

!" #&/
4;O <548G=>3> 70AB>AC20==O ;V:0@AL:>3> 70A>1C

"□ "□®
(TEVETEN®)

!,:04:

4VNG0 @5G>28=0: 5?@>A0@B0=;
1 B01;5B:0 A0@B0=C <578;0BC 735,8 <3 (I> 2V4?>2V40T 5?@>A0@B0=C 600 <3);
4>?>28=8: O4@> B01;5B:8 ;0:B>78 <=>3V4@0B, F5;N;>70 :@8AB0;VG=0, :@>E<0;L
?@565;0B8=V7>20=89, <03=VN AB50@0B, :@>A?>2V4=>; >1>;>=:0 ?04@V* 1V;89 OY-S-9603 (3V?@>
<5;>70 (464), <0:@>3>; 400, ?>;VA>@10B 80 (433), B8B0=C 4V>:A84 (171)).

V:0@AL:0 D>@<0. "01;5B:8, 2:@8BV ?;V2:>2>N >1>;>=:>N.
A=>2=V DV78:>-EVABV: 1V;V, C D>@2>N >1>;>=:>N B01;5B:8 7 <0@:C20==O< «5046» 7 >4=>3> 1>:C V
157 <0@:C20==O - 7 V=H>3>.

\$0@<0:>B5@0?52B8G=0 3@C?0. @>ABV ?@5?0@0B8 0=B03>=VABV2 0=3V>B5=78=C II.
>4 "% C09C A02.

\$0@<0:>;>3VG=V 2;0AB82>ABV.

\$0@<0:>48=0
?@>A0@B0= T ?>BC6=8<, =5?5?B84=8<, 0:B82=8< ?@8 ?5@>@0;L=>, =5B5B@07>;L=8< 1>;>0B>@<<
@5F5?B>@V2 0=3V>B5=78=C II, O:89 A5;5:B82=> 72'O7CTBLAO 7 AT1-@5F5?B>@0<8. =3V>B5=78= II T
?>BC6=8< 207>:>=AB@8:B>@<< V >A=>2=8< 0:B82=8< 3>@<=><< @5=V=-0=3V>B5=78=-
0;L4>AB5@>=>2>W A8AB5<8. V= 2V4V3@0T 206;82C @>;L C ?0B>35=57V 0@B5@V0;L=>W
3V?5@B5=7VW. =3V>B5=78= II 72'O7CTBLAO 7 AT1-@5F5?B>@0<8 C @V7=8E >@30=0E V B:0=8=0E
(=0?@8::04 C 3;04:8E <'O70E AC48=, =04=8@:>28E 70;>70E, =8@:0E, A5@FV) V 28O2;OT B0:V 206;82V
1V>;>3VG=V 5D5:B8, O: 72C65==O :@>2>=>A=8E AC48=, 70B@8AB5@>=C. =3V>B5=78= II 15@5
CG0ABL C D>@DVW A5@FO V AC48= G5@57 9>3> 2?;82 =0 @VAB 3;04:8E<'O7>28E ;VB8= AC48= V
A5@FO.
?@>A0@B0= ?5@5H:>460T 2?;82C 0=3V>B5=78=C II =0 0@B5@V0;L=89 B8A:, =8@:>289 :@>2>BV: V
284V;5==O 0;L4>AB5@>=C C 74>@>28E >AV1. =865==O 0@B5@V0;L=>3> B8A:C 715@V30TBLAO
?@>B03>< 24 3>48= 157 28=8=:5==O ?>ABC@0;L=>W 3V?>B5=7VW ?VA;O ?@89>W 4>78. @8?8=5==O
;V:C20==O 5?@>A0@B0=<=5 A?@8G8=OT @V7:>3> ?V428I5==O 0@B5@V0;L=>3> B8A:C (A8=4@>
E2>@8E =0 0@B5@V0;L=C 3V?5@B5=7VN 7=865==O 0@B5@V0;L=>3> B8A:C ?VA;O ?@89>
A0@B0=C =5 AC?@>2>46CTBLAO 7N G0AB>B8 A5@F528E A:>@>G5=L.
V4 G0A 2V4:@8B>3> A;V?>3> @0=4>20=>3> ?@>A?5:B82=>3> 4>A;V465==O MOSES
(70E2>@N20=VABL V A<5@B=VABL ?VA;O V=AC;LBC, 5?@>A0@B0= ?>@V2=N202AO 7
=VB@5=48?V=>< 4;O 2B>@8==>W ?@>DV;0:B8:8) 1405 E2>@8E =0 0@B5@V0;L=C 3V?5@B5=7VN 7
F5@51@>20A:C;O@=8<8 ?>4VO<8 2 0=0<=57V ;V:C20;8AO 01> 5?@>A0@B0=<, 01> =VB@5=48?V=>
A0@B0=C 78 % ?0FVT=BV2 >B@87C 600 <3 I>45==> V 12 % 4>7C 4> 800 <3 =0 4>1C. # 3@C?V
=VB@5=48?V=C 47 % >B@87V 10 <3 V 42 % 20 <3 =0 4>1C (11 % 4> 40 <3). 5@28==0 :<1V=>20=0
:V=F520 B>G:0 A::040;0AO V7 7030;L=>W A<5@B=>ABV, F5@51@>20A:C;O@=8E ?>4V9
(B@0=78B>@=0 VH54>265=89 >1>@>B=89 VH5;>3VG=89 45DVF8B, V=AC;LB), 0 B0:>6 A5@F52>-
AC48==8E CA::04=5=L (=5AB01V;L=0 AB5=>:0@4VO, V=D0@:B:0@40, A5@F520 =54>AB0B=VABL,
;535=520 5<1>;VO V ;5B0;L=V A5@F52V 0@8B4V9. ?L>289 0@B5@V0;L=89 B8A: 4>1@5 4>AO302AO 2
>1>E 3@C?0E ;V:C20==O B0 ?V4B@8B03>< 2AL>3> 4>A;V465==O. 0 ?5@28==>N :V=F52>N B>G:>N
1C;> ?>:070=> 4>AB>2V@=> :@0I89 @57C;LB0B 2 3@C?V 5?@>A0@B0=C (7=865==O @878:C =0 21 %).
?5@H>4V9 7=865==O @878:C AB0=>28;> 12 % 4;O F5@51@>20A:C;O@=8E V 30 % 4;O A5@F52>-
AC48==8E :V=F528E B>G>.. &V @57C;LB0B8 1C;8 4>AO3=CBV 3>;>2=8< G8=>< 70 @0EC=>: 7=865==O
G0AB>B8 B@0=78B>@=>W VH5W 0B0:8/?>4>265=>3> >1>@>B=>3> VH53> =52@>;>3VG=>3>
45DVF8BC, =5AB01V;L=>W AB5=>:0@4VW V A5@F52>W =54>AB0B=>ABV. 030;L=0 A<5@B=VABL
1C;0 <5=H>N 2 3@C?V =VB@5=48?V=C. "0:, 2 3@C?V 5?@>A0@B0=C ?><5@;> 57 ?0FVT=BV2 V7 681
?>@V2=O=> 7 52 ?0FVT=B0<8 V7 671 C 3@C?V =VB@5=48?V=C (2V4=>A=89 @878: 1,07; 95 % 0,73-
1,56; p=0,725). 5B0;L=89 V =5;5B0;L=89 V=D0@:B:0@40 @>728=C2AO C 18 ?0FVT=BV2 C 3@C?V

