

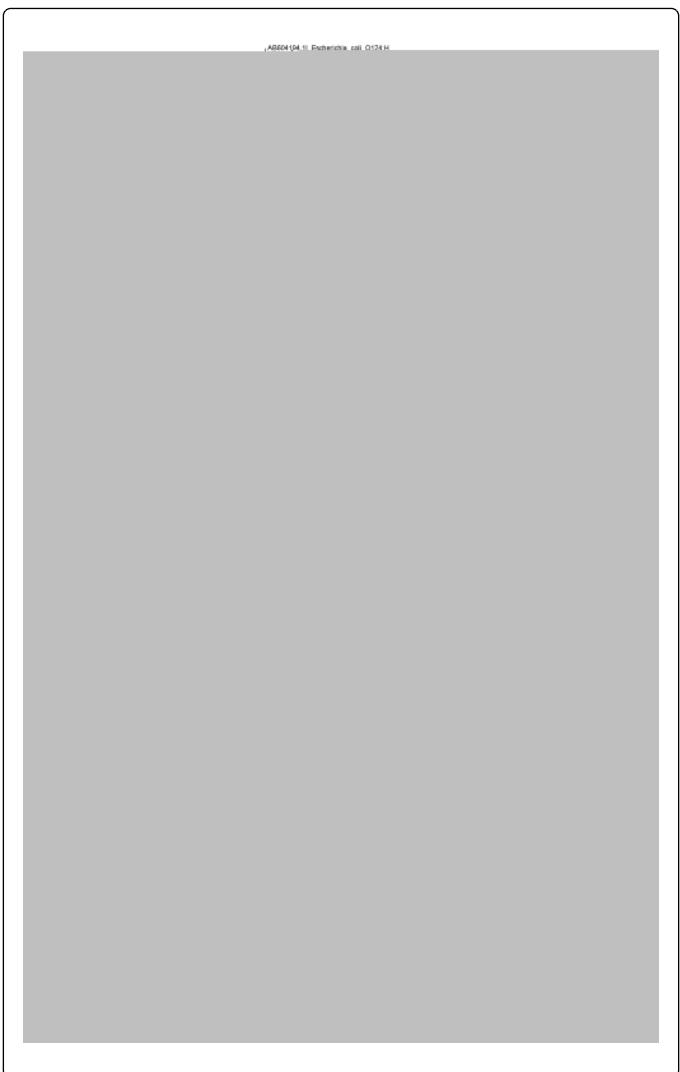


:KHUH,QLWLDO &RQFHQWUDWLRQ EHIRUH 7UHDWPHQW =HUR 7LPH
5& 5HVLGXDO &RQFHQWUDWLRQ DIWHU 7UHDWPHQW DW HDFK ([SRVXUH
7LPH

5HVXOWV

,VRODWLRQ DQG FKDUDFWHUL]DWLRQ RI LQGLJHQ RXV EDFWHULD
&RORQ\ PRUSKRORJ\ DQG WKH PLFURVFRSLF FHOO H[DPLQDWLRQ DV ZHOO DV
*UDP VWDLQ UHDFWLRQ RI WKH SXULILHG EDFWHULD GDWD QRW VKRZQ UHYHDO
WKDW WZR RI WKH LVRODWHG LQGLJHQ RXV EDFWHULD ZHUH *UDP SRVLWLYH UR
5] DQG 5] DQG WKH UHVW ZHUH *UDP QHJDWLYH URGV \$FFRUGLQJ WR WKH
\$3, ELRFKHPLFDO SURILOHV 7DEOH LQGLJHQ RXV EDFWHULD ZHUH PRVW
SUREDEO\ DIILOLDWHG W\RVH\YCHR P\RSQH\FL H\H\ S\QF\DX GLQJ
5] 5] DQG (5W\HUREDFWHU 5QJJ\J\H\RP\W\HDOQD
SQHXPRLD 6DOPRQHO 5QJ D\QJ\PRQHO QHULV
5]

Bacterial Code	API Identification
	(Q W H U R E D F W H U A I H F A R P H U D Q V
Ü:F	(FOR D A H D E A
	3 V H G X R P R Q D V A M E L A D F L D
Ü:G	3 P D O W A M S K L O L D
Ü:H	6DOPRQHOOD DUL]RQDH
Ü:I	. O H E V S I Q Q O D R V S I S D J A



