

PRILOG I.

UCfGVCM"QRKUC"UXQLUVXC"NKLGM

Lijek koji više nije odobren

Qxcl"lg"nklgm"rqf"fqfcvpk o "rtc gplg o 0"Vko g"ug"qoqi w wlg"dt|q"qvmtkxcpkg"pqxkj"uki wtpqupkj"
kphqt o cekl c 0"Qf" | ftexuvxgpkj"tcfpkmc"ug"vtfkl"fc"rtklcxg"uxcmw"uw o plw"pc"pwurqlcxw" | c"qxcl"nklgm 0" \ c"
rquvwrem"rtklcxnlkxepcl"pwurqlcx"xxflgvk"fkq 60: 0

1. PC \ KX"NKLGMC

P [ZV J TCEKU 100 o i l o n"mqpepvtcv" | c"qvqrkpw" | c"kphw | klw

2. MXCNKVCVKXPK"K"MXCPVKVCVKXPK"UCUVCX

Lgfc p" o n"mqpepvtcv"ucftflk"322 o i "qdknvqmucmuk o cdc 0"
Lgfpc"dq kec"qf"8 o n"ucftflk"822 o i "qdknvqmucmuk o cdc 0

Qdknvqmucmuk o cd"ug"rtqk | xqfk"vgj pqqi k lq o "tgmq o dkpcvpg" FPM"w"uvcpkc o c"ok-lgi" o klgq o c"
IU-PU20

Rq o q pc"vxct"u"rq | pcvk o "w kpmq o

Jedan ml koncentravc"ucftflk"58 o i "uqtdkvqnc 0

\ c"elgnqxkvk"rqrku"rq o q pkj"vxctk"xxflgvk"fkq 8030

3. HCTOCEGWUMK"QDNKM

Mqpepvtcv" | c"qvqrkpw" | c"kphw | klw"*uvgtkpk"mqpepvtcv+0"

P [ZV J TCEKU lg"dkvct"fq"qrcnguegpvc."dg | dqfpc"fq"dnklgfqflwvc"fq"dnklgfq"u o g mcuvqflwvc"qvqrkpc"
mqlc" o qlg"ucftflcxvk" o c nq"rtq | ktpkj"fq"dklgkj"dlgnep gxpvcvkj" guvkec"*mqlg" g"ug"wmnqpkvk"nkpklumk o "
hknvtkteplg o +u"r J"7.7"ku o qncnpq- w"qf"499- 52: o Qu o lmi 0

4. MNKPK MK"RQFCVEK

4.1 Vgtrklung"kp fkmceklg

P [ZV J TCEKU lg"kp fkektep"w"mq o dkpceklk"u"qf i qxctclw k o "cpvkdcmvgtklumk o "nklgmqxo c" w"uxkj" fqdpkj"
umwrkpc" | c"nklg gplg"kp j cneklumqi"cpvtcmuc" | dq i "dcmvgtklg" *Bacillus anthracis* *xxflgvk"fkq 703+0

P [ZV J TCEKU lg"kp fkektep"w"uxkj" fqdpkj"umwrkpc" | c"rtqhkncmuw"pcmqp" k | nc i cplc"kp j cneklumq o "
cpvtcmuw"mefc"ftw ig vgtrklg"pkuw"qf i qxctclw g"knk"pkuw"fqvwrpg"*xxflgvk"fkq 703+0

4.2 Fq|kteplg"pc kp"rtk o lgpg

P [ZV J TCEKU vtgdc"fcvk" k o "lg"vq"mnkpk mk"kp fkektecpq 0

Qf i qxctclw g" o g fkekpumq"nklg gplg" k"pc f | qt"wxklgm"vtgdclw"dkvk"fqvwrpk" w"unw clw"cpchknmcvk mq i "
fq i c clc"pcmqp"rtk o lgpg"nklgmc" P [ZV J TCEKU 0

Fq|kteplg

Rtgrqt w gpq"fq |kteplg"nklgmc" P [ZV J TCEKU w"qftcunkj"dqngupkmc"vgflkpg"qf"pcl o cplg"62 mi "lg"lgfpc"
kpvtxgpumc" kphw | klc"qf"38 o i lmi"vlngupg"vgflkpg 0" Rtgrqt w gpq"fq |kteplg"nklgmc" P [ZV J TCEKU w"
qftcunkj"dqngupkmc"vgflkpg" o cplg"qf"62 mi "lg"lgfpc"kvtxgpumc" kphw | klc"qf"46 o i lmi"vlngupg"vgflkpg 0"

Prije primjene lijeka P [ZVJTCEKU rtgrqtw wlg"ug"rtg o g fkmceklc"cpvkjkuvc o kpkmq o "*xkflgvk" fklgnqxg" 606"60: +0"

\c"rtknciqfdg"fq|g"w"unw clw"tgemeklc"rqxg|cpkj"u"kphw|klq o "xkflgvk"vcdnkew 30

Vcdnkew 30<"Rtknciqfdg"fq|g qdknvqmucmuk o cdc w"unw clw"tgemeklc"rqxg|cpkj"u"kphw|klq o

Vgflkpc"tgemeklc" r q x g c p k j " u " k p h w k l q o "	Rtknciqfdc"fq g
Tgemeklc" r q x g c p c " u " k p h w k l q o 30 - 50 uvwrnja	Kphw klw"qdknvqmucmuk o cdc"rqvtgdpq"lg"privremeno rtgmkpwwk"r"rtwflkvk"rqvrtqpw" vgtcrklw0 Rtk"rtxql"rqlcxk"rkumepc."dtqppjqure o c"knk"i gpgtcnk ktepg"wtvkmctklg"50 uvwrplc." nklg gplg"qdknvqmucmuk o cdq o "rqvtgdpq"lg"vtclpq"rtgmkpwwk0 Rtk"rqpxnlclw g o "rkumepw"knk"wtvkmctklk"40 uvwrplc"knk"rtk"rqpxnlcplw"dknq"mqkij" uk o rvq o c"50 uvwrplc."nklg gplg"qdknvqmucmuk o cdq o "rqvtgdpq"lg"vtclpq"rtgmkpwwk0" W"uwrtqvpq o ."pcmq"rqvwpqi"rxnc gplc"uk o rvq o c."kphw klc"ug" o qflg"pcuvcxkvk" u"72 % dt kpg"rqvki pwwg"rtklg"rtgmfc0" Cmq"pg o c"uk o rvq o c"rqxg cpkj"u" kphw klq o ."dt kpc"kphw klg"qrkucpc"lg"w"vcdnkew"50" Rqvtgdpq"lg"rtk o klgpvk"rtg o g fkmceklw0
Tgemeklc" r q x g c p c " u " k p h w k l q o 60 uvwrplc	Kphw klw"qdknvqmucmuk o cdc"rqvtgdpq"lg"qf o e j"rtgmkpwwk0 Rqvtgdpq"lg"rtwflkvk"rqvrtqpw"nklg gplg" Nklg gplg"qdknvqmucmuk o cdq o "rqvtgdpq"lg"vtclpq"rtgmkpwwk0"

Rqugdpg"rqrwneclg

Starije osobe

Pklg"rqvtgdpq"rtknciq cxcplg"fq|g"|"c"dqngupkmg"× 87 iqfkcpc"*xkflgvk"fkq"704+0

Pedijatrijska populacija

Rtgrqtw gpc"fq|c|"c"rgfklcvtklumg"dqngupkmg"vg o gnk"ug"pc"vgflkpk"mcq"-vq"lg"rtkmc|cpq"w"vcdnkew 40"w" pcuvcxmw0

Vcdnkew 40<"Rtgrqtw gpc"rgfklcvtklumc"fq|c"qdknvqmucmuk o cdc"*fq|kteplg"pc"vg o gnw"vgflkpg-

Vlgnupc"vgflkpc"jmi _	Fq c"j o i l m i " vlgnupg"vgflkpg
> 40	38
@37"fq"62	46
37"knk" o cplg	54

Pc kp"rtk o lpgg"

Qdknvqmucmuk o cd"ug" o qtc" fxcvkc"kpvtcxgpumq o "kphw|klq o "vklgmq o "; 2 o kpwvc0

Mjere opreza koje je potrebno poduzeti prije rukovanja ili primjene lijeka

Dq kec"ug"pg"u o klg"vtguvk0"Qdknvqmucmuk o cd"ug" o qtc"tc|tklgfkvk"w"qvqrkpk"pcvtklgxqi"mnqtkfc|"c"injekcije qf"; o i l o n"*2.; ' + "rtklg"rtk o lpgg"w"qdknw"kpvtcxgpumg"kphw|klg"*xkflgvk"fkq"808+0"

Tc|tklg gpc"kpvtcxgpumc"kphw|klc"qdknvqmucmuk o cdc" o qtc"ug" fxcvkc"vklgmq o "; 2 o kpwvc"dt|kpg o "kphw|klg" qrkucpq o "w"vcdnkew 50."u"rqo q w"xtg keg"knk"-vtecnlmg"|"c"kphw|klw"vg"nkpklumqi"hnkvtc"qf"2.44 o kmtqpc0"

Dqngupkmg"lg"rqvtgdpq"rcfnlkxq"pcf|ktevk"|dq i"rqlexg"|pcmqxc"uk o rvq o c"rtgqulgvlkxquvk"vklgmq o" kphw|klg"pcl o cplg"lg fcp"ucv"pcmqp"rtk o lpgg"*xkflgk"fkq"606+0" Pc"tgcmeklg"rqxg|cpg"u"kphw|klq o" rqtgdpq"lg"tgc iktevk"mcq"-vq"lg"pcxgfgpq"w"vcdnek 30

Nkplw"vtgdc"kurtevk"qvqrkpo"pcvtklxqi"mnqtkfc"|c"plgmeklg"qf"; o ilon"2.; ' + "rq|cxt-gvmw" kpvtxgpung"kphw|klg0

Vcdnek"50<"Fq|c"qdknvqmucmuk o cdc."wmwrpk"xqmw o gp"kphw|klg"dt|kpc"kphw|klg"rtg o c"vlngupql"vgflkpk

Vlmgupc"vgflkpc"mi_ *fq ktcplg"pc"vg o gnlw"vgflkpg+	Wmwrpk"xqmw o gp"kphw klg"]on_]xtg kec"knk"-vteclmc" c" kphw klw_ ,	Dt kpc"kphw klg]onlj_
> 40 mi"knk"qftcuk"*38 o ilmi"vlngupg"vgflkpg+		
> 40	472	389
> 15 mi"fq"62 mi"*46 o ilmi"vlngupg"vgflkpg+		
qf"53"fq"62	472	389
qf"38"fq"52	100	89
15 mi"knk" o cplg"*54 o ilmi"vlngupg"vgflkpg+		
qf"33"fq"37	100	89
qf"7"fq"32	72	55.5
qf"5.3"fq"6.;	47	39
qf"4.3"fq"5	42	35.5
qf"3.3"fq"4	37	10
1 ili manje	9	6.9

, "Wrwvg"q"tc|tlg kxcplw"nklgmc"kw"rqvtgdk"xtg keg"knk"-vteclmg"|c"kphw|klw"rtklg"rtk o lpgg"xkflgk"w" fklgnw 8080

4.3 Mqpvtekp fkmceklg

Rtgqulgvlkxquv"pc"flgncvpw"vxct"knk"pgmw"qf"rq o q pkj"vxctk"pcxgfgpkj"w" fklgnw 8030

4.4 Rqgdpc"wrq|qtgplc"o lgtg"qrtg|c"rtk"wrqtcdk

Unlgfkxquv

Mcmq"dk"ug"rqdqnl-cnc"unlgfkxquv"dkqng-mkj"nklgmxc."pc|kx"kdqtl"ugtklg"rtk o klgpplgqi"nklgmc"rqvtgdpq"lg" lcupq"gxkfgpvktevk0