5?@>A0@B0=C ?>@V2=O=> 7 20 C 3@C?V =VB@5=48?V=C, V=AC;LB C 36 ?>@V2=O=> 7 42
?0FVT=B0<8 2V4?>2V4=>, I> A2V4G8BL ?@> 5D5:B82=VABL 5?@>A0@B0=C ?>@V2=O=> 7
=VB@5=48?V=>W :V=F52>W B>G:8 5D5:B 5?@>A0@B0=C 28O282AO 1V;LH 28@065=8< C ?0FVT=BV2,
O:V =5 >B@81;>:0B>@V2.
?0FVT=BV2 7 0@B5@V0;L=>N 3V?5@B5=7VTN 5?@>A0@B0= =5 2?;820T =0 @V25=L 3;N:>78,
B@83;VF5@84V2, E>;5AB5@8=C B0 ;V?>?@>B5W=V2 =87L:>W IV;L=>ABV C ?;072V =0BI5.
?@>A0@B0= =5 ?>@CHCT <5E0=V7: C ?0FVT=BV2 V7 5A5=V70;L=>N 3V?5@B5=7VTN B0 ?0FVT=BV2
V7 =8@:>2>N =54>AB0B=VABN. ?@>A0@B0= =5 7<5=HCT H284:VABL ;C1>G:>2>W DV;LB@0FVW C
74>@>28E G>;>2V:V2, C ?0FVT=BV2 V7 0@B5@V0;L=>N 3V?5@B5=7VTN B0 ?0FVT=BV2 V7
=8@:>2>N =54>AB0B=VABN @V7=>3> ABC?5=O B06:>ABV. ?@>A0@B0= 28;;8:0T
=0B@V9C@5B8G=89 5D5:B C 74>@>28E >AV1, O:V 7=0E>4OBLAO =0 =87L:>A>;L>2V9 4VTBV.
?@>A0@B0= <>6=0 157?5G=> 70AB>A>2C20B8 ?0FVT=B0< V7 5A5=V70;L=>N 3V?5@B5=7VTN V
E2>@8< V7 =8@:>2>N =54>AB0B=VABN @V7=>3> ABC?5=O B06:>ABV 157 A?@8G8=5==O 70B@8
?>3V@H5==O DC=:FVW =8@>:..
?@>A0@B0= VAB>B=> =5 2?;820T =0 284V;5==O A5G>2>W :8A;>B8.
?@>A0@B0= =5 ?>A8;NT 5D5:BV2, ?>2O70=8E 7 0:B820FVTN 1@048:V=V=C, >?>A5@54:>20=>N
0=3V>B5=78=?5@5B2>@NNG8< D5@<5=B>< (\$), =0?@8;;04 :0H;N. # E>4V 4>A;V465==O, <5B>N O:>3>
1C;> ?>@V2=O==O G0AB>B8 28=8:=5==O :0H;N C E2>@8E, O:V ;V:C20;8AO 5?@>A0@B0=×, V
E2>@8E, O:V ?@89<0;8 V=3V1VB>@ \$, G0AB>B0 ACE>3> ?>ABV9=>3> :0H;N C ?0FVT=BV2, O:V
;V:C20;8AO 5?@>A0@B0=× (1,5 %), 1C;0 4>AB>2V@=> =86G>N (@< 0,05), =V6 C ?0FVT=BV2, O:V
?@89<0;8 V=3V1VB>@ \$ (5,4 %). '0AB>B0 :0H;N 1C4L-O:>3> E0@0:B5@C 1C;0 B0:>6 7=0G=> =86G>N
(@< 0,05) C E2>@8E, O:V ?@89<0;8 5?@>A0@B0= (21,2 %), =V6 C E2>@8E, O:V ?@89<0;8 V=3V1VB>@
\$ (29,9 %). E>4V V=H>3> 4>A;V465==O, A?@O<>20=>3> =0 282G5==O G0AB>B8 28=8:=5==O :0H;N C
?0FVT=BV2, O:V @0=VH5 AB@0640;8 2V4 :0H;N ?@8 70AB>AC20==V V=3V1VB>@V2 \$, G0AB>B0
ACE>3> ?>ABV9=>3> :0H;N 1C;0 2,6 % C 3@C?V 5?@>A0@B0=C, 2,7 % C 3@C?V ?;0F51> V 25 % C
3@C?V V=3V1VB>@C \$. V7=8FO G0AB>B8 28=8:=5==O ACE>3> ?>ABV9=>3> :0H;N C 3@C?V
E2>@8E, O:V ?@89<0;8 5?@>A0@B0=, V 3@C?V E2>@8E, O:V ?@89<0;8 V=3V1VB>@ \$, 1C;0
AB0B8AB8G=> 4>AB>2V@=>N (@< 0,01), B>4V O: B0:>W @V7=8FV ?@8 70AB>AC20==V
5?@>A0@B0=C V ?;0F51> =5 1C;>. '0AB>B0 28=8:=5==O :0H;N 1C4L-O:>3> E0@0:B5@C 1C;0 B0:>6
VAB>B=> =86G>N (@< 0,01) ?@8 70AB>AC20==V 5?@>A0@B0=C, =V6 ?@8 70AB>AC20==V
V=3V1VB>@C \$, V =57=0G=> 2V4@V7=O;0AO ?@8 ?@89>. @V< B>3>, 7030;L=89 0=0;V7
@57C;LB0BV2 6 ?>42V9=8E A;V?8E ;V=VG=8E 4>A;V465=L, O:V 2;;NG0;8 1554 E2>@8E, ?@>45
<=>AB@C202, I> G0AB>B0 :0H;N 70 A?>=B0==8<8 ?>2V4><;5==O<8 C E2>@8E, O:V ?@89<0;8
5?@>A0@B0=, 1C;0 B0:>3> 6 A0<>3> @V2=O (3,5 %), I> 9 C E2>@8E, O:V ?@89<0;8 ?;0F51> (2,6 %).
0 40=8<8 B@L>E ;V=VG=8E 4>A;V465=L (n=791), 5D5:B 5?@>A0@B0=C =0 7=865==O 0@B5@V0;L=>3>
B8A:C 1C2 ?@8=09<=V B0:8< 65 A0<8<, O: V 5D5:B V=3V1VB>@C \$ 5=0;0?@8;C, 7 B5=45=FVTN 4>
1V;LH>W 5D5:B82=>ABV 5?@>A0@B0=C.
0AB>AC20==O V=3V1VB>@C \$ C :<<1V=0FVW 7 1;>:0B>@< @5F5?B>@0 0=3V>B5=78=C II
4>A;V465=> C E>4V 42>E <0AHB01=8E @5=4>20=8E :=B@>;L>20=8E 4>A;V465=L (ONTARGET
(3;>10;L=V 4>A;V465==O 70AB>AC20==O B5;<) B0 VA NEPHRON-D (4>A;V465==O 4V015B8G=>W
=5D@>?0BVW).
ONTARGET F5 4>A;V465==O, I> ?@>2>48;>AO 70 CG0ABN ?0FVT=BV2 7 0=0<=57>< A5@F52>-
AC48==>3> 01> F5@51@>20A:C;O@=>3> 70E2>@N20==O, 01> FC:@>2>3> 4V015BC B8?C, I>
AC?@>2>46C20;8AO >7=0:0<8 C@065==O >@30=V2-A;V465==O, I> ?@>2>48;>AO 70 CG0ABN
?0FVT=BV2 7 FC:@>28< 4V015B>< B8?C B0 4V015B8G=>N =5D@>?0BVTN.
8I52:070=V 4>A;V465==O ?>:070;8 2V4ACB=VABL VAB>B=>3> ?>78B82=>3> 2?;82C ?@8
70E2>@N20==V =8@>: B0/01> A5@F52>-AC48==>W A8AB5<8 01> =0 7=865==O @V2=O A<5@B=>ABV,
B>4V O: A?>AB5@V302AO ?V428I5=89 @878: 3V?5@:0;VTAB@>W =8@:>2>W =54>AB0B=>ABV B0/01>
7=865==O 0@B5@V0;L=>3> B8A:C ?>@V2=O=> 7 <=>B5@0?VTN. @0E>2CNG8 ?>4V1=VABL D0@
<0:>48=0AB59, FV @57C;LB0B8 <0NBL 7=0G5==O B0:>6 4;O V=H8E V=3V1VB>@V2 \$ B0 1;>:0B>@V2
@5F5?B>@0 0=3V>B5=78=C II.
72O7:C 7 28I528;;045=8<, =5 @5:><5=4CTBLAO >4=>G0A=5 70AB>AC20==O V=3V1VB>@C \$ B0
1;>:0B>@0 @5F5?B>@V2 0=3V>B5=78=C II ?0FVT=B0< 7 4V015B8G=>N =5D@>?0BVTN.
>A;V465==O ALTITUDE (70AB>AC20==O 0;VA:V@5=C ?0FVT=B0< 7 FC:@>28< 4V015B>< B8?C 7
28:>@8AB0==O< :V=F528E B>G>: 70E2>@N20=L =8@>: B0 A5@F52>-AC48==>W A8AB5<8) 1C;>