)LJXUMHLJKERU -RLQLQJ 7UHH %DVH G RQ 6 U51\$ 6HTXHQFHV RI WKH ,QGLJHQRXV DQG ([RJHQRXV %DFWHULD (YROXWLRQDU\ GLVWDQFHV ZHUH FDOFXODWHG XVLQJ WKH .LPXUD PRGHO XVLQJ 0(*\$ VRIWZDUH 7KH QXPHUDOV VKRZ WKH UHVXOWV RI WKH ERRWWUDS DQDO\VLV YDOXHV IURP UHSOLFDWHV RQO\ ERRWWUDS YDOXHV DERYH ZHUH VKRZQ 7KH VHTXHQFH LQ EROG ZDV GHWHUPLQHG LQ WKLV ZRUN

'LVVROYHG R[\JHQ '2

'2 OHYHOV RI WKH VHHGHG DQG XQ VHHGHG ZDVWHZDWHU UHFRUGHG QRWLHFDEOH YDULDWLRQV ZLWK WLPH 'DWD QRW VKRZQ \$ UHPDUNDEOH GHFUHDVH LQ WKH UDZ ZDVWHZDWHU '2 PJ O ZDV UHFRUGHG ZLWK LQFUHDVLQJ H[SRVXUH WLPH FRPSDUHG WR WKH FRQWURO XQ VHHGHG ZDVWHZDWHU 7KLV LV PDLQO\ GXH WR LQFUHDVH '2 FRQVXPSSLRQ GXULQJ R[LGDWLRQ RI WKH LQFOXGHG RUJDQLF PDWWHU ZLWK WKH DGGLWLRQ RI EDFWH \$PRQJ WKH WHVWHG VSHFLHV WKH PL[HG FXOWXUH H[KLELWHG WKH KLJKHVW ELRGHJUDGDWLRQ DFWLYLW\ VKRZQ DV VKDUS GHFUHDVH LQ WKH '2 UHDFKLQJ

&KHPLFDO R[\JHQ GHPDQG &2' DXJPHQWHG EDFWHULD RQ WKH SDWKRJH
ZDVWHZDWHU
5DZ ZDVWHZDWHU UHFRUGHG UHODWLHYHO\ KLJK &2' OHYHO PJ O DW
WKH JHUR WLPH ZKLFK LV SURSRUWLRCDDQO\ DG/HFHUZDWHG ZKRVZH QV LKPLHJ KL OF RDQOQW
ZDVWHZDWHU VDPSOHV VHHGHG DQG &Q\ WPO U\ A\ Q\ K\ L\ Q\ J\ &W\ K\ P\ Q\ O\ R\ S\ M\ V\ R\ H\ Q\ U\ M\ H\ G\ G\ E\ D\ F\ O\ M\ F\ R\ Q\ F\ H\ Q\ W\ U\ D\ W\ L\ R\ Q\ 5& DW WKH ODVW H\ Q\ U\ R\ D\ X\ N\ P\ E\ O\ M\ D\ E\ L\ J\ Q\ U\ M\ \ P\ R\ U\ 7&HDQ\ G\ U\ H\ V\ M\ P\ &2' UHPRYDOV E\ DOO WUHDWPHQWV ZHUKH\ DPEKHA YFHGO\ W\ X\ W\ K\ L\ Q\ K\ W\ K\ E\ H\ W\ H\ G\ W\ K\ H\ K\ K\ R\ J\ K\ H\ [SRVXUH WLPH 6LJQLILFDQW YDULDW\ L\ R\ Q\ V\ LQ WKH &2' DQ\ P\ R\ 5Y\ M\ D\ G\ D\ Z\ H\ R\ U\ B\ H\ M\ S\ M\ R\ K\ UHFRUGHG DPRQJ WKH WHVWHG EDFWHULD Z\ D\ W\ K\ 3\ S\ D\ W\ R\ N\ H\ U\ L\ G\ H\ U\ D\ H\ V\ G\ U\ M\ K\ R\ U\ P\ R\ H\ G\ V\ I\ D\ F\ W\ 5(IROORZHG E\ WKH 5] P\ Z\ H\ W\ K\ F\ D\ X\ O\ P\ R\ X\ V\ H\ U\ H\ P\ R\ Y\ D\ O\ U\ H\ F\ R\ U\ G\ H\ G\ E\ 5] DQG ILQDOO\ WKH FRQWU\ K\ H\ O\ O\ Z\ L\ W\ K\ G\ D\ K\ H\ R\ I\ O\ R\ [Z\ H\ R\ W\ K\ U\ D\ F\ K\ L\ H\ Y\ H\ G\ H\ I\ I\ L\ F\ L\ H\ Q\ F\ \ D\ O\ O\ D\ I\ W\ H\ U\ G\ D\ V\ F\ R\ Q\ I\ L\ U\ P\ L\ Q\ J\ S\ R\ Z\ H\ U\ I\ X\ O\ D\ E\ L\ O\ L\ W\ \ R\ I\ W\ K\ H\ DXJPHQWHG EDFWHULD

