,CīPīP°PSPSCū PeP>C-PSC-CīPēCīC,P° PSP° PsCīPSPsPīC- P·PjC-PS P·P°
PīPsPeP°P·PSPēPeP°PjPē CIBIC-plus (p = 0,025); P·P° CēPeP°P>PsCḡ ADCS-ADLsev (p = 0,003)
C,P° PīPsPeP°P·PSPēPeP°PjPē SIB (p = 0,002)).
PḡCīPSPsPIPSḡPū PjPsPSPsC,PūCḡP°PīPūPīC,PēCāPSPū PrPsCīP>C-PrPḡPūPSPSCū
PjPūPjP°PSC,PēPSCf PīCḡPē P>C-PeCīPīP°PSPSC- C...PīPsCḡPsP±Pē PḡP>CḡCīPīPūPḡPjPūCḡP°
P>PūPīPePsC- C,P° CīPūCḡPūPrPSCḡPsC- C,CūPḡPePsCīC,C- (P·P°PīP°P>CḡPSPēPḡ P±P°P> MMSE
P·P° PīPēC...C-PrPSPēPjPē PīPsPeP°P·PSPēPeP°PjPē PīC-Pr 10 PrPs 22) PīPeP>CḡCīP°P>Ps 403
PīP°CīC-C”PSC,C-PI. PūP°CīC-C”PSC,Pē, CūPeC- PsC,CḡPēPjCīPīP°P>Pē P>C-PeCīPīP°PSPSCū
PjPūPjP°PSC,PēPSPsPj, PīPsPeP°P·P°P>Pē CīC,P°C,PēCīC,PēCāPSPs P·PSP°CāPSPs PēCḡP°CḡPēPḡ
PūC,PūPeC,, PSC-Pḡ PīP°CīC-C”PSC,Pē, CūPeC- PīCḡPēPḡPjP°P>Pē PīP>P°CīPūP±Ps, P·P°
PīPūCḡPīPēPSPSPēPjPē PeC-PSCāPūPīPēPjPē PīPsPeP°P·PSPēPeP°PjPē P·P° CēPeP°P>PsCḡ ADAS-
cog (p = 0,003), P·P° CIBIS-plus (p = 0,004) PSP° 24 C,PēPīPSC- (P·P° LOCF). P’
C-PSCēPsPjCf PrPsCīP>C-PrPḡPūPSPSC- PjPsPSPsC,PūCḡP°PīC-C— PīCḡPē C...PīPsCḡPsP±C-
PḡP>CḡCīPīPūPḡPjPūCḡP° P>PūPīPePsPīPs C,P° PīPsPjC-CḡPSPsPīPs CīC,CīPīPūPSCū
P·P°PīP°P>PsPj P±CīP>Ps CḡP°PSPPrPsPjC-P·PSPIP°PSPs 470 PīP°CīC-C”PSC,C-PI
(P·P°PīP°P>CḡPSPēPḡ P±P°P> MMSE P·P° PīPsCāP°C,PēPsPīPēPjPē PīPsPeP°P·PSPēPeP°PjPē
1ḡḡḡ”23). Pj PīPūCḡCīPīPūPeC,PēPIPSPs PīPēP·PSP°CāPūPSPsPjCf PīPūCḡPīPēPSPSPsPjCf
P°PSP°P>C-P·C- CīC,P°C,PēCīC,PēCāPSP° P·PSP°CāPēPjC-CīC,Cḡ PSḡPū P±CīP>P°