?@>2545=> 7 <5B>N 282G5==O :>@8ABV 2V4 4>4020==O 0;VA:V@5=C 4> AB0=40@B=>W B5@0?VW
V=3V1VB>@>< \$ 01> 1;>:0B>@>< @5F5?B>@V2 0=3V>B5=78=C II ?0FVT=B0< 7 FC:@>28< 4V015B><
B8?C B0 E@>=VG=>N =8@:>2>N =54>AB0B=VABN, 70E2>@N20==O<8 A5@F52>-AC48==>W A8AB5<8
01> 7 :><1V=0FVTN F8E 70E2>@N20=L. >A;V465==O 70:V=G8;>AO 4>AB@>:>2> G5@57 ?V428I5==O
@878:C 28=8:=5==O ?>1VG=8E @50:FV9. V4AO 71V;LH5==O G0AB>B8 28?04:V2 V=AC;LBC B0
;5B0;L=8E =0A;V4:V2, 0 B0:>6 28=8:=5==O ?>1VG=8E 5D5:BV2, C B.G. A5@9>7=8E (3V?5@:0;VT
B5=7VO B0 ?>@CH5==O DC=:FVW =8@>:), C 3@C?V 70AB>AC20==O 0;VA:V@5=C ?>@V2=O=> 7
3@C?>N, I> 70AB>A>2C20;0 ?;0F51>.

\$0@<0:>:V=5B8:0.