%LRFKHPLFDO R[\JHQ GHPDQG %2'

7KH UHFRUGHG %2' OHYHO LQ WKH UDZ HIOXHQW PJ O ZDV SURSRUWLRCDDQO\ GHFUHDVHG ZLWK WLPH LQ DOO WKH VHHGHG ZDVWHZDWHU VDPSOHV RQO\ UHDFKLQJ WKH ORZHVW UHVLGXDO FRQFHQWUDWLRQ 5& DW WKH ODVW H[SRVXUH GD\)LJXUH & 'XULQJ WKH ILUVW K IRXU ZDVWHZDWHU VDPSOHV 6 36 PL[HG FXOWXUH DQG WKH FRQWURO VKRZHG LQFUHDVHV LQ WKH %2' ZLWK GLIIHUHQW SURSRUWLRCDDQO\ 36 DQG WKH PL[HG FXOWXUH ZDVWHZDWHVVKRZHG YHU\ VOLJKW %2' LQFUHDVHV ZKLOH 6 V\ OHG HG DQG FRQWURO ZDVWHZDWHVVKRZHG VLJQLILFDQWO\ KLJKHU LQFUHDVHV DQG UHVSHFWLYHO\ 7KHVH LQFUHDVHV UHIOHFW WR[LFLW\ RI WKH ZDVWHZDWHU WKDW OHG WR WKH GHDWK RI VRPH EDFWHULD FHOOV DGGLQJ WKHLU RUJDQLF P\ WR WKH ZDVWHZDWHU FRQWDLQHG WKHP 5] DQG 5] UHSUHVHQW WKH PRVW UHVLWDQW VWUDLQV IROORZHG E\ 36 WKH PL[HG FXOWXUH DQG ILQDOO\ 6 %DFWHULD LQKDELWLQJ WKH GRPHVWLF ZDVWHZDWHU UHSUHVHQW WKH PRVW VHQLWLYH VWUDLQV DQG VKRZHG UHJXODU GHDWK DQG LQFUHDVHV LQ WKH %2' ZLWK WLPH LQ WKH FRQWURO ZDVWHZDWHU UHDFKLQJ WKH KLJKHVW OHYHO DIWHU GD\ V

5HPRYDO IILFLHQF\ RI WKH %2' LQFUHDVHG UHJXODUO\ LQ DOO WKH VHHGHG ZDVWHZDWHU VDPSOHV XQWLO WKH ODVW H[SRVXUH GD\ 36 H[KLELWHG WKH KLJKHVW 5(RI %2' IROORZHG E\ WKH PL[HG FXOWXUH 5] 5] DQG ILQDOO\ 6 ZLWK WKH OHDVW 5(UHFRUGLQJ DQG UHVSHFWLYHO\ DIWHU GD\ V +RZHYHU FRQWURO ZDVWHZDWHU GLG QRW DFKLH DQ\ UHPRYDO EXW H[KLELWHG %2' DGGLWLRQ L H IROG RI WKH LQLWDO %2' DIWHU WKH VDPH WLPH