PrPsCīCūPīPSCīC,P° PI PeC-PSCāPūPīC-Pḡ C,PsCāCīC- PīPūCḡPīPēPSPSPsC-
PūC,PūPeC,PēPIPSPsCīC,C- PSP° 24 C,PēPḡPSC-
B PḡPūC,P°P°PSP°P>C-P· PīP°CīC-C”PSC,C-PI P·C...PīPsCḡPsP±PsCḡ PḡP>CḡCīPīPūPḡPjPūCḡP°
CīPūCḡPūPrPSCḡPsPīPs C,P° C,CūPḡPePsPīPs CīC,CīPīPūPSCū (P·P°PīP°P>CḡPSPēPḡ P±P°P> MMSE < 20)
P· CēPūCīC,Pē C,P°P· III PīP>P°CīPūP±Ps-PePsPSC,CḡPsP>CḡPsPīP°PSPēC... 6-PjC-CīCūCāPSPēC...
PrPsCīP>C-PrPḡPūPSCḡ (PīPeP>CḡCāP°CḡCāPē PrPsCīP>C-PrPḡPūPSPSCū PjPsPSPsC,PūCḡP°PīC-C— C,P°
PrPsCīP>C-PrPḡPūPSPSCū PīP°CīC-C”PSC,C-PI, CūPeC- PīCḡPēPḡPjP°P>Pē CīC,P°P±C-P>CḡPSCf PrPsP·Cf
C-PSPIC-P±C-C,P,CḡC-PI P°CīPūC,PēP>C...PsP>C-PSḡPūCīC,PūCḡP°P·Pē) PīPsPeP°P·P°P>Pē
CīC,P°C,PēCīC,PēCāPSPs P±C-P>CḡCē P·PSP°CāPēPjPēPḡ PūC,PūPeC, P>C-PeCīPīP°PSPSCū
PjPūPjP°PSC,PēPSPsPj C%PpsPrPs PePsPīPSC-C,PēPIPSPēC..., P·P°PīP°P>CḡPSPēC... C,P°
C,CīPSPeCīC-PsPSP°P>CḡPSPēC... CīC,PūCḡ.
PūCḡPē PīPēCūPIP>PūPSPSC- Cī PīP°CīC-C”PSC,P° CīCīPīCīC,PSCHsPīPs PīPsPīC-CḡCēPūPSPSCū Pī
CīCīC-C... C,CḡCHsPSC... CīC,PūCḡP°C... CḡPūP·CīP>CḡC,P°C,Pē PrPūPjPsPSCīC,CḡCīPīP°P>Pē
CīC,P°C,PēCīC,PēCāPSPs P·PSP°CāPēPjPēPḡ PūC,PūPeC, PjPūPjP°PSC,PēPSCf Pī PīPsPīPūCḡPūPrPḡPūPSPSC-
PīPsPīC-CḡCēPūPSPSCū Cī PīP°CīC-C”PSC,C-PI PīCḡCīPīPē PīP>P°CīPūP±Ps PīPsPīC-CḡCēPūPSPSCū Pī
CīCīC-C... C,CḡCHsPSC... CīC,PūCḡP°C... CīPīPsCīC,PūCḡC-PīP°P>PsCīCū PīPrPīC-CāC- CāP°CīC,C-CēPū,
PSC-Pḡ Cī C...PīPsCḡPēC..., C%Pps PsC,CḡPēPjCīPīP°P>Pē PjPūPjP°PSC,PēPS (21 % PīCḡPsC,Pē 11 %,
p < 0,0001).

