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

" ®
(ADVANTAN®)

!:;04:
4VNGV @5G>28=8: 1 3 <07V   <5B8;?@54=V7>;>=C 0F5?>=0BC;
4>?>28=8: 2VA: 1V;89, >;VODehymuls E), 2>40 >G8I5=0.

V:0@AL:0 D>@<0. 07L.
A=>2=V DV78:>-EVABV: =5?@>7>@0 <07L 1V;>;3> 01> 7 6>2BC20B8< 2V4BV=:>< :>;L>@C.

$0@<0:>B5@0?52B8G=0 3@C?0. >@B8:>AB5@>W48 4;O 70AB>AC20==O C 45@<0B>;>3VW.
>4 "% D07A C14.

$0@<0:>;>3VG=V 2;0AB82>ABV.
$0@<0:><0:>>48=0
420=B0=® ?@8 AC20==V ?@83=VGCT 70?0;L=V B0 0;5@3VG=V @50:FVW H:V@8, 0 B0:>6 @50:FVW,
?>>2O70=V 7 3V?5@?@>;VD5@0FVTN :;VB8=, I> A?@8@8OT CAC=5==N O: >1T:B82=8E A8
V4<><, I> <5B8;?@54=V7>;>=C 0F5?>=0B 157?>A5@54=L> 72O7CTBLAO 7 2=CB@VH=L>:;VB8==8<8
3;N:>:>@B8:>W4=8 @5F5?B>@0<8. &5 >A>1;82> AB>ACTBLAO >A>2=>2=>3> <5B01>;VBC   6α-
<5B8;?@54=V7>;>=C-17-?@>?V>=0BC, O:89 CB2>@NTBLAO ?VA;O @>7I5?;5==O C H:V@V.
2O7C20==O :>@-AB5@>W4 V7 ?52>>N 4V;O=:>N <>;5:C;8 V=VFVNT =87:C 1V;>>3VG=8E 5D5:BV2.
2O7C20==O :>@-AB5@>W4 ?@8872>48BL 4> V=4C:FVW A8=B57C <0:>>>>>3@C?8 01>>0EV4=8<8
V=3V1CT 282V;L=>>OV 0@0EV4>>>2>W :A;>B8 V B0:8< G8=>< 7=86CT CB2>@5==O <54V0B>@V2
70?0;5==O, B0:8E O: ?@>AB03;0=48=8 B0 ;59:>B@VT=8.
AC?@5A82=C 4VN 3V?5@B@>W48 H:V@8 @5O;V7CTBLAO H;OE>< V=3V1>2V=8 V=4C:FVW A8=B57C F8B>:V=V2
B0 0=B8B8G=8E 5D5:BV2, >:89 ?>:8 I> =54>AB0B=L> 282G5=89.
=3V1C20==O A8=B57C 207>48;0>FV9=8E ?@>AB03;0=48=V2 01> ?>B5=FVN20==O
207>:>AB@B8B>V=>=>>3> 5D5:BC 04@5=0;V=C 2@5HBV @5HB 7C<>2;NT 207>:>AB@8:B>@=8C
0:B8V20=VABL 3V?5@B@>W4V2.
$0@<0:>:V=5B8:0.
5B8;?@54=V7>;>=C 0F5?>=0B 3V4@>;V7CTBLAO 2 5?V45@<VAV B0 A>AB>=>>>>   6a-
<5B8;?@54=V7>;>=C-17-?@>?V>=0BC, 9>3> >A=>2=>3> <5B01>;VBC, O:89 A8;L=VH5 72O7CTBLAO 7
@5F5?B>@0<8, =V6 ?>G0B:>20 A?>;C:0, =V6 ?@>0B:>289 ?@>4C:B, GVB:0 >7=0G0 H:V@>N
«1V>0:B820FVO».
V4A>B>:25 B0 7030;L=5 2V4@>H5==O G5@57H:V@>N 01A>@1FVN 3> :>@B8:>AB5@>W4C
70;568BL 2V4 =87:8 D0:B>@V2, B0:8E O: EV;:C8, 4>?>28=, :>=F5=B@0FVO A?>;C:8 2 >A=>2V, C>>28
4VW (?;>I0 >1@>1;5==O >?>2@EW, ?>250@=V, B@@820;VABL 4VW, 2V4:@8B89 01> >:;N782=89 A?>>AV1
;V:C20==O) B0 E0>0:B5@8AB8:8 H:V@8 (284 B0 ABC?V=L BO6:>ABV >B>?@@>;>>3VW, 0=B>0>:0;V70FVW).
'5@57H:V@=8 01A>@1FVN <5B8;?@54=V7>;>=C 0F5?>=0BC C D>@@ ?V428I5=>. # 4@@>A;8E,
E2>@8E =0 ?A>@V0B, B0 C ?0FV@=T 7 0B>?VT=BN 0B>>>>?VTN G5@57H:V@=C 01A>@1FVO <5B8;?@54=V7>;>=C
0F5?>=0BC C D>@@   2,5 %.
VA;O ?>B@0?;O==O 4> :@>2>B>:C >A=>2=89 ?@@>4C:B 3V4@>;V7C <5B8;?@54=V7>;>=C 0F5?>=0BC,
6a-<5B8;?@54=V7>;>=C-17-?@>?V>=0B, H284> 72O7CTBLAO 7 3;N:C@>=>2>N :A;>B>N V B0:8< G8=>A>
< V=0:B82CTBLAO.
5B01>;VB8 <5B8;?@54=V7>;>=C 0F5?>=0BC (>A=>2=89 <5B01>V: 6a-<5B8;?@54=V7>;>=C-17-
?@>?V>=0B-21-3;N:>@>=V4) 282>4OBLAO 7 >@30=V7<C ?5@52065=> G5@57 V7 A5G5N, ?5@V>4
=0?V2282545==O AB0=>28BL ?@81;87=> 16 3>48=. VA;O 2=CB@VH=L>25==>3> 70AB>AC20==O
?@@>2 5 282545==O G5@57 A5GC B0 28?@@>6=5==O 2V41C20BLAO ?@@>B034>< 7 4V2. :C30=V7W
@5G>28=8 01> <5B01>;VB82 =5 A?>AB5@V30TBLAO.