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Tgcmeklg"rqxg|cpg"u"kphw|klq o l reakcije rtgqulgvlkxquv" guvq"uw"ug"qrcflng"vklgmq o"mnkpk mkj" kurkvkxpcplc"qdknvqmucmuk o cdc"w"|ftexkj"kurkvkxpcplc" \ dq i"tk|kmc"qf"vg-mkj"tgcmeklc"knk"cpchknemuklg."pwflpq" lg"fc"quqdnlg"qurquqdnlgpq"q"rtg o nlgpq"|c"nklg gplg"cpchknemuklg"rtk o lgpplwlg"qdknvqmucmuk o cd"w" pcf|ktevk"o"wxlgvk"o c0"Dqngupkmg"lg"rqvtgdpq"rcfnlkxq"pcf|ktevk"vklgmq o"tc|fqdnlc primjene kphw|klg"k" pcl o cplg"lg fcp"ucv"pcmqp"rtk o lpgg0

Dwfw k"fc"uw"mnkpk mc"kurkvkxpcplc"rtqxfgpc"pc"|ftexk o"fqdtqxqnek o c."kphw|klg"qdknvqmucmuk o cdc" |cwuvxnlgpg"uw"rtk"rqlexk"dknq"mcmxg"tgcmeklg0" Pc"vg o gnlw"kumwuvxc"u"ftwik o" o qpqmnqpmuk o" cpvkvlgknk o c"mqle"ug"rtk o lgpplw"nklg gplw"q|dknlpkj"|ftexuvxgpj"uveplc."kphw|klg"ug"qr gpkvq"oqi w" fqxt-kvk"cmq"ug"mqpvtnktclw"pc"qfiqxtclw k"pc kp0" Pc"tgcmeklg"rqxg|cpg"u"kphw|klq o"rqvtgdpq"lg" tgc iktevk"mcq"-vq"lg"pcxgfgpq"w"vcdnek 30"

Rtg o g fkmceklc"cpvkjkuvc o kpkmq o."prt0"fkhgpkjftc o kpk o."rtgrqtw wlg"ug"rtklg"rtk o lpgg" qdknvqmucmuk o cdc"*xkflgk"fkq"604+0" Fkhgpkjftc o kp"lg"rtk o klgpplg"52 o kpwvc"rtklg"nklg gplc" qdknvqmucmuk o cdq o"w"mnkpk mk o"kurkvkxpcplc"o c"rtqxfgpk o"u"qdknvqmucmuk o cdq o0"Rtg o g fkmceklc"

cpvkjkuvc o kpkmq o "pg"urtlg cxc"cpchknkmuklw" o qflg"rtkmtkvk"knk"qfiqfkvk"rqlcxw"uk o rvq o c"
rtgqulgvnlkxquvk0

Cpvtcmupk" o gpkpi kvku

Qdknvqmucmuk o cd"pg"rtgnc|k"mtxpq- o qffcpw"detklgtw"vg"pg"urtlg cxc"pkvk"nklg k"cpvtcmupk" o gpkpi kvku0"

Rgfklevtklumc" rqrwnceklc

Pklg"dknq"kurkvkxpcle"uk i wtpquvk"knk"het o cmqmkpgvkmw"qdknvqmucmuk o cdc" w"rgfklevtklumoj rqrwncekli *xkflgvk"
fkq"704+0

Kpvgtcmeklg"u ncdqtcvqtklumk o "vguvktanjima

K|nqflgpquv"nklgmw" P [ZVJTCEKU o qflg"q o gvcvk"ugtqnq-mg"vguvqxs"pc"cpvtcmu0

Uqtdkvqn

Jedan ml lijeka P [ZVJTCEKU ucftfk"58 o i "uqtdkvqnc"*xkflgvk"fklnqxs"40"803+0"
Nklgmqxl"mqll"ucftflg"uqtdkvqn" o q i w"dkvk"u o tvqpupk"cmq"ug"fcw"kpvtcxgpumk"quqdc o c"u"pcunlgfpg o "
pgvqngtcpeklq o "pc"htwmvq | w"*gpi n0" *hereditary fructose intolerance*. JHK+0"Qdknvqmucmuk o cd"ug"pg"u o klg"
rtk o lgpikxcvk" w"quqdc"u"pcunlgfpg o "pgvqngtcpeklq o "pc"htwmvq | w."qul o "cmq"rqvqqlk" | pcvpc"mnkpk mc"
rqvtgdc" k"pkw"fqvwrpg"cmvgtpevkxg0"Rtklg"fcxpcle"qxq i "nklgmc"rqvtgdpq"lg"uxcmq o "dqngupkmw" w | gvk"
fgvnlpw"cp o pg | w"u"qd | ktq o "pc"uk o rvq o g"pcunlgfpg"pgvqngtcpeklg"pc"htwmvq | w0

Fqlgp cf" k" o cnc" flgec" o nc c" qf"4 iqfkgp+ "rqvqngtcpeklq" u" k | nqflgp" tk | kmw" lgt" k o " o qffc" lq- "wxklgm" pklg"
fkli pquvkektcpc" pcunlgfpc"pgvqngtcpeklc"pc"htwmvq | w0

Natrij

Qxcl"nklgm"ucftfk" o cplg"qf"3 o o qn"*45 o i+ "pcvtklc"po dq kek"qf"8 ml."qfpqppq" | cpg o ctkxw"mqnk kpw"
pcvtklc0

4.5 Kpvgtcmeklg"u ftwik o "nklgmqxl o c" k" ftwik" qdnkek" kpvgtcmeklc

Ekrtqhnqmucekp

W"kurkvkxpcle"kpvgtcmeklc."rqlgfpc pc"fq|c"qdknvqmucmuk o cdc"primijenjena lg"uc o c"knk" kuvqfqpq"u"
ekrtqhnqmucekpq o "w"62"kurkvcpkmc0" Fxcfgugv"kurkvcpkmc" rtk o cnq"lg"uc o q"qdknvqmucmuk o cd."c"42"
kurkvcpkmc" rtk o cnq"lg"qdknvqmucmuk o cd k"ekrtqhnqmucekp"vklgmq o "; fcp0"Rtk o lgp"38" o ilmi"kpvtcxgpumg"
kphw | klg"qdknvqmucmuk o cdc" rtklg"kpvtcxgpumg" kphw | klg"ekrtqhnqmucekp"knk"ekrtqhnqmucekp" fxc" rwwc"
fpgxpq" rgtqtcnpk o "w" | k o cplg o "vcdngvc" pklg" rtk o klgpkic"het o cmqmkpgvkmw"qdknvqmucmuk o cdc0"Kuvq"vcmq."
qdknvqmucmuk o cd" pklg" rtk o klgpkq"het o cmqmkpgvkmw"ekrtqhnqmucekp" rtk o klgplgpq i " rgtqtcnpq"knk"
kpvtcxgpumk0

Pkuw"rtqxfpgpc"ftwic"kurkvkxpcle"kpvgtcmeklc0"Dwfw k"fc"lg"qdknvqmucmuk o cd" o qpqmnqppumq"cpvkvklgnq."
tk | km"qf"kpvgtcmeklg"lg"pk | cm0

608 Rnqfpquv.vtwfpg c" k" fqlgplg

Vtwfpg c

Pg o c"rqfcvcmc"q" rtk o lgpk"qdknvqmucmuk o cdc" w"vtwfpkcc." o g wwk o "rq|pcvq"lg"fc"nlwfumk Ki I "rtgnc | k"
mtq | "rqvgnlkew0
Kurkvkxpcle"pc"flkxqkplc o c"pg"wrw wlv"pc" k | texpg"knk"pgk | texpg"-vgvpg" w kpmg" w"rqingfw"tgtrqfwmwkxpg"
vqmuk pquvk"*xkflgvk"fkq"705+0
Mcq" o lgtc"qrtg | c."rtgrqtw wlg"ug" k | dlgiexcpplg" rtk o lgp"nklgmc" P [ZVJTCEKU vklgmq o "vtwfpq g0

Fqlgplg

Pklg"rq|pcvq"kl|nw wlg"nk"ug"qdknvqmucmuk o cd"w" o cl kpg"o nklgmq0"Rq|pcvq"lg"fc"ug"nlwfumk Ki I"kl|nw wlg"w" o cl kpg"o nklgmq"rtxkj"fcpc"pcmqp"rtqf"fc."c"wdt|q"pcmqp"vqic"u o cplwlg"ug"pc"pkumg"mqpegpvtceklg0" Unklg"fq o"vqic."pg"o qflg"ug"kumnlw kvk"tk|km|c"fqlgpw"flgew"vklgmq o"qxqi"mtcvmqi"tc|fqdnlc0" Pcmqp"vqic." wrqvtgdc"qdknvqmucmuk o cdc" o qic"dk"ug"tc|o qvtkvk"vklgmq o" fqlgplc"uc o q"cmq"lg"vq"mnkpk mk"rqvtgdpq0

Rnqfpquv

Kurkvkxcpk"rnqfpquv"pkuw"rtqxf"fgpc"u"qdknvqmucmuk o cdq o0

4.7 Wvlgecl"pc"urquqdpquv"wtexnlcplc"xq|knk o c"k"tcfc"uc"uvtqlgxo c

Qdknvqmucmuk o cd" o qflg"ko cvk" o cnk"wlgecl"pc"urquqdpquv"wtexnlcplc"xq|knk o c"k"tcfc"uc"uvtqlgxo c"lgt" ug"pcmqp"rtk o lpgg"nklgmc" P [ZVJTCEKU o qi w"rqlexkvk"incxqdqnlc."q o cinkec"wo qt"krqxte cplg" *xkflgkv"fkq"60: +0

4.8 Pwurqlcxg

Ucflgvcm"uki wtpquqji"rtqhknc

Uki wtpquv"qdknvqmucmuk o cdc"rtqw cxcpc"lg"uc o q" w" |ftexkj"qftcunkj"kurkvepkmc0

Uki wtpquv"qdknvqmucmuk o cdc"kurkvepc"lg"w"542" |ftexkj"kurkvepkmc"*w"fqdk"qf"3:"fq"9;"iqfkpc+"nklg gpkj" lgfppq o"knk"fxco c"kpvtexgpumk o"fq|co c"qf"38 o ilmi" w"vtk"mnkpk mc"kurkvkxcpk0

Wmwrpq"472"qf"542"kurkvepkmc"rtk o knq"lg"lgfpw"fq|w"qf"38 o ilmi"qdknvqmucmuk o cdc0" Pwurqlcxg" rqxg|cpg"u"rtgqulgvnkxq- w"*wmnlw wlw k"qukr+"rqlexkng"uw"ug"w"; ' "*441472+"vkj"kurkvepkmc."u"lgfpk o" unw clg o"cpchknemuklg"mqk"ug"fq i qfkq"vklgmq o" kphw|klg0" kphw|klc"lg"rtgmkpwvc" w"5 ' " *:1472+"kurkvepkmc" |dq i"rtgqulgvnkxquv"knk"cpchknemuklg0

Pcl g- "g"rtklcxnlpgg"pwurqlcxg"dkng"uw"incxqdqnlc"*6 ' .";1472+."rtwtkvwu"*6 ' .";1472+"k"wtvmctklc"*4 ' . " 81472+0

Pcl g- "g"wg gpg"pwurqlcxg" w"rtxc"vtk"ucvc"pcmqp"rq gvmc" kphw|klg"dkng"uw"rtwtkvwu"*p = 9="4.: ' +." wtvkmctklc"*p = 8="4.6 ' +." incxqdqnlc"*p = 6="3.8 ' +." qukr"*p = 5="3.4 ' +." mc-cn1"*p = 5="3.4 ' +." q o cinkec" *p = 5="3.4 ' +." *wmnlw wlg"q o cinkec" w"rqvwtcnpw"q o cinkec" w"0

Unlgfg g"vg-mg"pwurqlcxg"rqlexkng"uw"ug"vklgmq o"rtxc"vtk"ucvc"pcmqp" kphw|klg<"wtvmctklc"*p = 1= 2.6 ' +." rtwtkvwu"*p = 1= 2.6 ' +." "dqnxk" w"ng ko c"*p = 1= 2.6 ' +0

Pcl g- "g"wg gpc"pwurqlcxg"wpwvct"5"fq"46"ucvc"pcmqp"rq gvmc" kphw|klg"dknc"lg"incxqdqnlc"*p = 5=" 3.4 ' +0

Vednk pk"rqrkupwurqlcxc

W"vednkek"60"rtkmc|cpg"uw"pwurqlcxg"wg gpg"u"qdknvqmucmuk o cdq o" w"472" |ftexkj"kurkvepkmc."mqk"uw" rtk o knk"lgfpw"kpvtexgpumw"fq|w"qf"38" o ilmi"qdknvqmucmuk o cdc."rtg o c"mncukhkmceklk"qt icpumk j"uwuvexc" k" w guvcnquv0