PaP°CḡPjP°PePsPeC-PSPūC,PēPeP°.

ḡP>P±CīPpsCḡP>CīC-Cū

PḡP>CīPpsP>CḡC,PSP° P±C-PsPrPsCīC,CīPīPSC-CīC,Cḡ PjPūPjP°PSC,PēPSCf CīC,P°PSPsPīPēC,Cḡ
PīCḡPēP±P>PēP·PSPs 100 % CāP°CīC PrPsCīCūPīPSPSPSCū PīC-PeP° PePsPSCīPūPSC,CḡP°CīC-C— Cf
PīP>P°P·PjC- PeCḡPsPīC- (T_{max}) 5ḡ” PīC-Pr 3 PrPs 8 PīPsPrPēPS. PḡP·PSP°Pe PīPīP°PēPICf PīCḡPēPḡPsPjCf
C—PḡC- PSP° PīCīPjPsPeC,CīPīP°PSPSCū PSḡPūPjP°C”.

P PsP·PīPsPrC-P>

P°PsP±PsPīP° PrPsP·P° 20 PjPī P·CfPjPsPīP>CḡC” CīC,P°P±C-P>CḡPSCf PePsPSCāPūPSC,CḡP°CīC-Cḡ
PjPūPjP°PSC,PēPSCf Cf PīP>P°P·PjC- PeCḡPsPīC- Cf PjPūPḡP°C... PīC-Pr 70 PrPs 150 PSPI/PjP° (0,5ḡḡ”1
PjPePjPs>Cḡ) P·C- P·PSP°CāPSPēPjPē C-PSPrPēPIC-PrCīP°P>CḡPSPēPjPē PīP°CḡC-P°CīC-CūPjPē. PūCḡPē
P·P°CīC,PsCīCīPīP°PSPSC- PrPsP±PsPīPēC... PrPsP· PīC-Pr 5 PrPs 30 PjPī PīC-PrPSPsCēPūPSPSCū
PīPjC-CīC,Cf PīCḡPūPīP°CḡP°C,Cf Cf CīPūCḡPūP±CḡPsCīPīC-PSP°P>CḡPSC-Pḡ CḡC-PrPēPSC- C,P°
CīPēCḡPsPīP°C,CīC- PeCḡPsPīC- CīC,P°PSPsPīPēC,Cḡ 0,52. PḡP±5ḡḡ” C”Pj CḡPsP·PīPsPrC-P>CīC
CīC,P°PSPsPīPēC,Cḡ P±P>PēP·CḡPePs 10 P>/PePī. PūCḡPēP±P>PēP·PSPs 45 % PjPūPjP°PSC,PēPSCf
P·Pḡḡ” CūP·CīC”C,CḡCīCū P· PīCḡPsC,PūC—PSP°PjPē PīP>P°P·PjPē PeCḡPsPīC-.

P’C-PsC,CḡP°PSCīC,PsCḡPjP°CīC-Cū

P’ PsCḡPīP°PSC-P·PjC- P>CḡPPrPēPSPē P±P>PēP·CḡPePs 80 % PjPūPjP°PSC,PēPSCf CīPēCḡPeCīP>CḡC” Cf
PīPēPjP·CūPīC- PīPsCḡP°C,PēPsPīPsC— CḡPūCāPpsPīPēPSPē, PsCīPSPsPīPSC- PjPūC,P°P±PsP>C-C,Pē PSḡPū
PjP>CḡC,Cḡ NMDA-P°PSC,P°PīPsPSC-CīC,PēCāPSPēC... PīP>P°CīC,PēPīPsCīC,PūPḡ. PjCāP°CīC,C-
CīPēC,PsC...CḡPsPjCīP 450 Cf PjPūC,P°P±PsP>C-P·PjC- in vitro PSḡPū PīPēCūPIP>PūPSPs.
Pj PrPsCīP>C-PrPḡPūPSPSC- PīCḡPē PīPūCḡPsCḡP°P>CḡPSPsPjCf PīPīPūPrPūPSPSC- 14C-
PjPūPjP°PSC,PēPSCf Pī CīPūCḡPūPrPSCḡPsPjCf 84 % PrPsP·Pē P±CīP>Ps PūP>C-PjC-PSPsPīP°PSPs
PīCḡPsC,CūPīPsPj 20 PrPSC-PI, P±C-P>CḡCēPū 99 % PrPsP·Pē PūPēCīPēCḡPūC,CīPīP°P>PsCīCū
PSPēCḡPeP°PjPē.