;V=VG=V E0@0:B5@8AB8:8.
>:070==O.

B>?VG=89 45@<0B8B (5=4>35==0 5:75<0, =59@>45@=B0:B=0 5:75<0, 48A3V4@>B8G=0,
=5A?5F8DVG=0 5:75<0, 5:75<0 C 4VB59.

@>B8?>:070==O.
V428I5=0 GCB;82VABL 4><5B8;?@54=V7>;>=C 0F5?>=0BC 01> 4> 1C4L-O:>3> V=H>3> :>=5=B0
?@5?0@0BC; BC15@:C:C;L>7=V B0 A8DV;VB8G=V ?@>F5A8 C 4V;;O=FV =0=5A5==O<07V; 2V@@CA=V
70E2>@@N20==O (=0?@>8.;;:04, 2VB@@@0=0 2VA?@0, >?5@V7VC20;L=8H09 ;8H09), @>70F50, =02:>;;:>@@>B>289
45@@<0B8B, 28@@07:8, 0:=5 728G09=5, 0B@0>DVG=89 45@@<0B8B 01> @50:FVW ?VA;O I5?;;5==O C
4V;;O=FV =0=5A5==O <07V.

70T<>4VO 7 V=H8<8 ;V:0@ALL:8<8 70A>10<8 B0 V=HV 2848 270T<>4V9.
5<0T 40=8E.