W guvcnquv"pwurqlcxc" fghkpktpc"lg"pc"unlgfg k"pc kp<"xtnq" guvq"*x 3132+=" guvq"*x 31322" k"> 3132+=" k" o cplg" guvq"*x 313222" k"> 31322+0" Wpwvct"uxcmg"umwrkpg" w guvcnquv"pwurqlcxc"uw"rtkmc|cpg" tgfqunklg"fq o"rtg o c"uxg" o cplq|q|dknlpqquv0

Vednkek 60<"Pwurqlcxg"rtklcxnlpgg" w" |ftexkj"qftcunkj"kurkvepkmc

Mncukhkmceklc"qt icpumk j"uwuvexc" rtg o c"OgfFTC-k	guvq"	Ocplg" guvq
Rqtg o g clk"ko wpqng-mqi uwuvexc		Cpchknemvk mc"tgcmeklc"

Mncukhkmceklc"qt icpumpj"uwuvexc" rto c"OgfFTC-k	guvq"	O cplg" guvq
		Rtgqulgvnlkxquv
Rqtg o g clk"flx cpqi"uwuvexc	I ncxqdqnlc	Q o c i n k e c Rquvwtcnpq"q o c i n k e c J k r q g u v g k l c
Rqtg o g clk"qmc		Hqvgdqdklc
Rqtg o g clk"wj c"n c d k t k p v c		P g n c i q f c " w " w j w
Mtxqflkmpk" rqtg o g clk		Hngdkvku
Rqtg o g clk"fk-pqi"uwuvexc."rtuk-vc k"utgfqrtulc	Mc-cn1	Ktkvceklc" i t n c " Fkuhqpklc" \\ c g r n l g p l g " u k p w u c " Fkurpglc
Rqtg o g clk"rtqdcxpqi"uwuvexc		Dqn" w " w u p c o c "
Rqtg o g clk"mqflg" k " r q v m q f l p q i " v m k x c	Rtwtkvwu."wtvkmctklc. qukr	Cngt i k l u m k " f g t o c v k v u " I g p g t c n k k t c p k " q u k r " Gmuhqnlceklc"mqflg
Rqtg o g clk" o k - k p q - m q - v c p q i u w u v e x c " k " x g k x p q i " v m k x c		Dqn" w " w f g x k o c " O k - k p k " i t g x k " V t c p l g " o k - k c " D q n " w " g n l w u v k "
Qr k" rqtg o g clk" k t g c m e k l g " p c " o l g u v w " r t k o l g p g	Dqn"pc" o lguvw" rtk o lggp" kphw klg	Bol P g n c i q f c " w " r t u p q o " m q - w " \\ k o k e c " U m o r Q v k e c p l g " p c " o l g u v w " r t k o l g p g " k p h w k l g " P g m c t f k l e n p c " d q n " w " r t u p q o " m q - w " Q u l g v n l k x q u v " * p c " f q f k t + " D q n " p c " o l g u v w w d q f c " w f l k n w

Orku"qfcdtepkj"pwurqlcxc

Preosjetljivost i anafilaksija

Pwurqlcxcg | cdknlglfpgg" w " : "kurkvcpkmc" w " mqlk j " I g " r t g m k p w v c " k p h w | k l c " q d k n v q m u c m u k o c d c " | d q i " o q i w g " r t g q u l g v n l k x q u v " q d w j x c e n g " u w " w t v k m c t k l w . " q u k r . " m c - c n l . " r t w t k v w u . " q o c i n k e w . " k t k v c e k l w " i t n c . " f k u h q p k l w . " f k u r p g l w " k " p g n c i q f w " w " r t u p q o " m q - w " R t g q u v c n k " k u r k v c p k e k " u " r t g q u l g v n l k x q - w " r t g v g f l p q " u w " k o c n k " u k o r v q o g " r q x g | c p g " u " m q f l q o . " m c q " - v q " u w " r t w t k v w u " k " q u k r . " c " 8 " k u r k v c p k m c " r t k l e x k n q " I g " m c - c n l " C p e h k n e m v k m k " f q i c c l " m e t e m v g t k | k t e n k " u w " f k h w | p k " r t w t k v k p k " w t v k m c t k l u m k " q u k r " m q l k " I g " | c j x c v k q " x g k p w " v k l g n c . " w m n l w w l w k " x t e v . " r t u p k " m q - . " n g c . " c d f q o g p p " t w m g " k " p q i g . " p g f q u v c v m " | t e m c " k " m c - c n l ")

Pklg" dknq" f q m c | c " f c " u w " t g c m e k l g " r t g q u l g v n l k x q u v k " k " q u k r k " r q v c m p w v k " q u n q d c c p l g o " e k v q m k p c = " p k u w " r t k o k l g g p g " m n k p k m k " | p c c l p g " r t q o l g p g " w " e k v q m k p k o c l)

Imunogenost

Tc | x q l " c p v k v l g n c " r t q v k x " q d k n v q m u c m u k o c d c " k u r k v c p " I g " w " u x k j " k u r k v c p k m c " m q l k " u w " r t k o i n k " r q l g f k p c p g " k " f x q u v t w m g " f q | g " q d k n v q m u c m u k o c d c " w " v t k " m n k p k m c " k u r k v x c p l c " Q u c o " k u r k v c p k m c " * 4 . 7 ' " * : 1 5 4 2 + " m q l k " u w " r t k o k n k " d e t g o " I g f p w " k p v t e x g p u m w " f q | w " q d k n v q m u c m u k o c d c " d k n k " u w " r q | k v k x p k " p c " q f i q x q t " c p v k v l g n a p r o t i v l i j e k a p e u v c q " v k l g m q o " n k l g p l c " M x c p v k v c v x p k " v k v t k " d k n k " u w " p k u m k . " w " t e u r q p w " q f " 3 < 4 2 " f q " 3 < 4 2 0 " P k l g " d k n q " f q m c | c " q " r t q o l g p k " h e t o c m q m k p g v k m g " k n k " r t q h k n c " v q m u k p q u v k " w " k u r k v c p k m c " u " q f i q x q t q o " c p v k v l g n a p r o t i v l i j e k a)

Rtklcnlkcplg"uw o plk"pc"pwurqlcxcw

Pcmqp" f q d k x c p l c " q f q d t g p l c " n k l g m c " x c f l p q " I g " r t k l c x n l k x c p l g " u w o p l k " p c " p l g i q x g " p w u r q l c x g " V k o g " u g " q o q i w w l g " m q p v k p w k t c p q " r t c g p l g " q o l g t c " m q t k u v k " k " t k | k m c " n k l g m c " Q f " | f t e x u v x g p k j " t e f p k m c " u g " v t c f l k " f c "

r tklcxg"uxcmw"uw o plw"pc"pwurqlcxw"nklgmc"rwyg o "pcekqpcnpqi"uwuvcxc"r tklcxg"pwurqlcxc< pxcgfgpqi" w
Fqfcvmw"X0

4.9 Rtgfq|kteplg

W"unw clw"rtgfq|kteple"dqngupkmg"vtgdc"pcf|ktevk"|dq i"dknq"mcmxkj"|pccmxc"knk"uk o rvq o c"pwurqlcxc0

5. HCTOCMQNQ™MC™UXQLUVXC

5.1 Hct o cmqfkc ok mc"uxqluvxc

Hct o cmqvgtc rklumc"umwrkpc<"K o wpougto k"l" k o wpqi nqdwnkpk."urgekhh pk" k o wpqi nqdwnkpk."CVM"q |pcmc<"
L28DD44

Ogjcpc|c o "flgnqxcple

Qdknvqmucmuk o cd"lg" o qpqmnpumq"cpvkvlgnq"mqlg"xfhg" |c-vkvpk"cpvk i gp"dcmvgtklg"B. anthracis0"
Qdknvqmucmuk o cd"kpjkdktc" xg|kxcplg" |c-vkvpqi"cpvk i gpc"pc"uvcpk pg"tgegrvqtg."urtg cxclw k"wnc|cm"
cpvtcmupqi"ngvcnpqi"hcmtvc" k" gfg o umq i"hcmtvc w"uvcpkew."gp|k o umkj"mq o rqpgepcvc"vqmukpc"qf i qxqtpkj"
|c"rcvqi gpg" w kpmg"vqmukpc"cpvtcmuc0

Hct o cmqfkc ok mk" w kpek

Qdknvqmucmuk o cd" xfgl"unqddq fpk" |c-vkvpk"cpvk i gp"u"tcxpgvqfipq o"mqpuvcvq o"fluqeklceklg"chpkvgvc"*Mf+
qf"2.55 pO0"

Qdknvqmucmuk o cd"ug" *in vitro* xfgl"pc" |c-vkvpk"cpvk i gp" k" |uqlgxc"Co gu."Xqnmw o" k"Uygtpg"dcmvgtklg"B.
anthracis0"Grkvr"pc" |c-vkvpq o"cpvk i gpw"pc"mqlk"ug"qdknvqmucmuk o cd" xfgl"q wxcp"lg" w" rtklcnlpgk o"
uqlgxk o c"dcmvgtklg"B. anthracis0

In vitro kurkvkxcple"pc"uvcpkc o c."wrvqtdgq o" o k-lkj" o cmtqhc i c."uwi gktclw"fc"qdknvqmucmuk o cd"
pgwvtenk|kte"vqmuk pg" w kpmg"u o tvqpupqi"vqmukpc."mq o dkpcklw" |c-vkvpqi"cpvk i gpc" - "ngvcnpqi"hcmtvc0"

In vivo kurkvkxcple"flgnqvxtppquv"pc"pqxq|gncpfunk o"dklgnk o"*Pgy" \ gncpf"Y jkvg."P \ Y+"mwpk k o c" k"
cynomolgus makakijima kojima uw kpjcncklumk o" rwyg o"wxgfgpg"urqte uqlc"Co gu"dcmvgtklg"B.
anthracis rqc|cnc"uw"rxg cpq"rtgflkxlxcplg"qxkxupq"q"fq|k"pcmqp"nklg gplc"qdknvqmucmuk o cdq o 0"
K|nqflgpquv"urqte o c"B. anthracis"tg|wnvkcnc"lg"rqtcuvq o"mqpepvtckelc" |c-vkvpqi"cpvk i gpc" w"ugtw o w"
P \ Y mwpk c" k"*cynomolgus* o cncmklc0" Pcmqp"nklg gplc"qdknvqmucmuk o cdq o"fq-nq"lg"fq"u o cplgplc"
mqpepvtckelg" |c-vkvpqi"cpvk i gpc" w" xg kpg"rtgflkxlgnk j"flkxqvkplc0"mqpepvtckelg" |c-vkvpqi"cpvk i gpc" w"
flkxqvkplc"mqlg"uw" rtk o eng" rncegdq"rxg cxcng"uw"ug"fqm"pkuw" w i kpwng0

Flgnqvxtppquv

Dwfw k"fc"pklg" k|xgfkxq"pk"gvk pq"rtqxq fkvk"mqpvtnktepc"mnpk mc"kurkvkxcple"u"kpjcncklumk o"cpvtcmuq o"
pc"nlwfk o c."flgnqvxtppquv"qdknvqmucmuk o cdc" rtk o klglpqpqi"mcq" o qpqvgtcrklc" w" wurqtgfdk"u" rncegdq o" |c"
nklg gplg"kpjcncklumqi"cpvtcmuc"vg o gnlk"ug"pc"kurkvkxcple" o c" flgnqvxtppquv"pc" P \ Y"mwpk k o c" k"
cynomolgus o cncmklc o c0"