1A>;NB=0 1V>4>ABC?=VABL 5?@>A0@B0=C ?VA;O >4=>@07>2>3> ?5@>@0;L=>3> ?@89>78 300 <3
AB0=>28BL ?@81;87=> 13 % 2=0A;V4>: 9>3> >1<565=>W 01A>@1FVW. V: :>=F5=B@0FVW
5?@>A0@B0=C C ?;072V ?VA;O ?5@>@0;L=>3> ?@89>AB5@V30TBLAO G5@57 1-2 3>48=8. 0 40=8<8
4>A;V465==O D0@<0:>:V=5B8:8 5?@>A0@B0=C 70;56=> 2V4 4>78, :>=F5=B@0FVO 9>3> C ?;072V 1C;0
?@>?>@FV9=>N 4>7V ?@8 70AB>AC20==V 4>7 2V4 100 4> 200 <3 V <5=H ?@>?>@FV9=>N ?@8
70AB>AC20==V 4>7 2V4 400 4> 800 <3. 5@V>4 =0?V2282545==O 5?@>A0@B0=C ?VA;O ?5@>@0;L=>3>
70AB>AC20==O 70728G09 AB0=>28BL 5-9 3>48=. @8 B@820;>AC20==V 5?@>A0@B0= VAB>B=> =5
=0:>?8GCTBLAO. 0AB>AC20==O 5?@>A0@B0=C 7 W65N ?@872>48BL 4> =57=0G=>W 70B@8@1FVW
7V 7N <5=H =V6 =0 25 % ?>:07=8:V2 C_{max} V AUC, I> =5 <0T VAB>B=8E ;V=VG=8E =0A;V4:V2.
2O7C20==O 7 1V;:0<8 ?;07<8 :@>2V 5?@>A0@B0=C T 28A>:8< (?@81;87=> 98 %) V ?>ABV9=8< C
4V0?07>=V B5@0?52B8G=8E :>=F5=B@0FV9. 2O7C20==O 7 1V;:0<8 =5 70;568BL 2V4 AB0BV, 2V:C,
?>@CH5==O DC=:FVW ?5GV=:8 01> =0O2=>ABV ;53:>W/?>W =8@:>2>W =54>AB0B=>ABV C
?0FVT=B0, 0;5 7<5=HCTBLAO C 45O:8E ?0FVT=BV2 V7 BO6:>N D>@<N =8@:>2>W =54>AB0B=>ABV.
;N48=8 ?VA;O ?5@>@0;L=>3> B0 2=CB@VH=L>25==>3> 22545==O 14!-5?@>A0@B0=C 5?@>A0@B0=
1C2 T48=>N A?>;C>N, 7=0945=>N C ?;072V B0 :0;V. @81;87=> 20 % @04V>0:B82=>3> 5?@>A0@B0=C
5:A:@5BCTBLAO 7 A5G5N C 283;04V 0F8;3;N:C@>=V4C V 80 % C 283;04V =573> 5?@>A0@B0=C.
1□T< @>7?>4V;C 5?@>A0@B0=C AB0=>28BL ?@81;87=> 13;. 030;L=89 ?;07<>289 ;V@5=A 4>@V2=NT
?@81;87=> 130 <;/E2. ?@>A0@B0= 5:A:@5BCTBLAO 7 A5G5N V 6>2GN. VA;O 2=CB@VH=L>25==>3>
22545==O 14!-5?@>A0@B0=C ?@81;87=> 61 % @04V>0:B82=>ABV 28O2;OTBLAO C :0;V V ?@81;87=>
37 % C A5GV. VA;O ?5@>@0;L=>3> ?@89>14!-5?@>A0@B0=C ?@81;87=> 90 % @04V>0:B82=>ABV
28O2;OTBLAO 2 :0;V V ?@81;87=> 7 % C A5GV.
=0G5==O AUC V C_{max} 5?@>A0@B0=C 71V;LHCNBLAO 2 >AV1 ;VB=L>3> 2V:C (C A5@54=L>C 2
@078), 0;5 F5 =5 ?>B@51CT 77C20==O.
=0G5==O AUC 5?@>A0@B0=C 71V;LHCNBLAO (2 A5@54=L>=0 40 %) 157 7_{max} C E2>@8E V7
?5GV=:>2>N =54>AB0B=VABN, I> B0:>6 =5 ?>B@51CT 77C20==O.
=0G5==O AUC V C_{max} 5?@>A0@B0=C 1C;8 ?@81;87=> =0 30 % 28I8<8 C E2>@8E V7 ?>N =8@:>2>N
=54>AB0B=VABN (:;V@5=A :@50B8=V=C 30-59<;/E2) V =0 50 % 28I8<8 C E2>@8E V7 BO6:>N
=8@:>2>N =54>AB0B=VABN (:;V@5=A :@50B8=V=C 5-29<;/E2) ?>@V2=O=> 7 >A>10<8 7 =>@<0;L=>N
DC=:FVTN =8@>: .>4=8E 77C20==O =5 ?>B@V1=>.
6V=>: V G>;>2V:V2 D0@<0:>:V=5B8:0 5?@>A0@B0=C =5 2V4@V7=OTBLAO.

;V=VG=V E0@0:B5@8AB8:8.

>:070==O.

A5=FV0;L=0 3V?5@B5=7VO.

@>B8?>:070==O.

V4><0 3V?5@GCB;82VABL 4> 0:B82=>W @5G>28=8 01> 4> 1C4L-O:>3> =50:B82=>3> :>=5=B0
?@5?0@0BC.

03VB=VcBL 01> ?;0=C20==O 203VB=>ABV (482. @>74V; «0AB>AC20==O C ?5@V>4 203VB=>ABV 01>
3>4C20==O 3@C44N»).

2>AB>@>=V @5=>20A:C;O@=V 70E2>@N20==O 7V 7=0G=8<8 ?>@CH5==O<8 35<>48=0 BO6:89
AB5=>7 =8@:>2>W 0@B5@VW T48=>W DC=:FV>=CNG>W =8@:8.

4=>G0A=5 70AB>AC20==O ;V:0@AL:>3> 70A>1C "525B5=® 7 ?@5?0@0B0<8, I> 2) (482. @>74V;8 «70T
<>4VO 7 V=H8<8 ;V:0@AL:8<8 70A>10<8 B0 V=HV 2848 270T<>4V9» B0 «\$0@<0:>;>3VG=V
2;0AB82>ABV»).

70T<4VO 7 V=H8<8 ;V:0@AL:8<8 70A>10<8 B0 V=HV 2848 270T<4V9.