A>1;82>4T>ABV 70AB>AC20==O.
;N:>:>@@@B8:>W48 A;V4 70AB>A>2C20B8 BV;L:8 C 3> B5@@@0?52B8G=>3> 5D5:BC.
@07V >1@>1;;5==O 25;;8:8E 4V;;O=>: H:V@@@8 B@@@820;;VABL ;V:C20==O ?>28==0 1CB8 28>7>G5=0
2V4?>2V4=8< G8=><
@8 10:B5@@V0;;;:8E V=D5:FV9=8E ?@>F5A0E H:V@@@8 V/01> C @07V 3@@@81:>2>3> C@06==O
=5>1EV4=> 4>40B:>25 A?5FVW;;:0 ;V:C20==O.
VAF52V H:V@@=V V=D5:FVW<>6CBL 703>AB@@N20B8AO C @07V 3> 70AB>AC20==O
3;N:>:>@@@B8:>W4V2.
5>1EV4=> C28=0B8 70AB>AC20==O ;V:0@@:>3> 70A>1C 2 7=>=CS ?@8 @>70F50 01>
=02:>@@>B>289
@8 70AB>AC20==V 420=B0>= A;V4 C28=0B8 ?>B@@@0?;;@0==O ?@0590@@0B2 2 >GV, =0 3;;810:>V 2V4:@@@8BV
@08, =0 A;87>2V >1>;>=:8.
/:I> H:V@@0 =04 ?5@@5A8E0T ?@8 B@@@820;;;:>@®, @5:>><5=4CTBLAO ?5@@59B8 =0 70AB>AC20==O V=H>W
;V:0@@:>W D>@<8 ?@5?0@@0BC 7 28I8< 2< 68@@@2 (68@@@0 <07L 420=B0>=®).
@8 =0AB>AC20==V ?@5?0@@0BC 420=B0>=® =0 25;;8:C ?>25@@E=N H:V@@8 (40-60 % ?>25;@E=V H:V@@8) 01>
=02VBL ?@8 28:>@@8AB0==V =5>;;N7V9=8E ?>2O7:8 =5 C 4>@>A;8E, =5 C 4VB59 =5
A?>@@>1;;V2VO;;;;>AO ?>@@@@CH5==O DC::FVW :>@@8 =04=8@@:>>28E 70>;7. V4 G0A 70AB>AC20==O
<5B8;?@54=V7>;>=C 0F5?>=0BC =0 60 % ?>25;@5=V BV;;0 ?V4 :;:N782==C ?>2O7:C ?@>BO3><   22
3>48= C 74>@@>28E 4>@>A;8E ;N459 A?>@@>1;;V2VO;;;;>AO 7=865==O @V2=O >:>B87>7V B0
2:;82 =0 9>3> 4>1>289 @8B
@8 =0AB>AC20==V :>@@@80B5;L=>W 4;O 3> 70AB>AC20==O C 28A>:8E 4>70E =04 25;;8:C ?;;>IV BV;0
01> C?@0>425 B@@@820;>3> ?5@@V>4C :5;;5@N7V9=V ?>2O7:8, 7=0G=> ?V428I5CTBLAO
@8878: 28=8:=5==O ?>1VG=8E 5D5:BV2. 5>1EV4=> C28=0B8 ;V:C20==O V7 28:>@@8AB0==O<
>;;N7V9=>W ?>2O7:8, O:I> 1>;5 =5 ?@@870G5==O. !;V4 ?0<O70B8, I> ?5;NH8 B0 ?V43:7=:8:8, B0: A0<>,
O: 7>=8 ?>?@0V;;;;>2O7:8,<>6CBL A0<@8G=82=>0 ?>1VG=>.
@07V ;V:C20==O 25;;8:8E 4V;;O=>: ?>25;@5=V BV;;0 B@@@820;>3V ;V:C20==O ?>28==0 1CB8 O:><>30
<5=H>N, >A:V;;:8 =5<>6;82> ?>20=VABN 28::NG8B8 2V@@@0=0 420=B0>=B01A0 @08A0AB5<=8H
5D5:BV2.
"0: A0<>, O: 7 CAV<0 :>@@@@8:>W40<8, =52V4@?>2V4=5 70AB>AC20==O FL>3> ;V:0@@:>@@>3>
70A>1C <>65 <0;@:0=B201B8 ;:V:>=VG=C :0@@B8=C A8<?0<.
@8 3;;0C:><V WW =0O2=>AB>ABV 2 0=0<=57V A;V4 ?>2O7:0 H:V@@8 ;V:0@@:>W.
/: 2V4<><>=, ?@8 A0<@0B5L<V B5;@0>?VW :>@@@@8:>W40<8 B0: A0<> 8 ?@8 7028E>@@N20==>@?H07V
(=0?@>8.;;:04, ?VA;O 70AB>AC20==O C 28A>:8E 4>70E 01> =0 25;;8:V9 ?>25;@5=V ?@>BO3>@><
?5@@V>4C, 28>6:N0@@0==O >;;;N7V9=>8E ?>2O7:8 01> =0=5A5==O =0 H:V@@C =00 >1;8G8V ?>B@@@0)
.
0AB>AC20==O, >A;V4 4>23>B@@@820;>3>, 70A1@V2O 4;O 3> ;V:C20==O, <>65 A?@@0V>7C20B8 @>72V@=V
A5A=81V;@V70FVW. # B>;;;V:C20==O ?@@8?8=8T BL>3> 70AB>AC20==O B0 ?@@870G5=8T 2V4?>2V4=N B5;@0?VN.

0AB>AC20==O C ?5@@V>4 203VB>>ABV 01> 3@>4C20==O 3@C44N.
03VB=VABL.

0 40==89 G0A =5<0T 4>AB>2V@=8E 40=8E I>4> 70AB>AC20==O ?@5?0@0BC 420=B0=B0=B0® 203VB=8<
6V=:0

:A?5@8<<5=B0;L=V 4>A;V465==O =0 B20@@8=0@0 V7 70AB>AC20==O< <5B8;;?@54=V7>;=0 0F5?>=0BC V7 70AB>AC20==O< 4>7, O:V ?5@8528IC20;8 B5@0?52B8G=C 4>7C, 28O28;8 5<1@V>B>:A8G=89 B0 /01> B5@0?V4>A>@0?32835==89 5D5:B.