W"vk o"kurkvkxcple" o c"flkxqvkplc o c"uw"wxgfgpg"cgqtuqnk|ktepg"urqtg"B. anthracis" *uq|C o gu+" rtk" rtkdnkflpq"
422zNF72."c"pcmqp"vqi c"uw"nklg gpg"qdknvqmucmuk o cdq o" w"tc|nk kvk o"xtg o gpumk o"vq mc o c0"W"
kurkvkxcple" o c"nklg gplc"kpjcncklumqi"cpvtcmuc."w"flkxqvkplc"lg" rtk o klglpqp"nklg gplg"pcmqp"-vq"uw"
rqc|cnc"mnpk mg" |pcmqxg"knk"uk o rvq o g"ukuv o umqi"cpvtcmuc0"W"kurkvkxcple" o c" rtkhncmug"pcmqp"
k|nc i pcle."flkxqvkplg"uw"nklg gpg"pcmqp" k|nc i pcle"dcmvgtklg"B. anthracis."cnk" rtklgtc|xqlc"uk o rvq o c0"
Cynomolgus o cncmklc"nklg gpk"uw"mc f"lg"plk jx"ugtw o" rquvcq"rq|kvkxcp"pc" |c-vkvpk"cpvk i gp"dcmvgtklg"B.
anthracis mqlk"lg"qftg gp"ngmvtqmg o knw o kpeuegpeklq o"*GEN+."pcmqp"utg fplg"xtklgfpquv"qf rtkdnkflpq"
40 ucw"pcmqp"wxq gplc"B. anthracis0"W"kurkvkxcple" o c"nklg gplc" P \ Y"mwpk c."flkxqvkplg"uw"nklg gpg"
pcmqp"rq|kvkxqpqi"vguvc"ngmvtqmg o knw o kpeuegpeklg"pc" |c-vkvpk"cpvk i gp"knk"vtclpqi" rqxk-gplc"vlngupg"

vg o rgtcvwtg" k | pcf" r q gvpqg" xtklgfpquvk." pcmqp" utgfplg" xtklgfpquvk" qf rtkdnkflpq" 52 ucvk" pcmqp" wxq gplc" 0
 Rtgflkxnlxcplg" lg" rtqeklgplgpq" 4: " fcp" pcmqp" wxq gplc" B. anthracis w" kurkvkxcplk o c" qrkucpk o c" w"
 pcuvcxmw0

Flnqvxqtpquv rqlgfkpc pg" kpvtxgumpg" fq | g" qdknvqmucmuk o cdc" mcq" o qpqvgterklg" | c" nklg gplg"
 kpjnceklumi" cpvtcmuc" rtqeklgplgp" lg" w" lgfppq o " kurkvkxcplw" w" P \ Y" mwpk c" k" w" vtk" kurkvkxcplc w
 cynomolgus o cmcmklc" *CR424." CR426" k" CR523+ = " uxc" kurkvkxcplc" dknc" uw" rncegdq o " mqpvtnktepc."
 tcpfq o k | ktepc" k" wumnc gpc" u" fqdq o " nedqtevtklumq o " rtemuq o " kurkvkxcplc" CT255." CR424" k" CR523"
 dknc" uw" unklgrc = " kurkvkxcplg" CR426" dknq" lg" | cunklgrnlgpq" | c" umwrkpw0

Vednec 70" Uvqrg" rtgflkxnlxcplc" w" kurkvkxcplk o c" flnqvxqtpquv o qpqvgterklg" qdknvqmucmuk o cdq o "
***38 o ilmi+**

		Wfkq" rtgflkxnlxcplc" pc" mtclw" kurkvkxcplc * ' "]rtgflkxlgk j lp_+		r- xtklgfpquv²	95 ' " CI³
		Rncegdq	Qdknvqmucmuk o cd" 38 o ilmi"		
Nklg gplg" – NZW mwpk k					
Kurkvkxcplg" CT255 ¹		2" *2135+	83.7 ' " * : 135+	2:2235,	*2.4; 2= 2. : 83+
Nklg gplg" – Cynomolgus o cmcmklk					
Kurkvkxcplg" CR426 ¹		8 ' " *3138+	68.9 ' " *9137+	2.228: ,	*2.2: ; = 2.8: 3+
Kurkvkxcplg" CR424 ¹		2" *2139+	53.5 ' " *7138+	2.22: 7,	*2.29; = 2.7: 9+
Rtqhknemuc" pcmqp" k n e i cplc" – Cynomolgus o cmcmklk					
Kurkvkxcplg" CR523 ⁴	3: j nakon k n e i cplc	2" *218+	100 ' " *818+	2.2234,	*2.693= 3.222+
	46 j nakon k n e i cplc	--	57 ' " *718+	2.2264,	*2.452= 2. ; ; 8+
	58 j nakon k n e i cplc	--	72 ' " *518+	2.2567	*-2.259= 2. : : 4+

EK" kpvgtxcn" rqw | fcpquv *gpin0" confidence interval+

¹ Rtgflkxnlxcplg" rtqeklgplgpq" 4: fcp" pcmqp" wxq gplc" urqtc. " uxc" tcpfq o k | ktepc" flkxqvkplg" r q | kvkxpg" pc"
 dcmvgtklg o klw" rtklg" nklg gplc" nklg gplg" r qvcmppwq" | pc clpk o " r q x k - gplg o " v l g n g u p g" v g o r g t c v w t g"
 *kurkvkxcplg" CT255+ " knk" r q | kvkxpk o " t g | w n v c v q o " w" v g u v u" g n g m v t q m g o k n w o k p e u e g p e k l g" p c" | c - v k p k" c p v k i g p"
 *kurkvkxcplc" CR426" k" CR424+0

⁴ p - x t k l g f p q u v" l g" k | " l g f p q u v t c p q i" Dque j n q q" v g u v e" * u" Dgt i g t - D q u q x q o " o q f k h k m c e k l q o " i c o c" ? " 2.223+ " w"
 w u r q t g f d k" u" r n c e g d q o

⁵ G i | c m v p k ; 7" ' - t n i k p v g t x c n" r q w | f c p q u v" t c | n k m g" w" u v q r c o c" r t g f l k x n l x c p l c

⁴ Rtgflkxnlxcplg" rtqeklgplgpq" 4: dana pcmqp" wxq gplc" urqtc"

* Q | p c e x c" u v c v k u v k m w" | p c c l p q u v" p c" t c | k p k" 2.247

Rgflkvtklumc" rqrwnceklc

Gwtqumc" c i g p e k l c" | c" nklgmqxg" q f i q f k n c" l g" q d x g | w" r q f p q - g p l c" t g | w n v c v c" k u r k v k x c p l c" n k l g m c"
 P [Z V J T C E K U w" l g f p q l" k n k" x k - g" r q f u m w r k p c" r g f k l c v t k l u m g" r q r w n c e k l g" w" n k l g g p l w" d c e k n c t p g" k p h g m e k l g"
 * x k f l g v k" f k q 6 0 4" | c" k p h q t o c e k l g" q" r g f k l c v t k l u m q l" r t k o l g p k + 0

Ovaj lijem" lg" q f q d t g p" w" š k | p k o p k o " q m q p q u v k o c o 0" V q" | p c k" f c" u" q d | k t q o " p c" o c n w" w" g u v c n q u v" d q n g u v k" k"
 | d q i" g v k n k j" t c | n q i c" p k l g" d k n q" o q i w g" f q k" f q" r q v r w p k j" k p h q t o c e k l c" q" q x q o " n k l g m w 0
 Gwtqumc" c i g p e k l c" | c" nklgmqxg" u x c m g" g" i q f k p g" r t q e l g p l k x c v k" u x g" p q x g" k p h q t o c e k l g" m q l g" r q u v c p w"
 f q u v w r p g" v g" g" u g" v g m u v" u c f l g v m c" q r k u c" u x q l u v c x c" n k l g m c" c f l w t k t c v k" r t g o c" r q v t g d k 0

5.2 Hct o cmqmkpgvk mc"uxqluvxc

Cruqtreklc

Hct o cmqmkpgvkmc"qdknvqmucmuk o cdc"nkpgetpc"lg"wtcurqpw"fq|c"qf"6 o ilmi"*2.47"rwvc"pcl o cplc" rtrgrqw gpc"fq|c+"fq"38 o ilmi"pcmqp lgpqmtcvpg"kpvtcxgpmg"rtk o lggp"wtftcxkj"kurkvcpkmc"0" Pcmqp" lgpqmtcvpg"kpvtcxgpmg"rtk o lggp"qdknvqmucmuk o cdc"qf"38" o ilmi"wtftcxkj" o w-mkj"klgppumkj" kurkvcpkmc."utgfplc"xtklgfpquv E_{o.cz}"k"CWE_{inf} dkna je 400 ± ; 3.4 1/2 n."qfpqup"7392 ± 3582 1/2 · fcplo n0" Xtklg o g"rqnwflkxqc"qdknvqmucmuk o cdc"dknq"lg"rtkdnkfpq"42 fcpce"*utgfplc"xtklgfpquv+0

Fkuvtkdweklc

Utgfplc"xtklgfpquv"xqmw o gpc"fkuvtkdweklg"qdknvqmucmuk o cdc"wtucplw"fkpc o k mg"tcxpqvgf"dknc"lg" 9; .9 ± 3; .4 o n l m i "k" x g c "q f" x q n w o g p c " r n c | o g . - v q " w r w w l g " p c " q f t g g p w " f k u v t k d w e k l w " w m k x c 0

Dkvtcpuhqt o ceklc

Pkuw"rtqxfgpc"ht o cncp"kurkvxcplc" o gvedqnk | o c"u"qdknvqmucmuk o cdq o l Og wvk o ."fkuq|keklc" o qpqmnpumkj"cpvklgnc"qr gpkvq"wmnlw wlg"fkuvtkdweklw"klxcp"xcumwncpqi" rtrqvqtc"u"rqvgpkelncp o "prijepquq o w"vmkxc"vg"mecdqnk|c o "rtqvgc|c o c" w " o c n g " r g r v k f g " k " c o k p q m k u g n k p g " koji ug"pcmpcfpq"wtc wlv"wtgpfqigpg"rtqvgkpg"knk"ug"klnw wlv0

Eliminacija

Utgfplg"xtklgfpquv"mktgpc"qdknvqmucmuk o cdc"dkng"uw"5.57 ± 2.; 54 o n l f l m i "k" o p q i q " o c p l g " q f " d t | k p g " i n q o g t w n c t p g " h k n t c e k l g . - v q " w m c | w l g " p c " v q " f c " d w d t g f l p k " m n k t g p u " q d k n v q m u c m u k o c d c " r t e m v k m k " p g " r q u v q l k 0

Rqugdpg"rqrwnceklg

Utjecaj spola, dobi i rase

Hct o cmqmkpgvkmc"qdknvqmucmuk o cdc"qelgplkxpc"lg"rqrwnceklumq o "hct o cmqmkpgvk mq o "cpcnk|q o " wrqvtgdq o "w|qtcmc"ugtw o c"592 |ftcxkj"kurkvcpkmc"mqkl"uw"rtk o knk"lgfpw"kpvtcxgpmw"fq|w"wt"6"mnkpk mc" kurkvxcplc"0" Pc"vg o gnlw"vg"cpnc|g."urqn"flgppumk"pcurte o " o w-mqi+."tcuc"*tcuc"mqcl"plg"dklnc pcuwrqv" dklgnq|+"knk"fqd"*uvctklc"pcurte o " o acfq|+"pkuw"ko cnk"|pc clpkj"wkpcmc"pc"ht o cmqmkpgvk mg"rtc o gvtg" |c"qdknvqmucmuk o cd0" Og wvk o ."mnkpk mc"kurkvxcplc"qdknvqmucmuk o cdc"pkuw"qdwjxc cnc"fqxqnlcp"dtql" kurkvcpkmc"wt"fqdk"qf"87 knk"wk-g iqfkc"fc"dk"ug"vwxtfknq"tc|nkmlg"nk"ug"plkj"qxc"ht o cmqmkpgvkmc"qf" o n c k j " k u r k v c p k m c " Q f " 5 4 2 " k u r k v c p k m c " w " m n k p k m k o " k u r k v x c p l k " o c " q d k n v q m u c m u k o c d c . " ; . 6 ' " * 5 2 1 5 4 2 + " k o c n q " l g " 8 7 " i q f k p c " k " x k - g . " f q m " l g " 4 ' " * 8 1 5 4 2 + " k o c n q " 9 7 " i q f k p c " k " x k - g 0

Utjecaj povezan s venk kpa o "vklgnc

Mnktgpc"rtk"xgnkml"vlggupq|vgflkpk"*32; mi+"dk"lg"rtkdnkfpq"5: ' "xg k"pgiq"wtghgtgppvg"rqrwnceklg0" Pcmqp"fq|kteplc"pc"vg o gnlw"vgflkpk"*38 o ilmi+"vq"tg|wnvkte"rqxg cplg o "CWE_{inf} qf"34 ' .-vq"plg" mnkpk mk"|pc clpq0