7207:C 7 B8<, I> C ?;0F51>-:=B@>;L>20=8E 4>A;V465==OE A?>AB5@V30;>AO ?V428I5==O @V2=O
:0;VN C A8@>20BFV :@>2V, B0 107CNG8AL =0 4>A2V4V 70AB>AC20==O V=H8E ?@5?0@0BV2, I>
2?;820NBL =0 @5=V=-0=3V>B5=78=-0;L4>AB5@>=>2C A8AB54=>G0A=5 70AB>AC20==O
5?@>A0@B0=C 7 :0;V9715@V30NG8<8 4VC@5B8:0<8, E0@G>28<8 4>102:0<8 B0 70 V=H8<8
?@5?0@0B0<8, O:V <>6CBL ?V428I8B8 @V2=V :0;VN C A8@>20BFV (=0?@8:;04 7 35?0@8=×), <>65
?@8725AB8 4> ?V428I5==O @V2=O :0;VN C A8@>20BFV.
0=V :;V=VG=8E 4>A;V465=L A2V4G0BL, I> ?>42V9=0 1;>;:040 @5=V=-0=3V>B5=78=-
0;L4>AB5@>=>2>W A8AB5<8 ?@8 :<1V=>20=>AC20==V V=3V1VB>@V2 \$, 1;>;:0B>@V2
@5F5?B>@V2 0=3V>B5=78=C 01> 0;VA:V@5=C ?@872>48;0 4> ?V428I5==O G0AB>B8 28=8:=5==O
?>1VG=8E @50:FV9, B0:8E O: 3V?>B5=7VO, 3V?5@:0;VT: (2 B>AB@0 =8@:>20 =54>AB0B=VABL),
?>@V2=O=> 7 <=>B5@0?VTN ?@5?0@0B>< 7 4VTN =0 @5=V=-0=3V>B5=78=-0;L4>AB5@>=>2C
A8AB574V;8 «@>B8?>:070==O», «A>1;82>ABV 70AB>AC20==O» B0 «\$0@<0:>;>3VG=V 2;0AB82>ABV»),
=B83V?5@B5=782=89 5D5:B 5?@>A0@B0=C <>65 ?>A8;N20B8AO V=H8<8 0=B83V?5@B5=782=8<8
?@5?0@0B0<8.
?>2V4><;5==O ?@> B>:A8G=VABL B0 >1>@>B=5 ?V428I5==O @V2=O ;VBN C A8@>20BFV :@>2V C
@07V >4=>G0A=>3> 70AB>AC20==O ?@5?0@0BV2 ;VBN 7 V=3V1VB>@0<8 \$. 5 <>6=0 28;;NG8B8
<>6;82VABL ?>4V1=>3> 5D5:BC ?@8 70AB>AC20==V 5?@>A0@B0=C, B>3> >4=>G0A=>3>
70AB>AC20==O 7 ?@5?0@0B0<8 ;VBN @5:><5=4CTBLAO :>=B@>;N20B8 @V2=V ;VBN C :@>2V.
?@>A0@B0= =5 28O2;OT *in vitro* 1;>;:CNG>3> 5D5:BC =0 D5@<5=B8 A8AB5<8 F8B>E@> CYP1A, 26, 2!9/8,
2!19, 2D6, 2B0 3.
/: V ?@8 70AB>AC20==V V=3V1VB>@V2 \$, >4=>G0A=5 70AB>AC20==O 1;>;:0B>@V2 @5F5?B>@V2
0=3V>B5=78=C 7 =5AB5@>W4=8<8 ?@>B870?0;L=8<8 ?@5?0@0B0<8 <>65 ?@872>48B8 4> ?V428I5=>3>
@878:C ?>3V@H5==O DC=:FVW =8@>;, 2;;NG0NG8 <>6;82VABL 3>AB@>W =8@:>2>W
=54>AB0B=>ABV B0 ?V428I5==O @V2=O :0;VN C A8@>20BFV, >A>1;82> C ?0FVT=BV2 7 C65
VA=CNG8<8 ?>@CH5==O<8 DC=:FVW =8@>.. "0:C :<1V=0FVN A;V4 70AB>A>2C20B8 7
>15@56=VABN, >A>1;82> ?0FVT=B0< ;VB=L>3> 2V:C. 0FVT=B8 ?>28==V 1CB8 045:20B=>
3V4@0B>20=V, =0 ?>G0B:C B5@0?VW B0 ?5@V>48G=> ?>B>B@>;N20B8 DC=:FVN =8@>..
!C?CB=O B5@0?VO ;>70@B0=>< 7 =5AB5@>W4=8< ?@>B870?0;L=8< ?@5?0@0B>< V=4><5B0F8=><
?@872>48;0 4> 7=865==O 5D5:B82=>ABV 1;>;:0B>@0 @5F5?B>@V2 0=3V>B5=78=C , =5 <>6=0
28;;NG0B8 :;0A>289 5D5:B 1;>;:0B>@0 @5F5?B>@V2 0=3V>B5=78=C .

A>1;82>ABV 70AB>AC20==O.

>@CH5==O DC=:FVW ?5GV=:8
@8 70AB>AC20==V 5?@>A0@B0=C ?0FVT=B0< 7 ?>@CH5==O< DC=:FVW ?5GV=:8 A;V4 1CB8
>A>1;82> >15@56=8<8 C 7207:C 7 >1<565=8< 4>A2V4>< 70AB>AC20==O 40=V9 ?>?C;OFVW.
>@CH5==O DC=:FVW =8@>:
>@5:FVO 4>78 4;O ?0FVT=BV2 7 =8@:>2>N =54>AB0B=VABN 2V4 ;53:>3> 4> ?>3> ABC?5=O (;V@5=A
:@50B8=V=C ≥ 30 <;/E2) =5 ?>B@V1=0. 5:><5=4CTBLAO 7 >15@56=VABN 70AB>A>2C20B8 ?@5?0@0B
?0FVT=B0< 7 ;V@5=A>< :@50B8=V=C < 30 <;/E2 01> ?0FVT=B0< =0 4V0;V7V.
0FVT=B8 7 @878:>< ?>@CH5==O DC=:FVW =8@>:
5O:V ?0FVT=B8, C O:8E DC=:FVO =8@>: 70;568BL 2V4 0:B82=>ABV @5=V=-0=3V>B5=78=-
0;L4>AB5@>=>2>W A8AB5<8 (=0?@8:;04 ?0FVT=B8 7 BO6:>N A5@F52>N =54>AB0B=VABN□;0A IV 70
:;0A8DV:0FVTN NYHA□ 42>1VG=8< AB5=>7>=8@:>2>W 0@B5@VW 01> AB5=>7>=8@:>2>W
0@B5@VW T48=>W =8@:8), <0NBL @878: @>728B:C >;V3>C@VW B0/01> ?@>3@5ACNG>W 07>B5
3>AB@>W =8@:>2>W =54>AB0B=>ABV =0 D>=V B5@0?VW V=3V1VB>@0<8 \$. V@>3V4=VABL 40=8E
?>4V9 28I0 C ?0FVT=BV2, O:V >4=>G0A=> ?@89<0NBL 4VC@5B8:8. 5<0T 4>AB0B=L>3>
B5@0?52B8G=>3> 4>A2V4C 287=0G5==O 0=0;>3VG=>3> @878:C @>728B:C ?>@CH5=L DC=:FVW
=8@>: C F8E 2@07;828E ?0FVT=BV2 ?@8 70AB>AC20==V 1;>;:0B>@V2 @5F5?B>@V2 0=3V>B5=78=C ,
B0:8E O: 5?@>A0@B0=. /:I> 5?@>A0@B0= ?@87=0G0NBL ?0FVT=B0< V7 ?>@CH5==O< DC=:FVW
=8@>;, =5>1EV4=> >FV=8B8 DC=:FVN =8@>: ?5@54 ?>G0B:>< ;V:C20==O V ?5@V>48G=> ?@>BO3>
< 70AB>AC20==O ?@5?0@0BC. /:I> ?@>BO3>< B5@0?VW A?>AB5@V30TBLAO ?>3V@H5==O
DC=:FVW =8@>;, 4>FV;L=VABL ;V:C20==O 5?@>A0@B0=× A;V4 ?5@53;O=CB8.
86G5707=0G5=V 70AB5@565==O A;V4 2@0E>2C20B8, 107CNG8AL =0 4>A2V4V 70AB>AC20==O
V=H8E ?@5?0@0BV2 FL>3> :;0AC B0 B0:>6 V=3V1VB>@V2 \$.
V?5@:0;VT