0728G09 C B@8<5<5AB2;O 203VB@8=>ABV A;V4 C=8:0B8 7@2=8>VH=L>3> 70AB>AC20==O ?@5?0@0BB2, I>@@8:>>AB5@@3>W48 (7>:@5<0, ?@5?0@0BC 420=B0=B0®). 5O:V 5?V45;>3VG=V 4>A;V465==O 4>72>;O;;ONBL ?@8?CA:0:0B8 <>6;82V;LH5==O @878:C ?0;>B@AE87C C =>7>0@>@465=8E, O:I> WE <0B5@V 70AB>A>2C20;8 A8AB5<<V <>6;82>;V4@8:>>AB5@@3>W48 C B@8<5AB2@V 203VB@8=>ABV.

?5@V>4 203VB@8=>ABV 01> 3>4C20==O 3@C44N =5>1EV4V4=> C=8:0B8 A8>=0=0 ?=:1@5==O 25;8:8E 4V;O5>>: ?>25@E2V H:V@8, 4>23@8202;>3> ;V:C20==O 01> 28:>@8AB0==O >:;N7V9=8E ?>2O7>:.
5>1EV4V4=> 2@0E>2C20B8 A?V;L22V4>H5==O @878:->:>8>ABL ?@8 ?@87>V35==V ?@5?0@0BC 420=B0=B0®
203VB=8

>4C20==O 3@C44N.

>A;V465==O =0 B20@@8=0@0 28O28;8, I> <5B8;;?@54=V7>;>=0 0F5?>=0BC V7 70AB>AC2 25?@540@5O40TBLAO
=>2>;L4@8:C >7V< G5@57 3@C4=5 <>;>:>. "8< =5 <5=H5, =5 28@02;5=>, G8 ?@>>=8:0B
<5B8;;?@54=V7>;=0 0F5?>=0BC V7 70AB>AC20==O< C 3@C4=5 <>;>:> ;N48=8, I>
70AB>A>2C2CNBL40O A8AB5<<>, 28?2;ONBL40O C 3@C4=5 70AB>AC20==O ?@5?0@0BC 420=B0=B0®
?@8@8725AB8 4> 4>@8;>L=>3>W A8AB5<<>W 01A>@2F?VW <5B8;;?@54=V7>;=0 0F5?>=0BC, O:0
4>7V :8;V 1 28>28B8 9>3> C 3@C4=5 <>;>:>

7 4>@8;>L=>3> ?@8A8V=8 =5>1EV4V4=> 28AB8 C8B8<>:C @5?@5@0@0BC @?@87>V35==O 420=B0=B0® 6V=:0<, O:V 3>4C4NBL
3@C44N.

# ?5@V>4 3>4C20==O 3@C44N =5 A;V4 =0>=>><A8B8 <07L 420=B0=B0® =0 <>:@V;V4 70;>3>V20 70;0@0. A>1;82>> A;V4
C=8:0B8 B@8<5<8AB2;V O: >1>A>8AB2 70AB>AC20==O ?@5?0@0BC, =>=0:5A5==O 9>3> =0 25;8:V 4V;O5>V 4V; H:V@8 01>
28:>@8AB0==O >:;N7V9=8E ?>2O7>:.
$5@8>;L=VABL. 0=8:> I>4> 2?;82C ?@5?0@0BC =0 D5@8;V=VABL =5<0T.

***40B=VABL 2?;82020B8 =0 H284:VABL @50:FVW ?@8 :5@C20==V 02B>B@0=A?>@B>< 01> V=H8<8 <5E0=V7<0<8
<5E0=V7<0<8.***
5 2AB0=>2;5=0.