P·P>C-PjC-PSP°CīC-Cū

PḡPūPjP°PSC,PēPS PūP>C-PjC-PSCf”C,CḡCīCū PjPsPSPsPūPeCīPīPsPSḡPūPSCīC-P°P>CḡPSPēPj CāPēPSPsPj
P· PīCḡPsPjC-PḡPePsPj t_{1/2} PīC-Pr 60 PrPs 100 PīPsPrPēPS. Pj PrPsP±CḡPsPīPsP>CḡCīC-PI P·
PSPsCḡPjP°P>CḡPSPsPīC,CīPSPeCīC-C”Cḡ PSPēCḡPsPe P·P°PīP°P>CḡPSPēPḡ PeP>C-CḡPūPSCf (CL_{ss})
CīC,P°PSPsPīPēC,Cḡ 170 PjP>/C...PI/1,73Pj”C,P° CāP°CīC,PēPSP° P·P°PīP°P>CḡPSPsPīPs PSPēCḡPePsPīPsPīPs
PeP>C-CḡPūPSCfCf PrPsCīCūPīP°C”C,CḡCīCū P·P° CḡP°C...CīPSPsPe PeP°PSP°P>CḡCīPūPīPsC—
CīPūPeCḡPūCīC-C—, PēPēCḡPePsPīP° CīC,P°PrC-Cū C,P°CḡPjP°PePsPeC-PSḡPūC,PēPePē PjPūPjP°PSC,PēPSCf
PīPeP>CḡCāP°C” C,P°PePsPḡ PeP°PSP°P>CḡCīPūPīCf CḡPūP°P±CīPpsCḡP±CīC-Cḡ CūPeP°, PjPsPḡP>PēPīPs,
PīPsPīCīPūCḡPūPrPePsPīP°PSP° PeP°C,C-PsPSPSPsCḡ C,CḡP°PSCīPīPsCḡC,PSPsCḡ CīPēCīC,PūPjPsCḡ.
PēPīPēPPrPeC-CīC,Cḡ PSPēCḡPePsPīPsC— PūP>C-PjC-PSP°CīC-C— PjPūPjP°PSC,PēPSCf Pī CīPjPsPīP°C...
P·CīPīPSPsC— CḡPūP°PeCīC-C— CīPūCāC- PjPsPḡPū P·PSPēPḡCīPīP°C,PēCīCū Cf 7ḡḡ”9 CḡP°P·C-PI.
PūC-PrP>CīPīPsPīCīPīP°PSPSCū CīPūCāC- PjPsPḡPū PīC-PrP±CīPīP°C,PēCīCū Cf CḡPūP·CīP>CḡC,P°C,C-
PīP>PēP±PsPePēC... P·PjC-PS PrC-C”C,Pē, PSP°PīCḡPēPeP>P°Pr P·PjC-PSPē P±P°PīP°C,PsPīPs
Pjḡḡ” CūCīPSPēPjPē CīC,CḡP°PīP°PjPē CḡP°CīC-PsPSCf PīPūPīPūC,P°CḡC-P°PSCīCḡPēPēPj CāPē
PīPSP°CīP>C-PrPsPe C-PSC,PūPSCīPēPīPSPsPīPs PīCḡPēPḡPsPjCf P°PSC,P°CīPēPrPSPēC...
CēP>CīPSPePsPīPēC... P·P°CīPpsP±C-PI.

P·C-PSC-PḡPSC-CīC,Cḡ

P—P° PrP°PSPēPjPē PrPsCīP>C-PrPḡPūPSCḡ PSP° PrPsP±CḡPsPīPsP>CḡCāCūC...,
C,P°CḡPjP°PePsPeC-PSPūC,PēPēP° PjPūPjP°PSC,PēPSCf PjP°C” P>C-PSC-PḡPSPēPḡ C...P°CḡP°PeC,PūCḡ Cf
PrC-P°PīP°P·PSPSC- PrPsP· 10ḡḡ”40 PjPī.