@>BO3><;V:C20==O;V:0@AL:8<8 70A>10<8, I> 2?;820NBL =0 @5=V=-0=3V>B5=78=-0;L4>AB5@>=>2C A8AB5A>1;82> ?@8 =002=>ABV ?>@CH5==O DC=:FVW =8@>: B0/01> A5@F52>W =54>AB0B=>ABV. 0FVT=B0< 3@C?8 @878:C @5:><5=4CTBLAO 2V4?>2V4=89 <=>VB>@8=3 :0;VN C A8@>20BFV :@>2V. >3;O4C =0 4>A2V4 70AB>AC20==O ;V:0@AL:8E 70A>1V2, I> 2?;820NBL =0 @5=V=-0=3V>B5=78=-0;L4>AB5@>=>2C A8AB5A=>G0A=5 70AB>AC20==O 5?@>A0@B0=C 7 :0;V9715@V30NG8<8 4VC@5B8:0<8, 70 V=H8<8 ;V:0@AL:8<8 70A>10<8, I> <>6CBL ?V428I8B8 @V25=L :0;VN (=0?@8::04 7 35?0@8=><), <>65 ?@872>48B8 4> ?V428I5==O :0;VN C A8@>20BFV :@>2V V B> >42V9=0 1;>:040 @5=V=-0=3V>B5=78=-0;L4>AB5@>=>2>W A8AB5<8 A=CNBL 4>:078, O:V A2V4G0BL, I> >4=>G0A=5 70AB>AC20==O V=3V1VB>@V2 \$, 1;>:0B>@V2 @5F5?B>@V2 0=3V>B5=78=C 01> 0;VA:V@5=C ?V428ICT @878: 3V?>B5=7VW, 28=8=:5==O 3V?5@:0;VT : (C B>AB@>W =8@>:2>W =54>AB0B=>ABV). ">42V9=0 1;>:040 @5=V=-0=3V>B5=78=-0;L4>AB5@>=>2>W A8AB5<8 V7 70AB>AC20==O< :><1V=0FVW V=3V1VB>@V2 \$, 1;>:0B>@V2 @5F5?B>@V2 0=3V>B5=78=C 01> 0;VA:V@5=C =5 @5:><5=4>20=0 (482. @>74V;8 «□70T<>4VO 7 V=H8 <8 ;V:0@AL:8<8 70A>10<8 B0 V=HV 2848 270T<>4V9» B0 «\$0@<0:>;>3VG=V 2;0AB82>ABV»). /:I> B5@0?VO 7 ?>42V9=>N 1;>:04>N 01A>;NB=> =5>1EV4=0, WW A;V4 70AB>A>2C20B8 ?V4 @5B5;L=8 <=03;O4><;V:0@O B0 7 G0AB8< <=>VB>@8=3>< DC=:FVW =8@>:, 5;5:B@>;VBV2 B0 0@B5@V0;L=>3> B8A:C. 4=>G0A=5 70AB>AC20==O V=3V1VB>@V2 \$ B0 1;>:0B>@V2 @5F5?B>@V2 0=3V>B5=78=C =5 @5:><5=4CTBLAO ?0FVT=B0< 7 4V015B8G=>N =5D@>?0BVTN. 5@28==89 3V?5@0;L4>AB5@>=V7< 5 @5:><5=4CTBLAO ;V:C20==O 5?@>A0@B0=× ?0FVT=BV2 7 ?5@28==8< 3V?5@0;L4>AB5@>=V7<> V?>B5=7VO !8<0B8G=0 3V?>B5=7VO <>65 28=8=:CB8 C ?0FVT=BV2 7V 7=0G=8< 45DVF8B>< @V48=8 B0/01> A>;V (=0?@8::04, 2=0A;V4>: 70AB>AC20==O 28A>:8E 4>7 A5G>3V==8E 70A>1V2). 5:><5=4>20=> 2V4=>28B8 F59 45DVF8B ?5@54 ?>G0B:<<;V:C20==O ;V:0@AL:8< 70A>1><<"525B5=®. 0E2>@N20==O :>@>=0@=8E 0@B5@V9 >A2V4 70AB>AC20==O 5?@>A0@B0=C ?0FVT=B0< V7 70E2>@N20==O< :>@>=0@=8E 0@B5@V9 >1 <565=89. !B5=>7 0>@B0;L=>3> B0 3> :;0?0=V2/3V?5@B@>DVG=0 :0@4V>?0BVO /: V I>4> 2AVE 207>48;0B0B>@V2, 5?@>A0@B0= A;V4 70AB>A>2C20B8 7 >15@56=VABN ?0FVT=B0< 7V AB5=>7>< 0>@B0;L=>3> B0 3> :;0?0=V2 01> 3V?5@B@>DVG=>N :0@4V>?0BVTN. "@0=A?;0=B0FVO =8@:8 >A2V4 70AB>AC20==O 5?@>A0@B0=C ?0FVT=B0< 7 =5I>402=L>N B@0=A?;0=B0FVTN =8@>: 2V4ACB=V9. 0AB>AC20==O C ?5@V>4 203VB=>ABV. 5 A;V4 @>7?>G8=0B8 B5@0?VN 1;>:0B>@0<8 @5F5?B>@V2 0=3V>B5=78=C ?V4 G0A 203VB=>ABV. 0FVT=B:C, O:0 ?;0=CT 203VB=VABL, A;V4 ?5@525AB8 =0 0;LB5@=0B82=5 0=B83V?5@B5=782=5 ;V:C20==O 7 2V4><8< ?@>DV;5< 157?5:8 70AB>AC20==O 203VB=82;5==V 203VB=>ABV ;V:C20==O 1;>:0B>@>< @5F5?B>@V2 0=3V>B5=78=C A;V4 =5309=> ?@8?8=8B8 B0 70 ?>B@51>N @>7?>G0B8 0;LB5@=0B82=C B5@0?VN (482. @>74V; «@>B8?>:070==O» B0 «0AB>AC20==O C ?5@V>4 203VB=>ABV 01> 3>4C20==O 3@C44N»). =HV 70AB5@565==O /: A?>AB5@V30;>AO I>4> V=3V1VB>@V2 \$, 5?@>A0@B0= B0 V=HV 1;>:0B>@8 @5F5?B>@V2 0=3V>B5=78=C >G5284=> <5=H 5D5:B82=V 4;O 7=865==O B8A:C :@>2V C ?0FVT=BV2 =53@>W4=>W @0A8 ?>@V2=O=> 7 V=H8<8 @0A0<8, <>6;82>, 2=0A;V4>: ?5@52060==O AB0=C 7 =87L:8< 2< @5=V=C C ?>?C;OFVW =53@>W4=>W @0A8 7 3V?5@B5=7VTN. 0FVT=B0< 7 @V4:VA=8<8 A?04:>28<8 70E2>@N20==O<8, B0:8<8 O: =5?5@5=>A878, =54>AB0B=VABL ;0:B078 01> <0;L01A>@1FVO 3;N:>78-30;0:B>78, =5 A;V4 ?@89<0B8 F59 ?@5?0@0B. 0AB>AC20==O C ?5@V>4 203VB=>ABV 01> 3>4C20==O 3@C44N. V:0@AL:89 70AV1 =5 A;V4 70AB>A>2C20B8 203VB=8< 01> 6V=:0<, O:V ?;0=CNBL 70203VB=VB8. ?V45;>3VG=V 40=V ?@> @878: B5@0B>35==>ABV 2=0A;V4>: 5:A?>78FVW V=3V1VB>@V2 \$?@>BO3> < ?5@H>3> B@8<5AB@C 203VB=>ABV =5 T ?5@5:>=:828<8, >4=0: =57=0G=5 ?V428I5==O @878:C =5 <>6=0 28::NG8B8. 72O7:C 7 2V4ACB=VABN :>=B@>;L>20=8E 5?V45;>3VG=8E 4>A;V465=L AB>A>2=> @878:C, ?>2O70=>3> 7 1;>:0B>@0<8 @5F5?B>@V2 0=3V>B5=78=C , =5 <>6=0 28::NG8B8 F59 @878: 4;O ?@5?0@0BV2 40=>3> :;0AC. 0FVT=B:C, O:0 ?;0=CT 203VB=VABL, A;V4 ?5@525AB8 =0 0;LB5@=0B82=5