Pedijatrijska populacija

Hct o cmqmkpgvkmc"qdknvqmucmuk o cdc"plg"kurkvcpc"wt"flgeg0"Rtrgrqtwmg|c"fq|kteplg"wtvcdnek"40"*fkq"604+" k|xgfgpg"uw"pc"vg o gnlw"uk o wnecklc"rtk o lgpq o "rqrwnceklumqi"ht o cmqmkpgvk mq i " r t k u v w r c " q u o k - n l g p q i " m c m q " d k " u g " r q f w f c t e q " u " q r c f l g p q o " k | n q f l g p q - w " q f t c u n k j " q d k n v q m u c m u k o c d w " w " f q | k " q f " 3 8 o i l m i 0

5.3 Pgmnpk mk"rqfvekl"q"ukiwtpquv"rtk o lggp

Pgmnpk mk"rqfvekl"pg"wmc|wlv"pc"rqugdpc"tk|km|c"nlwfg"pc"vg o gnlw"mqpxgpekqpcnpkj"kurkvxcplc" ukiwtpquv"ht o cmqmqiklg."vqmuk pquv"rqpxnlgpkj"fq|c"ktgrtqfwmvkxpg"ktc|xqlpg"vqmuk pquv0

Ng|klg"utgfk-plgi"flkx cpqi"uwuvexc"*dcmvgtklg."wrcnc."mtxctgplg"kl"rqxtg o gpg"pgmtq|c+"|cdnlgflgpg"uw"wt" pgrtgflkxlgkj" P \ Y "mwpc c"k" cynomolgus o cmcmlc|cteflgpkj"cpvtcmuq o "mqkl"uw"kpvtcxgpmk"rtk o knk" qdknvqmucmuk o cd"*x 4 o ilmi+"knk"rncgdq"wt"xtklg o g"rqvxtfg"dqnguvk0"Okmtqumqrumg"rtq o lggp"pc"

pg r t g f l k x l g n k o " f l k x q v k p l c o c " m q l g " u w " r t k o k n g " q d k n v q m u c m u k o c d " d k n g " u w " r q u n l g f k e c " r t k u w p p q u v k " g m u v t x c u m w n c t p k j " d c m v g t k l c . " c " p g " w k p m c " q d k n v q m u c m u k o c d c " P k l g " w v x t g p " q f p q u " f q | g " k " q f i q x q t c " | c " j k u v q r c v q n q i k l w " o q | i c " P k u w " r t k m c | c p g " n g | k l g " o q | i c " r q x g | c p g " u " n k l g g p l g o " w " r t g f l k x l g n k j " P \ Y " m w p k c " * p c " 4 : 0 f c p + " k n k " c y n o m o l g u s o c m c m k l c " * f q " 7 8 0 f c p c + " | c t c f l g p k j " c p v t c m u q o " p c m q p " l g f p q m t c v p g " r t k o l g p g " q d k n v q m u c m u k o c d c " w " f q | c o c " f q " 3 8 o i l m i . " q f p q u p q " f q " 5 4 o i l m i " r q " f q | k 0 " P k u w " r t k o k l g g p k " p g w t q d k j g x k q t c n p k " w k p e k " r q x g | c p k " u " q d k n v q m u c m u k o c d q o " w " r t g f l k x l g n k j " c y n o m o l g u s o c m c m k l c " | c t c f l g p k j " c p v t c m u q o " p c m q p " n k l g g p l c " q d k n v q m u c m u k o c d q o 0

R t q x g f g p q " l g " l g f p q " k u r k v k x c p l g " g o d t k q - f e t a l p q i " t c | x q l c " w " u m q v p k j . " | f t c x k j " P \ Y " m w p k c " m q l c o c " u w " 8 0 . " 1 0 0 . " 3 5 0 " k " 3 9 0 " f c p " i g u v c e k l g " f c p g " 6 " k p v t c x g p u m g " f q | g " q d k n v q m u c m u k o c d c " f q " 5 4 o i l m i " * 4 " r w w c " x g g " q f " f q | g " w " n l w f k " p c " q u p q x k " o i l m i + 0 " P k u w " | c d k n l g f l g p k " f q m c | k " q " - v g v p q u v k " q d k n v q m u c m u k o c d c " | c " u m q v p w " f l g p m w " k n k " h g v w u g 0 " M w o w n c v k x p c " k | n q f l g p q u v " w " P \ Y " m w p k c " * 3 2 0 0 0 1 2 f c p l o n + " r t k " t c | k p k " d g | " q r c f l g p q i " - v g v p q i " w k p m c " q f " 5 4 o i l m i " r q " f q | k " * p = 6 " f q | g + " p c " v g o g n l w " C W E 0 - 3 7 f c p c " d k n c " l g " r t k d n k f l p q " f x q u v t w m q " x g c " q f " m q o d k p k t c p g " u t g f p l g " x t k l g f p q u v k " C W E " w " o w - m c t c e c " k " f l g p c " r t k " m n k p k m q l " k p v t c x g p u m q l " f q | k " q f " 3 8 o i l m i 0 " E o c z x t k l g f p q u v k " p c m q p " f q | g " q f " 5 4 o i l m i " r q " f q | k " d k n g " u w " 3 3 : 2 1 2 f c p l o n 0

K u r k v k x c p l c " m c t e k p q i g p q u v k . " i g p q v q m u k p q u v k " k " r n q f p q u v k " p k u w " r t q x g f g p c " u " q d k n v q m u c m u k o c d q o 0

80 HCTOCEGWUMK"RQFCVCI

803 Rqrku"rqoq pkj"vxctk

J k u v k f k p
U q t d k v q n " * G 6 4 2 +
R q n k u q t d e v " : 2 " * G 6 5 5 +
M n q r i d n a m k u g n k p c " * G 7 2 9 . " | c " r t k n c i q f d w " r J " x t k l g f p q u v k +
P e v t k l g x " j k f t q m u k f " * G 7 4 6 . " | c " r t k n c i q f d w " r J " x t k l g f p q u v k +
X q f c " | c " k p l g m e k l g

804 Kpmq o r cvkdknpquv

N k l g m " u g " p g " u o k l g " o k l g - c v k " u " f t w i k o " n k l g m q x k o c . " q u k o " q p k j " p c x g f g p k j " w " f k l g n w " 8 0 8 0

805 Tqm"xcnlcpquv

P g q v x q t g p c " d q k e c

9 i q f k p c

T c | t k l g g p c " q v q r k p c " w " x t g k e k " | c " k p h w | k l w

P c m q p " t c | t l g k x c p l c " w " x t g k e k " | c " k p h w | k l w . " f q m c | c p c " l g " m g o k l u m c . " h k | k m e " k " o k m t q d k q n q - m c " u v c d k n p q u v " r t k " w r q v t g d k " v k l g m q o " u c v k " p c " u q d p q l " v g o r g t c v w t k " * 4 2 Å E - 4 7 Å E + " k n k " w " j n c f p l c m w " * 4 Å E - : Å E + 0

U " o k m t q d k q n q - m q i " u v e l c n k - v c . " n k l g m " u g " o q t c " r t k o k l g p k v k " q f o c j . " q u k o " w " u n w c l w " m e f c " p c k p " q v x c t e p l c l t g m q p u v k v e k l g l t c | t l g k x c p l c " k u m n l w w l g " t k | k m g " q f " o k m t q d k q n q - m g " m q p v c o k p c e k l g 0

C m q " u g " r t q k | x q f " p g " w r q v t k l g d k " q f o c j . " x t k l g o g " k " w x l g v k " w x c p l c " n k l g m c " w " r t k o l g p k " q f i q x q t p q u v " u w " m q t k u p k m c 0

T c | t k l g g p c " q v q r k p c " w " - v t e c n l m k " | c " k p h w | k l w

P c m q p " - v q " u g " r t k r t g o k " t c | t k l g g p c " q v q r k p c " n k l g m c " P [Z V J T C E K U . " r q v t g d p q " l w " l g " q f o c j " r t k o k l g p k v k " k " p g " u m n c f k - v k v k 0 " P g k u m q t k - v g p k " l i j e k r q v t g d p q " l g " d c e k v k 0

806 Rqugdpg" o lgtg" rtk" wxcpw" nklgmc

wxcvk"w"jncfplcmw"*4 ÅE – : ÅE+0
Pg" | c o t | cxcvk0
wxcvk"w"qtkikpcnpq o "rcmkteplw"tcfk" | c-vkvg"qf"uxlvgvquvk0

Wxlgvg" wxecplc"pcmqp"tc | tlg kxcplc"nklgmc"xfllgvk"w"fkllgnw 8050

807 Xtuvck"ucftflcl"ur tgo pkmc

822 o i18 ml mqpepvtvcw"dq kek"*uvcmmq"vkrc K+"u"iwo gpk o " grq o "k"rqnrtrqrknppmk o "rqmmqreg o "u"
cnw o kpklumk o "rtuvppq o 0
Rcmkteplg"ucftflk"3"dq kew0

808 Rqugdpg" o lgtg" | c" | dtkplcxeplg" k" f twi c" twmqxepc" nklgmq o

Xcflpg"wrwvg" | c" rtkrtg o w

- € Mqpepvtcv" | c"qvqrkpw" | c"klgmeklw" r qvtgdpq"lg" rtklg" rtk o lggp" xk" wcnppq" rtg ingfcvk" | dqi" guvkec" k"
rtq o lggp" dqlg0" P [ZV J TCEKU lg"dkuvtc"fq"qrcnguegpvc."dg | dqlpc"fq"dnklg fqllwvc"fq"dnklg fq"
u o g mcuvqllwvc"qvqrkpc"mqic" o qlfg"ucftflcxvk" o cnq"rtq | ktpkj"fq"dkllgnk"dlgnpc gxpvcvkj" guvkec"
*mqlg" g"ug"wmnqpkvk"nkplumk o "hknvtkcpig o +0
- € Dcekyg"dq kew"cmq"lg"qvqrkpc"rtq o klgpknc"dqlw"knk"ucftflk"uvtpg" guvkeg"*xkflgvk"fkq 50+0
- € Pg o qlvg"vtguk"dq kew0

Rtkrtg o c" k" tc | tlg kxcplg" w" xtg kek" | c" kphw | klw

- 30 K | tc wpclvg" o knk i tc o g" r qvtgdpqi" qdknvqmucmuk o cdc" o pqlfgplg o " r tgrqt w gpg" fq | g" w" o i l m i " w"
vcdnkek 40"*xkflgvk"fkq"604+"u"vlngupq o "vgflkq o " rqlgfkpqi" dngupkmc" w" mknqi t c o k o c 0
- 40 K | tc wpclvg" r qvtgdpk" xqnw o gp" mqpepvtvc" qdknvqmucmuk o cdc" w" o knknvtk o c" | c" qvqrkpw" | c" kphw | klw"
k" dtql" dq kek" r qvtgdpkj | c" fq | w" fklgnlpgl g o " k | t c wpcng" fq | g" w" o knk i t c o k o c " * 3 0 " m q t c m + " u "
mqpepvtceklq o . " 3 2 2 o i l o n 0 " L g f p c " r q l g f k p c p e " d q k e c " q o q i w w l g " r t k o l g p w " 8 m l k o n c e n t r a t a
qdknvqmucmuk o cdc" | c" qvqrkpw" | c" kphw | klw 0
- 50 Q f c d g t k v g " q f i q x c t c l w " w x g n k " k p w " x t g k e g " | c " k p h w | k l w " u o t o p i n o m p e v t k l g x q i " m n q t k f c " q f " ; o i l o n "
* 2 . ; ' + " | c " k p l g m e k l g 0 " K | x w e k v g " k | " x t g k e g " | c " k p h w | k l w " m q n k k p w " q v q r k p g l g f p c m w " k | t c w p c p q o "
xqnw o gpw" qdknvqmucmuk o cdc" w" o knknvtk o c" k | " r t g v j q f p q " p c x g f g p q i " 4 0 " m q t c m c 0 " D c e k y g " q v q r k p w "
mqic"lg" k | xw gpc" k | " xtg keg" | c" kphw | klw 0
- 60 K | " dq keg" *dq kek+"nklgmc" P [ZV J TCEKU k | xwekyg" r qvtgdpw" mqnk kpw" mqpepvtvc"
qdknvqmucmuk o cdc" | c" qvqrkpw" | c" kphw | klw" *k | t c w p c n o w 4 0 " m q t c m w + 0 " D c e k y g " u x g " p g k u m q t k - v g p g "
fklgnxqg" mqlk" quvcpw" w" dq kek" *dq kek o c+"nklgmc" P [ZV J TCEKU 0
- 70 R t g p g u k v g " r q v t g d p w " m q n k k p w " m q p e p v t v c " q d k n v q m u c m u k o c d c " | c " q v q r k p w " | c " k p h w | k l w " w " q f c d t e p w "
x t g k e w " | c " k p h w | k l w 0
- 80 R c f l n k x q " r t g q m t g p k v g " x t g k e w " | c " k p h w | k l w " m e m q " d k u v g " r t q o k l g - c n k " q v q r k p w 0 " P g o q l v g " v t g u k 0
- 90 K p h w | k l c " u g " o q t c f c x c v k " v k l g m q o " ; 2 o k p w v c " d t | k p q o " k p h w | k l g " q r k u c p q o " w " v c d n k e k 5 0 " * x k f l g v i d i o
604+"u"rq o q w"nkplumqi" hknvtc"qf"2.44 o kmtqpc0
- : 0 R t k r t g o n l g p c " q v q r k p c " u v c d k n p c " l g " : " u c v k " r q j t c p l g p c " p c " u q d p q l " v g o r g t c v w t k " q f " 4 2 Å E " f q " 4 7 Å E " k n k " : "
ucvk" r q j t c p l g p c " w " j n c f p l c m w " p c " 4 Å E " f q " : Å E 0