!?>AV1 70AB>AC20==O B0 4>78.
0728G09 <07L =0=>A8B8 =0 C@065=V 4V;O5>:> H:V@8, A>@065=V 4V;O5>:> 3@8 4>1V 1 @07 =0 4>1C B0 >>:@>8202;>3> 70AB>AC20==O 0728G09
  =5 <5=H5 ?5@V528IC20==O 4;O 4>A?O35==O 12 B86=V2, 4;O 4>22HV=9 4 B86=V2.
@07V ?>>?CA:C 28:>@8AB0==O ?@5?0@0BC =5>1EV4V4=> =0=>A8B8 =0 =0ABC?=V 4>7V, =5 =0=>AOG8 2?>4>V9=V 4V;O5>>: 4>7C ?@5?0@0BC =0 =0ABC?=89 4>7C.
7@8<=5 B86=V2 4 A;V4 28:>@8AB0B8 ?@5?0@0B <5=H5 4V;O5V H:V@8 B0 A;V4 @7E2>@8282;L4>>: 4>>A0>>6O =>@8@725AB8 4> A?;;L4@8:5@865=89 5D5:B2.
28?>2V>=> 28:>@8AB0B8 ?@5?0@0B (70AB>AC20==O =0 25;8:V 4V;O5V H:V@8 25?@540@5O40TBLAO 2V4 25;8:8E 4V;O5>>:,
I> A?@8O4NONBL 01A>@2FVW) 01> ?@8A8>V35==O =0 B@82>V> 3@8 4>@8:>>AB5W@3>W48) 01> ?@8@8725AB8 4> =5>1EV4V4=>ABV 28AB8>8B8.

>1VG=V @50:FVW.

'0AB>BC ?>1VG=8E 5D5:BV2, I> A?>AB5@V30NBLAO C E>4V :;V=VG=8E 4>A;V465=L V =02545=V 2 B01;8FV =86G5, 287=0G5=> 2V4?>2V4=> 4> :>=25=FVW G0AB>B8 ?>1VG=8E 5D5:BV2 MedDRA: 4C65 G0AB> (>1/10); G0AB> (>1/100, <1/10); =5G0AB> (>1/1000,<1/100), @V4:> (>1/10000, <1/1000); 4C65 @V4:> (<1/10000), =52V4><> (G0AB>B0 =5 <>65 1CB8 >FV=5=0 =0 >A=>2V =0O2=8E 40=8E; <>6;82V ?>1VG=V G=V @50:FVW, I> =5 A?>AB5@V30;8AO ?V4 G0A :;V=VG=8E 4>A;V465=L). ;O :>4C20==O 1C;> 28:>@8AB0=> 25@AVN 12.0 MedDRA.

@30=8 V A8AB5<8<8	'0AB>	5G0AB>	52V4><>
030;L=V @>7;048 B0 @50:FVW C	5GV==O, A25@1V6 C	!CEVABL, 5@8B5<0, 2578:C;8, ?>4@07=5==O, 5;75<0, ?0?C;8 2	$>;V:C;VB, 3V?5@B@@8E>7, 28A8?0==O, ?CABC;8, 1V;L, ?0@5AB57VO C
>@CH5==O 7 1>:C H:V@8 B0 ?V4H:V@=>W :;VB:>28=8		B@>DVO H:V@8, 5EV<>7, V, =04 68@@=0 H:V@0	:=5, B5;50=3V5:B07VO, ?>O20 AB@VW, =02:>;> B@>289 45@<0B8B, H:V@V=V ?@>O28 B0 VI8=8, 7;O@@C C H:V@8, 10:B5@V0;L=89 F5;N;VB, :>=B0:B=89 45@<0B8B, ?V>45@<0 45@2V C@065==O, 7=510@2;5==O H:V@8, H:V@=0 @50:FVO =0 A>=F5 @3VG=89 :>=B0:B=89 45@<0B8B, 45@<0B8B @>7;048 B0 @50:FVW
>@CH5==O 7 1>:C H:V@8 VW A8AB5<8<8			V?5@3V4@>7, 82VABL 4> ;V:0@L:>3> 70A>1C C

>:@5<5=8 28?04:8 ?@8 B0 B@820;>AB>AVO ==V <>6;8289 A8AB5<=5 89 2?;82 :>@B8:>AB5@>WV4V2 C 72O7:C 7 WE 01A>@@1FVTN.

"5@A@ABV. 3 @>:V2:8.

#<>28 715@V30==O.
15@V30B8 ?@8 B5<@5@0BC@V =5 2814 40>ABC?=89 4;O 4VB5289

#?0:>2:0. > 15 3 C BC1V, ?> 1 BC1V 2 :0@B>==V9 :>@>1FV.

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

8@>1=8:.
09T@ %5;A:5@ 0=CD0:BC@V=3 !. .. .., B0;VO/Bayer Healthcare Manufacturing S.R.L., Italy.

VAF57>0E>465==O 28@>1=8:0 B0 9>3> 04@5A0 20465==O 4VO;L=>ABV.
V0 (5@V=3 21, 20090, !53@0B5, V;0=, B0;VO/
Via E Schering, 21, 20090, Segrate, Milano, Italy.