0=B83V?5@B5=782=5 ;V:C20==O 7 2V4<8< ?@>DV;5< 157?5:8 70AB>AC20==O 203VB=82;5==V
203VB=>ABV ;V:C20==O 1;>:0B>@< @5F5?B>@V2 0=3V>B5=78=C A;V4 =5309=> ?@8?8=8B8 B0 70
?>B@51>N @>7?>G0B8 0;LB5@=0B82=C B5@0?VN.

V4><>, I> C ;N48=8 B5@0?VO 1;>:0B>@< @5F5?B>@V2 0=3V>B5=78=C ?@>BO3>< 4@C3>3> B0
B@5BL>3> B@8<5AB@C A?@8G8=OT D5B>B>:A8G=VABL (7=865==O =8@:>2>W DC=:FVW,
>;V3>3V4@0<=V>=, 70B@8A8DV:0FVW :VAB>: G5@5?0) B0 =5>=0B0;L=C B>:A8G=VABL (=8@:>20
=54>AB0B=VABL, 0@B5@V0;L=0 3V?>B5=7VO, 3V?5@:0;VTIEV4=>ABV 70AB>AC20==O 1;>:0B>@0
@5F5?B>@V2 0=3V>B5=78=C C 4@C3>
>2>=0@>465=8E, <0B5@V O:8E 70AB>A>2C20;8 1;>:0B>@8 @5F5?B>@V2 0=3V>B5=78=C , ?>B@V1=>
@5B5;L=> >1AB568B8 I>4> @>728B:C 0@B5@V0;L=>W 3V?>B5=7VW (482. @>74V;8
«@>B8?>:070==O» B0 «A>1;82>ABV 70AB>AC20==O»).

"525B5=® =5 @5:><5=4>20=89 4;O 70AB>AC20==O C ?5@V>4 3>4C20==O 3@C44N, B> 2V4ACB=V
2V4?>2V4=V 40=V. 5@5203C A;V4 =040B8 0;LB5@=0B82=8< <5B>40< ;V:C20==O 7V 2AB0=>2;5=8<
?@>DV;5< 157?5:8 70AB>AC20==O ?@>BO3>< ?5@V>4C 3>4C20==O 3@C44N, >A>1;82> ?@8
3>4C20==V =>2>=0@>465=8E 01> =54>=>H5=8E 4VB59.

**40B=VABL 2?;820B8 =0 H284:VABL @50:FVW ?@8 :5@C20==V 02B>B@0=A?>@B< 01> V=H8<8 <5E0=V7
<0<8.**

?;82 5?@>A0@B0=C =0 740B=VABL :5@C20B8 02B>B@0=A?>@B< V ?@0FN20B8 7 V=H8<8 <5E0=V7<0
<8 =5 4>A;V46C202AO, 0;5 7 >3;O4C =0 D0@<0:>48=0ABV ?@5?0@0BC <0;>9<>2V@=>, I> 2V=
=530B82=> 2?;820T =0 FN 740B=VABL. 4=0: A;V4 4>B@8I5@56=>ABV, >A:V;L:8 V=>4V ?@8 ;V:C20==V
0@B5@V0;L=>W 3V?5@B5=7VW <>6CBL 28=8:0B8 2V4GCBBO AB<;5=>ABV V 70?0<>@>G5==O.

!/?>AV1 70AB>AC20==O B0 4>78.

5:><5=4>20=0 4>70 4;O 4>@>A;8E AB0=>28BL 600 <3 5?@>A0@B0=C 1 @07 =0 45=L (2@0=FV).
1V;LH>ABV E2>@8E <0:A8<0;L=5 7=865==O 0@B5@V0;L=>3> B8A:C 4>AO30TBLAO G5@57 2-3
B86=V ;V:C20==O.