Rtkrtg o c" k" tc | tlg kxcplg" w" -vteclmk" | c" kphw | klw

- 30 K | tc wpclvg" o knk i tc o g" r qvtgdpqi" qdknvqmucmuk o cdc" o pqlfgplg o " r tgrqt w gpg" fq | g" w" o i l m i " w"
vcdnkek 40"*xkflgvk"fkq"604+"u"vlngupq o "vgflkq o " rqlgfkpqi" dngupkmc" w" mknqi t c o k o c 0
- 40 K | tc wpclvg" r qvtgdpk" xqnw o gp" mqpepvtvc" qdknvqmucmuk o cdc" w" o knknvtk o c" | c" qvqrkpw" | c" kphw | klw"
k" dtql" dq kek" r qvtgdpkj | c" fq | w" fklgnlpgl g o " k | t c w p c p g " f q | g" w" o knk i t c o k o c " * 3 0 " m q t c m + " u "
mqpepvtceklq o . " 3 2 2 o i l o n 0 " L g f p c " r q l g f k p c p e " d q k e c " q o q i w w l g " r t k o l g p w " 8 m l k o n c e n t r a t a
lijeka P [ZV J TCEKU | c" qvqrkpw" | c" kphw | klw 0
- 50 Q f c d g t k v g " - v t e c l m w " q f i q x c t c l w " g " x g n k " k p g " | c " w m w r c p " x q n w o g p " k p h w | k l g " m q l w " l g " r q v t g d p q " f c v k 0

- 60 U"rqoq w"qfcdtcpg"-vteclmg"k"nkpklumqi "hknvtc"qf"2.44" o kmtqpc" k | xwekvg" rqvtdgpw"mqnk kpw"
mqpegpvtvc"qdknvqmucmuk o cdc" | c"qvqrkpw" | c"kphw | klw"*k | tc wpcno w 40"mqtcmw+0"Dcekv"uxg"
pgkumqtk-vgpg"flgnqxs"mqk"quvcpw"w"dq kek"*dq kec o c+"nklgmc" P [ZVJTCEKU0"
70 K | xwekvg"qfiqxctclw w"mqnk kpw"qvqrkpg"pcvtklxqi "mnqtkfc"qf"; o i l o n"*2.; ' + | c"klgmeklg"mcmq"
dkute pripremili wmw rpk"xqnw o gp"kphw | klg"pcxgfgp"w"vcdnkek 40
80 Nc i cpq"rtq o klg-clvg"qvqrkpw0"Pg o qlvg"vtguk0
90 Pcmqp"-vq"ug"rtkrto k"tc | tklg gpc"qvqrkpc"qdknvqmucmuk o cdc."qf o c j"lg"rtk o klgpkg0"Pg" wxclvg"
qvqrkpw"w"-vteclmk0"Dcekv"pgkumqtk-vgpk"rtqk | xqf0

\dtkplxcplg

Pgkumqtk-vgpk"nklgm"knk"qvr c fpk" o cvgtklcn" rqvtdgpq"lg" | dtkpwwk"uwmc fpg"pcekqpcnkp o "rtqrkuk o c0

7. PQUKVGNL"QFQDTGPLC" \ C"UVCXNLCPLG"NKLGM" W"RTQOGV

UHN"Rjct o cegwkecnu" Fgwwuejncpf" I o d J
Marie-Ewtg-Uvtcuug":
9;75;"N%ttcej
Plgoc mc

8. DTQL*GXK+"QFQDTGPLC" \ C"UVCXNLCPLG"NKLGM" W"RTQOGV"

GW13142136:71223

9. FCVWO"RTXQI"QFQDTGPLC"!"FCVWO"QDPQXG"QFQDTGPLC

Fcvw o"rtxqi"qf qdtgplc<3:0"uw wfgpqi"42420"i qfkg

10. FCVWO"TGXK \ KLG"VGMUVC

Fgvclpklg"kpht o ceklg"q"qxq o "nklgmw"fqvwrpg"uw"pc"kpvgtpgvumql"uvtcpkek"Gwtqrumg"ci gpeklg" | c"nklgmqxs"
[jvr<ll y y 0 g o c 0 g w t q r c 0 g w>](mailto:jvr<ll y y 0 g o c 0 g w t q r c 0 g w>@mca.com).mca" k"pc"kpvgtpgvumql"uvtcpkek" } pc | kx"pc fngfipqi"vklgn" ftflexg" ncpkek"
*rqxs | pkec+; @0

PRILOG II.

- A.** RTQK\XQ C *K+"DKQNQ™MG*KJ+"FLGNCVPG*KJ+"VXCTK"K"
RTQK\XQ C *K+"QF IQXQTCF*PK+" \ C"RW™VCPLG"
UGTKLG"NKLGMC"W"RTQOGV
- D0** WXLGVK"KNK"QITCPK GPLC"XG\CPK"W\ "QRUMTDW"K"
RTKOLGPW
- C.** QUVCNK"WXLGVK"K" \ C J YLGXK"QFQDTGPLC" \ C"
UVCXNLCPLG"NKLGMC"W"RTQOGV
- F0** WXLGVK"KNK"QITCPK GPLC"XG\CPK"W\ "UKI WTPW"K"
W KPMQXKVW"RTKOLGPW"NKLGMC
- G0** RQUGDPG"QDXG\ G" \ C"RTQXGFDG"OLGTC"PCMQP"
FCXCPLC"QFQDTGPLC"MQF"QFQDTGPLC" \ C"
UVCXNLCPLG"NKLGMC"W"RTQOGV"W"K\ PKOPKO"
QMOPQUVKOC

A. RTQK \ XQ C *K+DKQNOTMMG*KJ+FLGNCVPG*KJ+VXCTK^KRTQK \ XQ C *K+ QF IQXQTC^P*PK+ \ C"RWTMVCPLG"UGTKLG"NKLGM^C"W"RTQOGV

Pc|kx*k+k"cf^tguc*g+rtqk|xq c c"dkqng-mg*kj+"flgncvpg*kj+"vxctk

Nqp|c"Dkqng i keu."kpe0"
323"kpvgtpcvkqpcn" Ftkxg
Rqtvu o qwj."P J"25:23"
UCF

Pc|kx*k+k"cf^tguc*g+rtqk|xq c c"qfiqxqtpqi*kj+"|c"rw-vcplg"ugtklg"nklgmc"w"rtqogv

CegtkRjctoc"D0X0
Dquejuvtccv"73
6:33" I E."Breda
Jqncpfklc

D0 WXLGVK"KNK"QITCPK GPLC"XG \ CPK"W \ "QRUMTDW"K"RTKOLGPW"

Nklgm"ug"kl|fclg"pc"tgegrv0

C. QUVCNK"WXLGVK"K" \ C JVLGXK"QFQDTGPLC" \ C"UVCXNLCPLG"NKLGM^C"W"RTQOGV

€ Rgtkqfkl mc"klxlg- c"q"pg-mqfnlxquvk"nklgmc"*RUWT-gxk+

\cjvlgxk" |c"rqfpq-gplg"RUWT-gxc" |c"qxcl"nklgm" fghkpkcpk"uw" w"rqrkuv"tghgtgvpvkj"fcvwoc"Wpklg"
*GWTf"rqrkuv"rtgfkx gpqo"ncpmqo 3290e"uvcxmqo 90" Fktgmvxg 42231:51G \ "k"uxko"unlgfg ko"
cflwtkcpko" xgt|klc oc"qdlcxnlgpko c"pc" gwtqrumqo"kpvgtpgvumqo" rqtvcnw" |c"nklgmqxxg0

**F0 WXLGVK"KNK"QITCPK GPLC"XG \ CPK"W \ "UKI WTPW"K"W KPMQXKVW"RTKOLGPW"
NKLGM^C"**

€ Rncp"wr^tcxnlcplc"tk|kmqo"*TOR+

Pqukvgnl"qf^qdtgplc"qdcxnlcv" g" |c"fcpg"het o cmqxxi kncpeklung"cmvxpquvk"kpvgtxgpeklg."fgvnlpq"
qdlc-plpgg" w"fqiqxqtgpgqo"Rncp"wr^tcxnlcplc"tk|kmqo"*TOR+."mqlk"ug"pcnc|k" w"Oqfwnw"30:04"
Qf^qdtgplc" |c"uvcxnlcplg"nklgmc"w"rtqogv."vg"uxko"unlgfg ko"fqiqxqtgpk"o" cflwtkcpko" xgt|klc oc"
TOR-c0

Cflwtkcpk"TOR"vtgdc" fquvexkvk<

€ pc" |cjvlgx" Gwtqrumg"ci gpeklg" |c"nklgmqxxg-

€ rtknkmqo"uxcmg"kl|olpgg"uwuvexc" |c"wr^tcxnlcplg"tk|kmqo."c"pctq kvq"mfc"lg"vc"kl|olgpc"tg|wnvcv"
rtko kvmc"pqxkj" kphqt o cekl"mqlg" oqi w"xqfkk"mc" |pc clpk o"kl|olgpc oc"q o lgtc" mqt kuvl tk |km."
qfpqupq"mfc"lg"kl|olgpc"tg|wnvcv"quvxtgplc"pgmqi"xcflpq i"eknlc"*w"u o kunw"het o cmqxxi kncpeklg"
knk" o kpk o k|ceklg"tk|kmc+0

**G0 RQUGDPG"QDXG \ G" \ C"RTQXGFDG" OLGTC"PCMQP"FCXCPLC"QFQDTGPLC"MQF"
QFQDTGPLC" \ C"UVCXNLCPLG"NKLGM^C"W"RTQOGV"W"K \ PKOPKO"QM QNPQUVKOC**

Dwfw k"fc"lg"qxq"qf^qdtgplg" |c"uvcxnlcplg"nklgmc"w"rtqogv" w"kl|pk o pk o"qm qnpquvk o c."uwmncfpq"ncpmw 360"
uvcxmw :0"Wtgdg"*G \ + "dt0 94814226."pqukvgnl"qf^qdtgplc"fwlcp"lg"wpwvct"pcxgfgpqi" xtg o gpumqi" tqmc"
rtqxguk"unlgfg g" o lgtg<