PaP°CḡPjP°PePsFrPēPSP°PjC-CāPSPēPḡ/C,,P°CḡPjP°PePsPeC-PSPūC,PēCāPSPēPḡ P·Pḡḡḡ” CūP-PsPe
PūCḡPē PrPsP·C- PjPūPjP°PSC,PēPSCf 20 PjPī PSP° PrPsP±Cf PḡPsPīPs PīPjC-CīC, Cf
CīPūCḡPūP±CḡPsCīPīC-PSP°P>CḡPSC-Pḡ CḡC-PrPēPSC- PīC-PrPīPsPīC-PrP°C” PīPūP>PēCāPēPSC- k₁
(PePsPSCīC,P°PSC,P° PīP°P>CḡPjCfPīP°PSPSCū) PjPūPjP°PSC,PēPSCf, C%Pps CīC,P°PSPsPīPēC,Cḡ 0,5
PjPePjPs>Cḡ Cf PrC-P>CūPSCīC- C,CḡPsPSC,P°P>CḡPSPSC— PePsCḡPē PīPsP>PsPīPSPsPīPs PjPsP·PeCf
P>CḡPPrPēPSPē.

PfPIPsC_HPsP±P° PhP»CHC†PiPuPN:PjPuC_HP° PIC-Pr C†PuC_HPuPrPSC_HPsPiPs C†C,CrPiPuPSC_H
C,CUP¶PePsC†C,C- PrPs C,CUP¶PePëC... C,,PSc_HPj.

P⁺C-PiPmCfjCzCrC,P»PëPIC-CfC,CH PrPs PrC-CHCzPsC— CfjPmCzPsPIPëPSPë P°±Ps PrPs P±CrPrCh.
CljPePsPiPs PePsPjPiPsPSPmPSC,P° PiCfjPmPiP°CfjP°C,Cr.

P*CPpP, PpPSPiC-PrPSPiPp, PpPSPSPiC PiCpPpPSPeCpPpPpC- PiPpPiPpPrPePpP- PiC-PrPiPpPpCpPpPpPSPSPiC
PpP-PrPSPpCpPpPPrPSPiPpPSPSPCpPpPpP- C-PpPiPpPSPpSPiPpPpP CPrPiC- PiPpP-PrPSPpCpPpPSPSPiC (PpPpPpY)
PiCpPpP P-P*CPiC, PpSPCpPiPpPSPSPC- PpPpPiPpP*PSP, PpPSPpC PiP*CPiC-C*PSPC, PpPi, CUpPpC- PiCpPpPpPpPpP*CPiC, CbP
PiPpP*CPiC, C*CPpPpPpP, PiPpSPpCpPpPSPpSPpPpP-PpPiPpPiP CUpPpPpP PSPpPi PiCpCpP, PpPSPpPiPpPpPpPpPpPpPpP,
PSPpPiPpSPpPpC... C-PrPSPpPpPpP CbPpPpC, PpPpP*CbPSPpPpPpP PiPpSPSPC-C*CbPSPpPSP PiCpPpPpC, CbPpPpPiPpPpPpPpPpPpPpPpPpP
CpPpP*CPiCpP CPrPpPpPpPpP CPr PiPpP*CPiC-C*PSPC, C-Pi, CUpPpC- PpSPrPSPpSPpCpP*CPiPSPSP PiCpPpPpPpPpP*CPiC, CbP
PiPpPpCpPpPpPpP*CbPSPC- P*PSPC, PpPpPpPpP*PiCpPpC*CUlPSP, PpP.
PpP C...PpSPrC- C, P*CPpPpPiPpPpPpPpPpC-PSPpPiC, CbPpPSPpPpC... PrPSPpCpP C-PrPiPpPpPSPiCpP CPr C-PrPrPpPpPSPiPpC...
PrPSPpCpPSPiPpSPpP*CbPpPiC-PiPiCPiCpC, C*PiPpC... PpPpC, PiPpPpC, C-PiPiPpPpC* PpPpPpPpC C- PpPpPiPpP*PSP, PpSPpC PiP
P- PiPpC-PpCfCpPpPpPrPSPpPi/ PpPpC, C, PpSPpPiC-PSPpPiP, PrPrPSPpPiPpPiPpPpPpPpPiP P*PpPp PiP*PpP*PSPC, PpPiPpC-PSPpPiP
PSPpPiPiPpCUPiPpPpPpP.
PrPSPpPiPpP*PSP, PpPp *in vitro* PSPpPi C* C-PSPiC-PpP-C, PpSPpSPpPi CYP 1A2, 2A6, 2C9, 2D6, 2E1, 3A,
C, PpP*PiC-PSPiPiC-CPiPSPpC- PpPpSPSPpSPpPpCfPpPiPpPiPSP*PpP, PpPiPSPpSPpPpPpPiC-PrCPpPpPpP*PpPpP PpPpPpP
SPpPpP*CbPpC, C-P, C-PpSPpCf.