"525B5=® <>6=0 70AB>A>2C20B8 O: <>=>B5@0?VN 01> C :><1V=0FVW 7 V=H8<8
0=B83V?5@B5=782=8<8 70A>10<8. >:@5<0, 4>4020==O BV0784=8E 4VC@5B8:V2, B0:8E O:
3V4@>E;>@>BV0784, 01> 1;>:0B>@V2 :0;LFVT28E :0=0;V2, B0:8E O: =VD548?V= ?>4>265=>W 4VW,
?@>45<>=AB@C20;> 048B82=89 5D5:B ?@8 70AB>AC20==V 7 5?@>A0@B0=>
@5?0@0B <>6=0 ?@89<0B8 =570;56=> 2V4 ?@89>

0FVT=B8 ;VB=L>3> 2V:C

A>1;82>3> ?V41>@C 4>78 4;O ?0FVT=BV2 ;VB=L>3> 2V:C =5 ?>B@V1=>.

0FVT=B8 7 ?>@CH5==O< DC=:FVW ?5GV=:8

>A2V4 70AB>AC20==O ?0FVT=B0< 7 ?5GV=:>2>N =54>AB0B=VABN >1<565=89 (482. @>74V;

«A>1;82>ABV 70AB>AC20==O»).

0FVT=B8 7 ?>@CH5==O< DC=:FVW =8@>:

;O ?0FVT=BV2 V7 ?>

VB8. A:V;L:8 157?5:0 B0 5D5:B82=VABL 70AB>AC20==O ;V:0@AL:>3> 70A>1C "525B5=® 4;O 4VB59 =5
2AB0=>2;5=V, ?@87=0G5==O ?@5?0@0BC ?0FVT=B0< 40=>W :0B53>@VW =5 @5:><5=4CTBLAO.

5@54>7C20==O.

0=V I>4> ?5@54>7C20==O ;N4O< T >1<565=8<8. # ?>AB<0@:5B8=3>289 ?5@V>4 >45@60=V >:@52V4>
<;5==O ?@> 70AB>AC20==O 2=CB@VH=L> 4>7 4> 12 000 <3. V;LHVABL ?0FVT=BV2 ?>2V4><;O;8 ?@>
2V4ACB=VABL A8=>3> ?0FVT=B0 A?>AB5@V302AO F8@:C;OB>@=89 :>;0?A ?VA;O ?@89>
A0@B@0=C 2 4>7V 12 000 <3. 0FVT=B ?>2=VABN >4C602. 09V<>2V@=VH8< ?@>O2<< ?5@54>7C20==O
<>65 1CB8 0@B5@V0;L=0 3V?>B5=7VO. # @07V 28=8=:5==O A8<0B8G=>W 0@B5@V0;L=>W
3V?>B5=7VW =5>1EV4=> ?@87=0G8B8 2V4?>2V4=C B5@0?VN.

>1VG=V @50:FVW.

?0FVT=BV2, O:V >B@8A0@B0=><, =091V;LH G0AB> ?>2V4<;O;>AO ?@> B0:V ?>1VG=V @50:FVW,
O: 3>;>2=89 1V;L B0 =5A?5F8DVG=V ?>@CH5==O 7 1>:C H;C=:>2>:8H:>2>3> B@0:BC, I>
A?>AB5@V30;8AO C ?@81;87=> 11 % V 8 % ?0FVT=BV2 2V4?>2V4=>.

>1VG=V @50:FVW 2:070=V 70 G0AB>B>N: 4C65 G0ABV (□1/10), G0ABV (□1/100, <1/10), =5G0ABV (□1/1000, <1/100), @V4:VA=V (□1/10000, <1/1000), 4C65 @V4:VA=V (<1/10000), 2;;NG0NG8 ?>>48=>:V 28?04:8.

>7:048 7 1>:C VW A8AB5<8:

=5G0ABV: 3V?5@GCB;82VABL*.

>7:048 7 1>:C =5@2>2>W A8AB5<8:

4C65 G0ABV: 3>;>2=89 1V;L*;

G0ABV: 70?0<>@>G5==O*.

!C48==V @>7:048:

=5G0ABV: 0@B5@V0;L=0 3V?>B5=7VO.

5A?V@0B>@=V, B>@0:0;L=V B0<54V0AB8=0;L=V @>7:048:

G0ABV: @8=VB.

>7:048 7 1>:C H:V@8 V ?V4H:V@=8E B:0=8=:

G0ABV: 0;5@3VG=V H:V@=V @50:FVW (2;;NG0NG8 28A8?0==O, A25@1V6);

=5G0ABV: 0=3V>=52@>B8G=89 =01@O*.

>7:048 7 1>:C H:C=:>2>-.8H:>2>3> B@0:BC:

G0ABV: =5A?5F8DVG=V H;C=:>2>-.8H:>2V @>7:048: =C4>B0, 1;N20==O, 4V0@5O.

030;L=V ?>@CH5==O B0 @50:FVW C

G0ABV: 0AB5=VO.

*0AB>B0 =5 28I0, =V6 ?@8 70AB>AC20==V ?;0F51>.

@V< ?>1VG=8E @50:FV9, 28O2;5=8E ?V4 G0A ;;V=VG=8E 4>A;V465=L, ?@8 ?>AB<0@:5B8=3>2> AC20==V 5?@>A0@B0=C A?>=B0==> ?>2V4<;O;>AO ?@> =86G5707=0G5=V ?>1VG=V @50:FVW, G0AB>BC O:8E =5<>6;82> >FV=8B8 70 =0O2=8<8 40=8<8 (=52V4><0).

>7:048 7 1>:C =8@>: V A5G>284V;L=>W A8AB5<8:

?>@CH5==O DC=:FVW =8@>;, C B>20 =54>AB0B=VABL, C ?0FVT=BV2 3@C?8 @878:C (=0?@8;;04, 7V AB5=>7>< =8@>:2>W 0@B5@VW).

"5@ABV.3 @>:8.

#<>28 715@V30==O. 15@V30B8 2 >@83V=0;L=V9 C?0:>2FV ?@8 B5ABC?=>

#?0:>2:0.> 14 B01;5B>: C 1;VAB5@V, ?> 1 01> 2 1;VAB5@8 C :0@B>==V9 :>@>1FV.

0B53>@VO 2V4?CA:C. 0 @5F5?B><.

8@>1=8:. 11>BB %5;A:50 !!, \$@0=FVO/Abbott Healthcare SAS, France.

VAF57=0E>465==O 28@>1=8:0 B0 9>3> 04@5A020465==O 4VO;L=>ABV. CB 45 5;L2V;L, L9> 4V 5;0@, 01400, (0B89>--AN@-(0;0@>=, \$@0=FVO/Route de Belleville, Lieu dit Maillard, 01400, Chatillon-sur-Chalaronne, France.