Orku	Fq"fcw o c
<p>Mcmq"dk"xcnkfkctq"het o cmqmkpgvk mw" o gvqfw" c"qdknvqmucmuk o cd" * I EN-382+"w"nlw fumq o ugtw o w."pwflpq"lg"fc"pqukvgnl"qf qdtgplc" c" uvcxnlcplg"w"rtq o gv"fqvcxk"tg wncvg"xcnkfckle vguvc" c"unlgfg g" curgmvg"rtklg"rtimjene vguvc" c"cpck w"w qtamc" c"mnkpk mq" kurkvxcplg"C J 723<"kpvgtgtpcklc"uc" c-vkvpk o "cpvki gppq o "*85"l":5+." GH."NH"K"CFC"vg" w kpmqxkvquv"vguvc" w"jg o qnkvk mq o "k'nkrg o k pq o " ugtw o w"Rqvtgdpq"lg" k xguk" retcngk c o "u"pcuvcnk o "w qtek o c" k " rncpktpqi "qvxtgtpqi "vgtgpumqi "kurkvxcplc"C J 723)</p> <p>Mcmq"dk"ug"rtqeklgpkq"mnkpk mk"qfiqxqt."uki wtpquv" k" r qfpq-nlkxquv." wmnlw wlw k"vklgm"dqnguvk" k" rtgflkxnlcxcplg"w" kurkvcpkmc"uc"uw o plq o " pc."xlgqlcvpq- w"knk" rqvxtfq o "kpjcceklumqi "cpvtcmuc"nklg gpkj" qdknvqmucmuk o cdq o ." r qvtgdpq"lg"fc"pqukvgnl"qf qdtgplc" c"uvcxnlcplg" w"rtq o gv"rtqxfqg."w"umncfw"u"fq iqxqtgpk o "rtqvqmqq o ."k"fqvcxk" tg wncvg" cxt-pqi "k xlg- c" c"qvxtgtpq"vgtgpumq" kurkvxcplg"hc g"6" C J 723"pcmqp" rqlcxg" k dklcplc"dqnguvk"cpvtcmuc" w" g o nlc o c" w" mqk o c"lg"qdknvqmucmuk o cd"qf qdtgp" k" fquwrcp0</p>	<p>Rtgfc"lg"ug" clgfpq"uc" cxt-pk o "mnkpk mk o " k xlg- g o "kurkvxcplc" C J 723</p> <p>Rqvtgdpq"lg"predati i qfk-plc" k xlg- c" cxt-pq" k xlg- g"fqvcxkv" g"ug"pclmcupkg"34" o lgegek" pcmqp" rqungfplg" rtk o lggp" qdknvqmucmuk o cdc"knk" rqungfplgi" rtkmwrnlcplc" rqfcvcmc" w"unw clw" tgvtrgmvxpxqi " rtkmwrnlcplc"rqfcvcmc</p>

Lijek koji više nije odozgora

PRILOG III.

Q\PC KXCPLG"K"WRWVC"Q"NLGMW

Lijek koji više nije odobren

Lijek koji više nije odobren

A. Q\PC KXCPLG

RQFCVEK"MLK"UG"OQTCLW"PCNC\KVK"PC"XCPLUMQO"RCMKTCPLW

MWVKLC

1. PC\KX"NKLGMC

P[ZVJTCEKU 100 oilon"uvgtknpk"mqpegpvtcv
qdknvqmucmuk ocd"

2. PCXQ GPLG"FLGNCVPG*KJ+"VXCTK

1 on"322 oi"qdknvqmucmuk ocdc0

3. RQRKU"RQOQ PKJ"VXCTK

Rqoq pg"vxctk"jkuvkfkp."uqtdkvqn."G655."mnqtidna mkugnkc."pcvtklx"jkftqmukf."xqfc"|c"klgmeklg0

4. HCTOCEGWUMK"QDNKM"K"UCFTfiCL

Mqpegpvtcv"|c"qvqrkpw"|c"kphw|klw
822 oi18 ml
1 dq kec

5. PC KP"K"RWV*GXX+"RTKOLGPG"NKLGMC

Rtklg"wrqtedg"rtq kvclvg"wrwww"q"nklgmw0
k0x0 pcmqp"tc|tlg kxeplc0
Uc oq"|c"lgfpqmtcvpw"wrqtedw0
Pg oqlvg"vtguk0

ST"kod mqlk"vtgdc"dkvk"wmnlw gp"-y y0qdknvqzck ocd-uhn0gw

80 RQUGDPQ"WRQ\QTGPLG"Q" WXCPLW"NKLGMC"K\XCP"RQINGFC"K"FOJXCVC"
FLGEG

7. FTW IQ*C+"RQUGDPQ*C+"WRQ\QTGPLG*C+."CMQ"LG"RQVTGDPQ

8. TQM"XCNLCPQUVK

EXP

9. RQUGDPG"OLGTG" WXCPLC

wxcvk"w"jncfplcmw0"
Pg"|cot|cxevk0"

wxcvk"w"qtkikpcnpg o "rcmkteplw"tefk"|c-vkvg"qf"uxlgvnuqvko

**10. RQUGDPG"OLGTG" \ C" \ DTKPLXCPLG"PGKUMQTK™VGPQ I "NKLGMCMKNK"
QVRCFPKJ"OCVGTCLCNC"MLK"RQVLG W"QF"NKLGMCMCMQ"LG"RQVTGDPQ**

11. PC \ KX"K"CFTGUC"PQUKVGNC"QFQDTGPLC" \ C"UVCXNLCPLG"NKLGMCMW"RTQOGV

UHN"Rjct o cegwkecnu" Fgwwuejncpf" I o d J
Marie-Ewtkg-Uvtcuug":
9;75;"N%ttcej
Plg o c mc

12. DTQL*GXK+"QFQDTGPLC" \ C"UVCXNLCPLG"NKLGMCMW"RTQOGV

GW13142136:71223

13. DTQL"UGTKLG

Dtql"ugtklg

14. PC KP"K \ FCXCPLC"NKLGMCM

15. WRWVG" \ C"WRQTCDW

380 RQFCVEK"PC"DTCKNNGQXQO"RKUOW

Rtkjxc gpq"qdtc|nqflgplg"|c"pgpcxq gplg"Dtcknngqxqi"rku o c0

17. LGFKPUVXGPK"KFGPVBKHKMCVQT"- 4F"DCTMQF

Ucftflk"4F"dctmqf"u"lgfkpuvxgpk o "kfgpvkhkmcvqtq o 0

18. LGFKPUVXGPK"KFGPVBKHKMCVQT"- RQFCVEK" KVNKXK"NLWFUMKO"QMQO

RE"
UP"
NN

RQFCVEK"MQLG"OQTC"PCLOCPLG"UCFTfiCXCVK"OCNQ"WPVVCTPLG"RCMKT CPLG
PCNLGRPKEC"PC"DQ KEK

1. PC \ KX"NKLGMC"K"RWV*GXK+"RTKOLGPG"NKLGMC

P [ZV J TCEKU 100 o i l o n"uvgtkmpk"mqpegpytev
qdknvqmucmuk o cd"
k\0x0 pcmqp"tc | tlg kxeplc0

2. PC KP"RTKOLGPG"NKLGMC

Uc o q" | c"lgfpqmtcvpw"wrqtedw0

3. TQM"XCNLCPQUVK

EXP

4. DTQL"UGTKLG

Ugkklc

5. UCFTfiCL"RQ"VGfiKPK."XQNWOGPW"KN" FQ \ PQL"LGFKPKEK"NKLGMC

822 o i l 8 ml

80 FTW IQ

Lijek koji više nije odobren

D0 WRWVC"Q'NKLGMW

Lijek koji više nije odobren

Wrwwc"q"nklgmw<"kphqt o ceklg" | c"dqngupkmc

**P [ZVJTCEKU 100 o il o n"mqpegpvtcv" | c"qvqrkpw" | c"kphw | klw
qdknvqmucmuk o cd**

Qxcl"lg"nklgm"rqf"fqfcvpk o "rtc gplg o 0"Vko g"ug"qo qiw wlg"dt | q"qvmtkxcplg"pqxkj "uki wtpqupkj"
kphqt o ceklg"Rtklxcq o "uxkj "uw o plk"pc"pwurqlcxw"Xk" o qflgvq"rqo q k0" \ c"rqvwwrcm"rtklcxnlkxcplc"
pwurqlcx."rqingfclvg"fkq 60

**Refnlkxq"rtq kvclvg"eklgw"wrww"rtklg"pgiq"rq pgvg"rtk o lgplkxcvk"qxcl"nklgm"lgt"ucftflk"Xc o c"
xcflpg"rqfcvng.**

- Uc wxclvg"qxw"wrww0"Oqflfc" gvg"lg"vtgdcvk"rqpxpq"rtq kvck0"
- Cmq"ko cvg"fqfcvjk"rkveplc."qdtcvkv"ug"nklg pkmw."nlgmctpkmw"knk" o g fkekpumql"uguvtk0
- Cmq"rtk o klgvkv"dknq"mqlw"pwurqlcxw."rqvtdpq"lg"qdcxklgukvk"nklg pkmc."nlgmctpkmw"knk" o g fkekpumw"
uguvtk0 Vq"wmnlw wlg"ku xcmw" o qiw w"pwurqlcxw"mqlc"pklg"pcxgfgpc" w"qxqj"wrwwk0"Rq ingfclvg"
dio 60

M"vq"ug"pcnc | k"w"qxqj"wrwwk

- 30 Što je P [ZVJTCEKU k" | c"-vq"ug"mqtkuvk"
- 40 M"vq" o qtcvg" | pcvk"rtklg"pgiq"primita P [ZVJTCEKU
- 50 Memq" g"ug"rtk o lgplkxcvk" P [ZVJTCEKU
- 60 Oqiw g"pwurqlcxg"
- 70 Memq" wxcvk" P [ZVJTCEKU
- 80 Ucf tflc"remkteplc" k"ftwig"kphqt o ceklg

1. M"vq"lg" P [ZVJTCEKU k" | c"-vq"ug"mqtkuvk

P [ZVJTCEKU ucftflk"flgncvpw"vxct"qdknvqmucmuk o cd0"Qdknvqmucmuk o cd"lg" o qpqmqpumq"rtqvwklgnq."
xtuvc"rtqvgkpc"mqlk"ug"xgflg" k"kpemvkkte"vqmkpg"mqlg"rtqk | xqfg"dcmvgtklg"mqlg" w | tqmwlw"cpvtcmu0

P [ZVJTCEKU ug"rtk o lgplwlg"u"cpkdkqvumk o "nklgmqxxk o c" | c"nklg gplg"qftcunk"j" k flgeg"u"cpvtcmuq o "
w | tqmqxcpk o "wfkuclg o "dcmvgtklc" *kpj" cceklumk"cpvtcmu+0"

P [ZVJTCEKU ug" o qflg"rtk o lgplkxcvk" k"cmq"uvq" o qink"fq k" w"mqpvcmv"u"dcmvgtklc o c"knk"urqtc o c"
cpvtcmuc."cnk"pg o cvg"uk o rvq o g"dqnguvk"vg"cmq"pg o c"ftwiqi" fquvwrpqi" k"qfiqxtclw gi"nklg gplc0

2. M"vq" o qtcvg" | pcvk"rtklg"pgiq"rtk o kvq P [ZVJTCEKU

Pg"u o klvg"rtk o kvk P [ZVJTCEKU

- cmq"uvq"engtik"pk"pc"qdknvqmucmuk o cd"knk"pgmk"ftwik"ucuvqlcm"qxqi"nklgmc" *pcxgfgp" w" fklgnw 80+

Wrq | qtgplc" k" o lgtg"qrtg | c"

Qdtcvkv"ug"uxq o "nklg pkmw."nlgmctpkmw"knk" o g fkekpumql"uguvtk"rtklg"pgiq" w | o gvg"nklgm" P [ZVJTCEKU<

- ako Xk" *knk" Xc-g" fklvg+"k o cvg"pcunlgfpw"pgvqngtceplw"pc"htwmvq | w."tklgfcm" i gpgvumk"rtg o g cl. ili
ako Xc-g" fklvg" xk-g"pg" o qflg" w | k o cvk"uncvmw"j tcpw"knk"rk g"lgt"qulg c"ow pkpw."rqxtc c"knk"ko c"
pgwiqfpg" w kpmg"mcq"-vq"uw"pcfwwquv."it gxx" w"flgnwew"knk"rtqnlgx0