[illegible][illegible]

[illegible]

PYPeCfC, P _u PjP ^o , PsC ₇ PiP ^o PS, PeP _o PCf	PSP ^o CfC, PsC, P ^o	PuPsP±C-CfPSC- C ₇ PuP ^o PeCfC-C-
PfPSC _u , P _u PiPeCfC-C-	PkP _u CfP ^o CfC, Ps	P ^o C ₇ P _u P±PePsPIC-P ^o PC...PiPsC ₇ C ₆ PiP ^o PSPSC _u
PuPsC ₇ CfC ₆ PuPSPSC _u C-PjCfPSPSPsC- CfPeCfC, P _u PjP ₆	PSP ^o CfC, Ps	P ^o C-PiP _u C ₇ CfCfC, P _o PePIC-CfC, C ₇ H
PuCTPeC...C-CfPSC- PiPsC ₇ CfC ₆ PuPSPSC _u	PSP ^o CfC, Ps PkP _u CfP ^o CfC, Ps P ^o PC ₇ CfCfCfCfC, Ps PkP _u CfP ^o CfC, Ps	PYPsPSP _o PePIC-CfC, C ₇ H PYPiP _o CfC, P ^o PSC-CfC, C ₇ H CfPIC-PrPsPiPsCfC, C- P ^o PC ₇ CfCfCfCfCfC-C- ¹ PkCTPeC...PsC, PeCfPSC- C ₇ PuP ^o PeCfC-C- ²
PuPsC ₇ CfC ₆ PuPSPSC _u PSP _u C ₇ PiPsPiPsC- CfPeCfC, P _u PjP ₆	PSP ^o CfC, Ps PSP ^o CfC, Ps PkP _u CfP ^o CfC, Ps P ^o CfP _u P _u C ₇ C-PrPeC-CfPSC-	P-P ^o PiP ^o PiPsC ₇ P _u PSPSC _u PuPsC ₇ CfC ₆ PuPSPSC _u C ₇ C-PiPSPsPiP ^o PiP ₆ PuPsC ₇ CfC ₆ PuPSPSC _u C...PsPrP ₆ PVCfPrPsPiPSC- PSP ^o PiP ^o PrP ₆
PYPuP _u CfCfP _u PiC- PiPsC ₇ CfC ₆ PuPSPSC _u PYCfPrP ₆ PSPSC- PiPsC ₇ CfC ₆ PuPSPSC _u	PkP _u CfP ^o CfC, Ps PSP ^o CfC, Ps PSP ^o CfC, Ps PkP _u CfP ^o CfC, Ps	PYPuP _u CfCfP _u PiP ^o PSPuPrPsCfC, P ^o PC, PSC-CfC, C ₇ H P ₇ C ₇ C, PuCfC- P ^o PC ₇ C ₇ HSP ^o PiC-PiP _u C ₇ C, PuPSP-C-C _u P ^o PiPSPsP-PSPeP _u C, C ₇ PSPiP _u ±PsP _u /C, C ₇ PSPiP _u ±PsPuP _u ±PsP _u PC-P _u P _u
PuPsC ₇ CfC ₆ PuPSPSC _u PrPeC...P ^o PC ₇ C ₇ HSPSC- CfPeCfC, P _u PjP ₆	PSP ^o CfC, Ps	P-P ^o PrP ₆ C ₆ PeP ^o
PEP _o CfPSPePsPiPs- PiPsC ₇ CfC ₆ PuPSPSC _u	PSP ^o CfC, Ps PkP _u CfP ^o CfC, Ps PkP _u PiP ₆ PSP ^o PC ₇ PuPSC-	P-P ^o PiPsC ₇ P ^o PC ₇ PiP ^o PSPSC _u PuP ^o PSPeC ₇ PuP ^o PC, PeC, ⁷
PuPsC ₇ CfC ₆ PuPSPSC _u P _u P±PsPeCf PiP _u CfC-PSPeP ₆ C, P ^o PiPsPiCfPSPiP _u PiC-PrPSP ₆ C... C ₆ P _u C _u ...C-Pi	PSP ^o CfC, Ps PkP _u CfP ^o CfC, Ps PkP _u PiP ₆ PSP ^o PC ₇ PuPSC-	PuC-PrPiP ₆ C%PuPSPSC _u PiPsPeP ^o P-PSPeP ₆ -Pi C _u CfPSPeCfC-C- PiP _u CfC-PSPeP ₆ P ^o PuP _u PiP ₆ PSP ^o PC ₇ PuPSC-
P-P ^o PiP ^o PC ₇ HSPSC- PiPsC ₇ CfC ₆ PuPSPSC _u	PSP ^o CfC, Ps PkP _u CfP ^o CfC, Ps	P ^o PC ₇ PSPiPSP ₆ PeP _u PC-P _u C-P _u C ₇ H PuC-PrPiP ₆ C%PuPSP ^o PiC, PsPiP _u PC ₇ PiP ^o PSC-CfC, C ₇ H