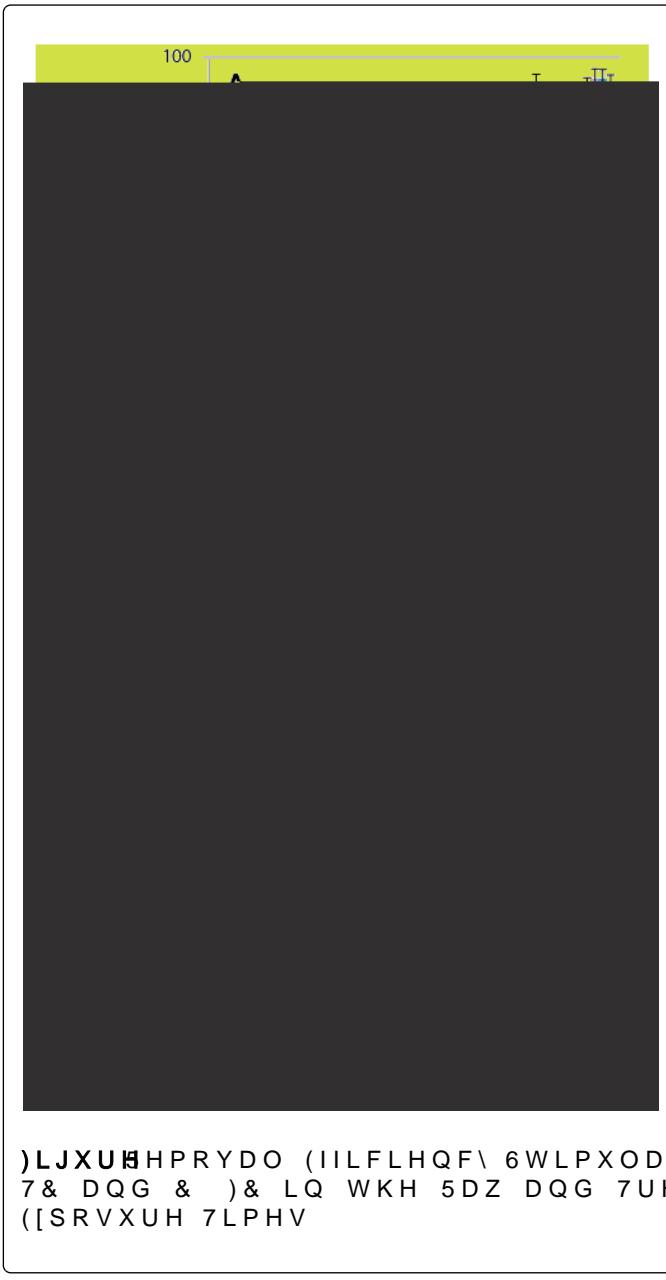
)DW RLO DQG JUHDVH)2*

)2* LQ WKH UDZ ZDVWHZDWHU UHFRUGHG PJ O ZKLFK FRQVLGHUHG KLJK OHYHO 7KH EXON UHPRYDOV RI)2* WRRN SODFH GXULQJ WKH ILUVW K DQG JUDGXDOO\ GHFUHDVHG ZLWK WLPH LQ DOO WUHDWPHQWV WLOO WKH H[SRVXUH GD\)LJXUH \$ 5] DQG 5] H[KLELWHG WKH KLJKHVW 5(IROORZHG E\ WKH PL[HG FXOWXUH 36 DQG ILQDOO\ 6 DQG UHVSHFWLYHO\ +RZHYHU FRQWURO ZDVWHZDWHU VKRZHG RQO\ FRQILUPLQJ WKH DELOLW\ RI WKH ELRDXJPHQWDWLRQ RI EDFWHULD

7RWDO FROLIRUPV 7& DQG IHFDO FROLIRUP)&

6LQFH WKH SUHVHQW VWXG\ GHDOV ZLWK GRPHVWLF ZDVWHZDWHU WRWDO DQIHFDO FROLIRUP 7& DQG)& DUH WKH PRVW LPSRUWDQW TXDOLW\ SDUDPHWHUV GXH WR WKHLU HIIHFW RQ WKH UHFHLYLQJ ZDWHU ZLWK WKHLU SDWKRJHQLF OR7KH\ DOVR GHWHUPLQH WKH IILFLHQF\ RI WKH WUHDWPHQW SURFHV (OLPLQDWLRQ RU PLQLPL]DWLRQ RI 7& DQG)& FRXQW LQGLFDWHV HILFLHQW WUHDWPHQW ZKLOH LQFUHDVLRQ 7& DQG)& LQGLFDWHV WKH RSSRVLWH 5(RI 7& DQG)&)LJXUH % DQG & ZHUH SRVLWLYHO\ FRUUHODWHG ZLWK WLPH LQ DOO VHHGHG ZDVWHZDWHVVKRZHG YHU\ VOLJKW LQFUHDVHV LQ WKH PRVW UHPRYDO EXW H[KLELWHG %2' DGGLWLRQ L H IROG RI WKH LQLWDO %2' DIWHU WKH VDPH WLPH