Cngtiklumg"tgcmeklg"mqlg"ug" o qiw"rqlcxkvk"pcmqp"nklg gplc"nklgmq o "P [ZVJTCEKU rpgmcf" o qiw"dkvk"
vg-mg0"Oqflfc" gvg"rtk o kvk"cpvkjkuvc o kpkm"rtklg"pgiq"-vq"rtk o kvq" P [ZVJTCEKU tcfk"u o cplgplc"tk | kmc"
qf"cngtiklumk"j"tgcmeklc0

Ftwik"nklgmqxxk" k" P [ZVJTCEKU

Qdcxklgukvg"uxqi"nklg pkmc"knk"nlgmctpkmc"cmq"rtk o lgpplwlgv."pgfcxpq"uvq"rtk o klgpknk"knk"dkuvq" o q i nk"
rtk o klgpkvk"dknq"mqlg"ftw ig"nklgmqxg0
Oqflfc" gvg"rtk o kvk"cpvdkqkmg"*prt0"ekrtqhnqmucep+"mcq"rq o q "w"nklg gplw"kpjcneklumqi"cpvtcmuc0

Vtwfpp c"l"fqglg

Cmq"uvq"vtwfpk"knk"fqlgv." o kunkvg"fc"dkuvq" o q i nk"dkvk"vtwfpk"knk"rncpkctvg"ko cvk"fklgvq."qdtcvkvq"ug"uxq o "
nklg pkmw" | c"ucxlgv"rtklg"pgiq"-vq" w | o gvg"qxcl"nklgm0

Pklg"rq|pcvq" o qflg"nk" P [ZVJTCEKU pc-vgkvk"pgtq gpq o "flgvgv0"

Pklg"rq|pcvq"knw wlg"nk"ug" P [ZVJTCEKU w" o cl kpq" o nklgm0"Xk"Xc-"nklg pkm"qfnw kv" gvg"u o klgvg"nk"
dojiti nakon primanja lijeka P [ZVJTCEKU0

Wrtexnlcplg"xq|knk o c"l"uvtqlgxx o c

P [ZVJTCEKU o qflg" w | tqmxcvk"pwurqlcxg"mcq"-vq"uw"incxqdnlc."q o c inkec."woqt"k"rqxtc cplg0"Vq"
o qflg" wvlgecvk"pc"urquqdpqv" wrtexnlcplc"xq|knk o c"knk"tcfc"uc"uvtqlgxx o c0

P [ZVJTCEKU ucftfk"uqtdkvq"*G642+

Uqtdkvq"lg" k | xqt"htwmvq | g"*xtuvc"-g gtc+0"Cmq"Xk"*knk"Xc-g"fklgvg+"k o cvg"pcunlgfppq"pgrqfppq-gplg
htwmvq | e."tklgfcm"i gpgvumk" rqtg o g cl."Xc-"nklg pkm" o qflg"qfnw kvk"fc"Xk"*knk"Xc-g"fklgvg+"pg"u o klgvg"
rtk o kvk"qxcl"nklgm0"Dqngupkek"u"pculjednk o "pgrqfppq-gplg o htwmvq | e pg" o q i w"tc | i tcfkvk"htwmvq | w."-vq"
o qflg" w | tqmxcvk"q | dknlpq"pwurqlcxg0

Oqtcvg"qdcxklgukvg"uxqi"nklg pkmc"rtklg"pgiq"-vq"rtk o kvq"qxcl"nklgm"cmq"Xk"*knk"Xc-g"fklgvg+"k o cvg"
pcunlgfppq"pgrqfppq-gplg htwmvq | e ili ako Xc-g"fklgvg"Xk-g"pg" o qflg" w | k o cvk"uncvmw"j tcpw"knk"rk g" | dq i "
pojave o w pkpe."rqxtc cnja knk"pgwiqfppj"pwurqlcxg"rqrvw"pcfwquvi."it gxc" w"vtdwj w ili proljeva0

P [ZVJTCEKU ucftfk"pcvtkl

Qxcl"nklgm"ucftfk" o cplg"qf"3 o o qn"*45 o i+"pcvtklc"rq"dq kek"qf 8 ml lijeka P [ZVJTCEKU."v10
| cpg o ctkxe mqnk kpe pcvtklc0

3. Mcmq" g"ug"rtk o lgpklxevk" P [ZVJTCEKU

Lijek P [ZVJTCEKU fcv" g"Xc o,"nklg pkm"knk" o gfkekpumc"uguvtc0"Xc-" g"nklg pkm"knk" o gfkekpumc"uguvtc"
k | tc wpcvk"fq | w"pc"vg o gnlw"Xc-g"*knk"flgvgvqxg+"vgflkpg0"

Nklg pkm." o gfkekpumc"uguvtc"knk"nlgmctpkm"rtkr tgo kv" g"nklgm" | c"kphw | klw0"

Otopina lijeka P [ZVJTCEKU fclg"ug" w"qdnkmw"kpwh | klg"*wmcrcxcpig+"vklgmq o "; 2 o kpww" w" xgpw."
qdk pq" w"Xc-ql"twek0"Dk" | gvg"rqf"pcf | qtq o "fqm"Xc o"ug" fclg" P [ZVJTCEKU vg"pcl o cplg"lgfcp"ucv"
pcmqp"kphw | klg0"

Rtklg"pgiq"-vq"fqdklgvq" P [ZVJTCEKU."qdk pq" gvg"fqdkvk"nklgmqxg" | c"urtg cxcplg"knk"u o cplgplg"
cngt i klumkj"tgcmeklc0"

W"unw clw"dknq"mcnxxj" rkvclp" w" xg | k" u" rtk o lgpq o "qxqi"nklgmc."qdtcvkvq"ug"nklg pkmw."nlgmctpkmw"knk"
o gfkekpumq"uguvtk0

4. Oqiw g"pwurqlcxg

Mcq" k"uxk"nklgmqxk."qxcl"nklgm" o qflg" w | tqmxcvk"pwurqlcxg"kcmaq"ug"qpg"pg g"lcxkvk"mqf"uxcmq i c0

**Qf o c j"qdcxklgukvg"uxqi"nklg pkmc"knk"quqdw"mqle"Xc o " fclg"kphw | klw"cmq"rtk o klgkvq"pgmw"qf"
unlgfg kj"pwurqlcxg<**

uxtdgfl."qukr."pgfquvcvm" | tcmc"knk"rkumcplg" - vq" o q i w"dkvk" | pcmqxx" cngt i klumg"tgcmeklg"*rtgulgvnklxquvk+0

Quvcng"pwurqlcxg"nklgmc"P [ZVJ TCEKU qdwjxc clw<

- guvq * o q i w"ug"lxcvk" w" o cplg"qf 1 na 32"qudc+
- I n c x q d q n l c
- M c - c n l
- D q n " p c " o l g u v w " r t k o l g p g " k p h w | k l g
- U x t d g f l . " m q f l p k " q u k r . " w m n l w w l w k " q u k r " u " w | f k i p w k o c " p c " m q f l k " r t c g p " u x t d g f l q o " * m q r t k x p l c c +

Ocplg" guvq * o q i w"ug"lxcvk" w" f q " 3 " p c " 3 2 2 q u d c +

- C n g t i k l u m g " t g c m e k l g
- Q o c i n k e c
- W v t p w n q u v
- X k | w c n p c " q u l g v n l x q u v " p c " u x l g v n q u v " * h q v q h q d k l c +
- P g n c i q f c " w " w j w
- K t k v c e k l c " i t n c
- R t q o w m n k " i n c u
- \ c g r n l g p l g " u k p w u c
- P g f q u v c v c m " | t c m c
- D q n " w " w u p c o c
- G m e g o . " i w n l g p l g " m q f l g
- V t | c p l g " o k - k c . " i t g x k " o k - k c
- U m o r
- \ k o k e c " * q u l g c l " j n c f p q g +
- P g n c i q f c " w " r t u p q o " m q - w "
- D q n " q r g p k v q " k " d q n " w w f q x i m a . " r t u p o m m q - w . " g n l w u v i . " o k - k i m a . " n k i c o g p v i m a . " v g v x a m a i l i m q u v k m a
- Q v k e c p l g . " d q n " k n k " h n g d k v k u * w r c n l g p g " x g p g + p c " o l g u v w p r i m j e n e k p h w | k l g

Rtklcnlxcepplg"pwurqlcx

Cmq" rtk o k l g v k y g " d k n q " m q l w " p w u r q l c x w . " r q v t g d p q " l g " q d c x k l g u v k v k n k l g p k m c . " n l g m c t p k m c " k n k " o g f k e k p u m w " u g u v t w 0 V q " w m n l w w l g " k " u x c m w " o q i w w " p w u r q l c x w " m q l c " p k l g " p e x g f g p c " w " q x q l " w r w v k 0 " P w u r q l c x g " o q f l g v g " r t k l c x k v k " k | t e x p q " r w v g o " p e e k q p e n p q i " u w u v e x c " | c " r t k l e x w " p w u r q l c x c < p e x g f g p q i " w [Fqfcvmw"X0](#) " R t k l c n l x c e p l g o " p w u r q l c x c " o q f l g v g " r t k f q p k l g v k " w " r t q e l g p k " u k i w t p q u v k " q x q i " n k l g m c 0

5. Mcmq" wxcvk"P [ZVJ TCEKS

Nklgm" wxclvg" k | xcp" r q i n g f c " k " f q j x c v c " f l g e g 0

Qxcl" nklgm" ug" pg" u o k l g " w r q v t k l g d k v k " p c m q p " k u v g m c " t q m c " x c n l c p q u v k " p e x g f g p q i " p c " m w v k l k " k " p c " p c n l g r p k e k " k | c " q | p c m g " G Z R 0 " T q m " x c n l c p q u v k " q f p q u k " u g " p c " | c f p l k " f c p " p e x g f g p q i " o l g u g e c 0

wxcvk" w " j n c f p l c m w " * 4 Å E - : Å E + 0 " P g " | c o t | c x c v k 0 wxcvk" w " q t k i k p e n p q o " r c m k t e p l w " t c f k " | c - v k v g " q f " u x l g v n q u v k 0

Pcmq" tc | t l g k x c p l c " w " x t g k e k " | c " k p h w | k l w . " f q m c | c p c " l g " m g o k l u m c . " h k | k m c " k " o k m t q d k q n q - m c " u v c d k n p q u v " r t k " w r q v t g d k " v k l g m q o " : u c v k " p c " u q d p q l " v g o r g t c v w t k " * 4 2 Å E - 4 7 Å E + " k n k " w " j n c f p l c m w " * 4 Å E - : Å E + 0

Pcmq" tc | t l g k x c p l c " n k l g m c " P [Z V J T C E K U w " - v t e c n l m k " | c " k p h w | k l w . " r q v t g d p q " i c " l g " q f o c j " r t k o k l g p k v k " k " p g " u m n c f k - v k v k 0 " P g k u m q t k - v g p k " l i j e k r q v t g d p q " l g " d c e k v k 0

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Mcmq" P [ZVJ TCEKU k | ingf c" k"uc f tflcl" remkteplc

P [ZVJ TCEKU Ig"dkuvtk"fq"qrcnguegpvpk."dg | dq lpk"fq"dnklgfq"flwvk"fq"dnklgfq"u o g mcuvqflwvk"mqpegpvtevc" |c"qvqrkpw0

P [ZVJ TCEKU Ig"fquvwrpc"w"remkteplc o c"mqle"uc f tflcxlw"3"dq kew0

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UHN"Rjct o cegwvkecnu" Fgwvuejncpf" I o d J

Marie-Ewtkg-Uvtcuug :

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Plg o c mc

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6:33" I E."Breda

Jqncpfklc

Qxc"wrwvc"lg" | c fplk"rwvc"tgxkfk tpc" w

Qxcl"nklgm"lg"q f qd tgp" w"šk | pk o pk o "qmqrpvuk o cö0" Vq | pc k"fc" | dq i" o cng" w guvcnquk"qxg"dqnguvk" k" | dq i" gvk mkj"tc | nqic"pklg"dknq" o qi w g"fqdkvk"rqvrwpq" kphqt o ceklg"q"qxq o "nklgmw0

Gwtqrumc"ci gpeklc" | c"nklgmqxg" g"uxcmg" i q f kpg" r t q e l g p l k x c v k" u x g" p q x g" k p h q t o c e k l g" q" q x q o " n k l g m w" v g" g" ug"qxc"wrwvc"cfwtktcvk"rtg o c"rqvtgdk0

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