$$^2\text{PhPeCfPuPiC}-\text{PiPsPIC}-\text{PrPsPiP}\gg\text{PuPSPSCII PiCfPe PiPuPrPeC}\pm\text{PSPsPiCf P}\cdot\text{P}^\circ\text{CfC.PsCfCfPIP}^\circ\text{PSPSC}-$$

PvPμCHPiC-PS PiCHPePrP°C,PSPsCfC,C-, 4 CHPsPePe.

PJpJPsPIPë P·P±PµCḡC-PiP°PSPSCII.

PŸPŸPµC†C-P°P»CHbPSPëC... CfPjPsPI P·P±PµCḡC-PiP°PSPSCII PSPµ PŸPsC,CḡPµP±C†C".
P—P±PµCḡC-PiP°C,Pë Cf PSPµPrPsC†C,C†PŸPSPsPjC† PrP»CII PrC-C,PµP№ PjC-C†C†C-.

PJPIP°PePsPIPeP°.

PµPs 14 C,P°P±P»PµC,PsPe Cf P±P»C-C†C,PµCḡP°C..., PŸPs 2 P±P»C-C†C,PµCḡPë Cf PeP°CḡC,PsPSPSC-P№
PePsCḡPsP±C†C-.

P¬P°C,PµPiPsCḡC-CII PIC-PrPiC†C†PeC†. P—P° CḡPµC†PµPŸC,PsPj.

P°PëCḡPsP±PSPëPe.

PI. P»C†PSPPrP±PµPe Pḡ/PŸ / H. Lundbeck A/S.

P¬C-C†C†PµP·PSP°C...PsPrP¶PµPSPSCII PIPëCḡPsP±PSPëPeP° C,P° P°PrCḡPµC†P° PjC-C†C†CII

PiCḡPsPIP°PrP¶PµPSPSCII P№PsPiPs PrC-CIIP»CHbPSPsC†C,C-.

PbC,C,C-P»C-P°PIPµP№ 9, 2500 P°P°P»P±C-, P°P°PSC-CII / Ottiliavej 9, 2500 Valby, Denmark.