GLUHFWO\ DIIHFWHG LWV SHUIRUPDQF MV KXHS PDRQ/GW GHR ZLQFL & Q W B R OG B B R Q/W D B WGDW
VDPSON VKRZHG WKH RSSRVLWH WUHQ SSUHWHQ FVR DW&COXROXVHA QBUQ B DORQ QXV@UAH pM
VWDUWLQJ IURP DIWHU RQH GD\ WR DIWHU GD\V DQG FRQWLQXRXV
LQFUHDVH LQ WKH)& UDQJHG EHWHQ D PLQLPXP RI DQG PD[LPXP
RI LQGLFDWLQJ WKH DEVHQFH RI DQWDJRQLVWLF EDFWHULD WKDW FDQ NLO
DQG DVVLPLQDWLQJ FROLIRUP EDFWHULD



)LJXUHPRYDO (IILFLHQF\ 6WLPXODWLRQ 5(6 RI \$)2* %
7& DQG &)& LQ WKH 5DZ DQG 7UHDWHG :DVWHZDWHU DW 'LIIHUVHQW
([SRVXUH 7LPHV

'LVFXVVLRQ

\$PRQJ WKH ZDVWHZDWHU LQGLJHQRXV LVRODWHV RQO\ WZR 5] DQG
5] DV ZHOO DV WZR H[RJHQRXV VVUDLQV 6 DQG 36 VKRZHG WKH DELOLW\ WR
FODULI\ GRPHVWLW ZDVWHZDWHU LQGLFDWLQJ WKDW WKH\ SRVVHVW WKH UHTXL
GHJUDGLQJ HQ]\PHV OROHFXODU FKDUDFWHUL]DWLRQ RI WKH PRVW DFWLYH
EDFWHULD UHYHDO HIGFKHVKHULFJEB OFRQLGQQLW
YHUPLPSRQD DFLVSDXV6 3 DQGG RPARSQD 186 7KH
PL[HG FXOWXUH FRPELQDWLRQ ZLWK D IRXU VHOHFWHG FXOWXUHV SURYHG WR

ZDV DFKLHYHG XV⁶ DIWHU GD\ ZKLFK LV IROG KLJKHU WKDQ LWV PD[LPXP
6

7LOOH\ ') \$HURELF :DVWHZDWHU 7UHDWPHQW 3URFHVVHV +LVWRU\ DQG
'HYHORSPHQW ,:\$ 3XEOLVKLQJ
+HQ]H YDQ /RRVGUHFKW O&O (NDPD *\$ %UGMDQRYLF ' HGV
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(O%HVWDZ\ ((O \$ODVU\ 0+ (O \$GO 1 7KH SRWHQWLDO\ RI IUHH
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(O%HVWDZ\ (+XVVHLQ + %DJKGDGL ++ (O 6DND 0) &RPSDULVRQ
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QLWURJHQ DQG SKRVSKRUXV - ,QG0LFURELRO%LRWHFKQRO
(O%HVWDZ\ ((O 6RNNDU\ , +XVVHLQ +)DURXN \$ 3ROOXWLRQ
FRQWURO LQ SXOS DQG SDSHU LQGXVWULDO HIOXHQWV XVLQJ LQWHJUDWLG FKHPLFDO
ELRORJLFDO WUHDWPHQW VHTXHQFHV -RXUQDO RI ,QGXVWULDO 0LFURELRORJ\ DQG
%LRWHFKQRORJ\
&OHVFBHU*WHHQEHUJ &* (DWRQ \$' 6WDQGDUG 0HWKRQG IRU WKH
([DPLQDWLRQ RI :DWHU DQG :DVWHZDWHU WKHGQ \$PHULFDQ 3XEOLF
+HDOWK \$VVRFLDWLRQ \$3+\$:DVKLQJWRQ
0D < 'LVVHQ *\$ 5DJH) 2MHGD 65 51DVH 3URWHFWLRQ \$VVD\
0HWKRGV
\$OWV6FKXDVK : 0LOOHU : 0\HUV (: /LSPDQ '- %DVLF ORFDO
DOLJQPHQW VHDUFK WRRO - 0RO%LRO
+DOO %. +RPRORJ\ ,QWURGXFWLRQ 1RYDUWLV)RXQG 6\PS
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