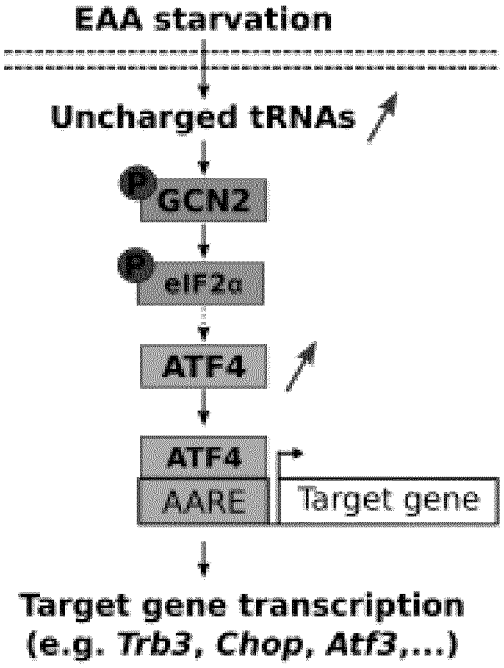


FIGURE 1

AAREs driven expression of a Cas nuclease

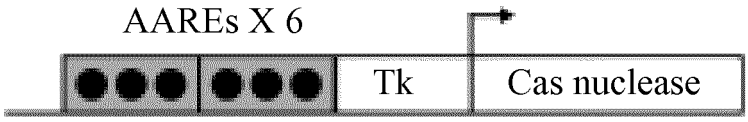


FIGURE 2

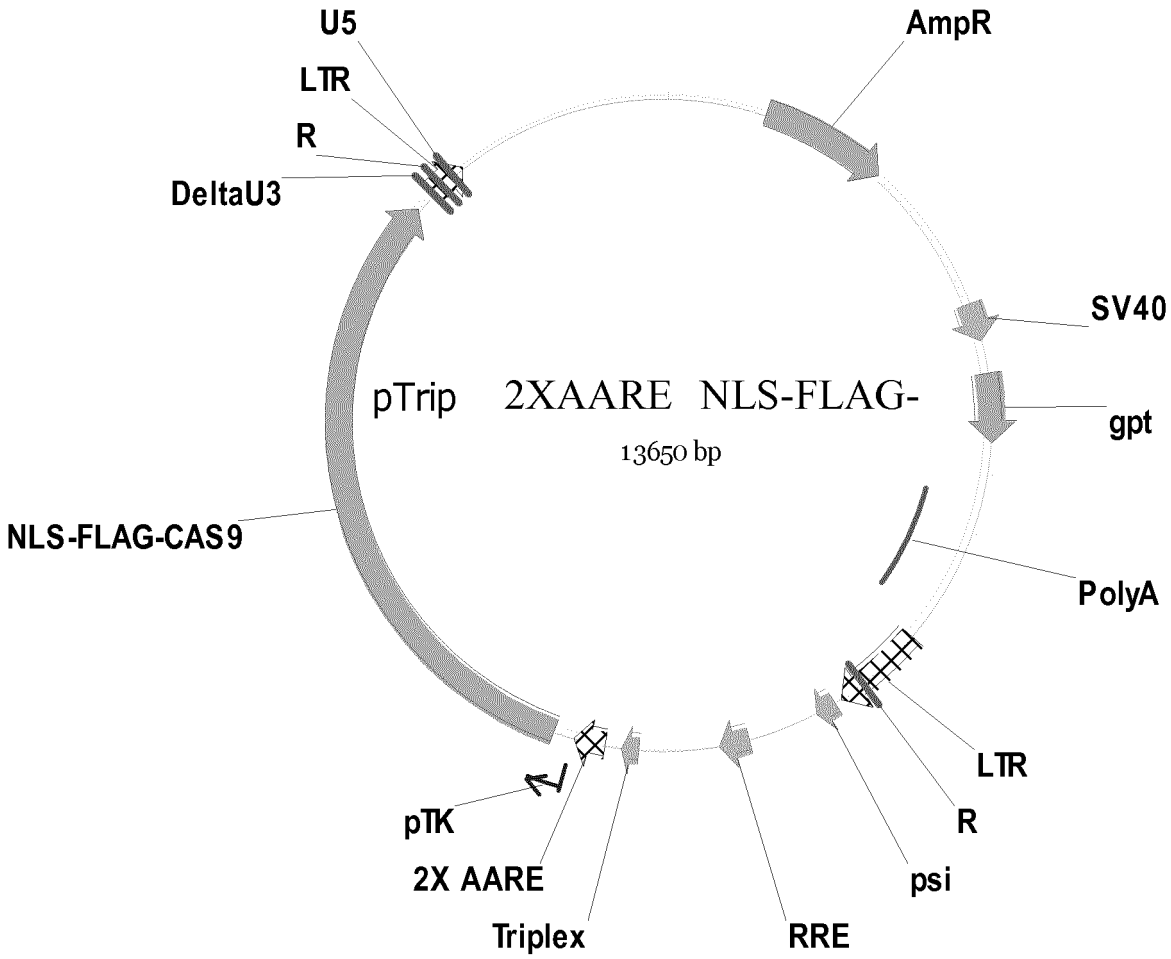

U5
LTR
R
DeltaU3
AmpR
SV40
gpt
PolyA
LTR
R
psi
RRE
Triplex
2X AARE
pTK
NLS-FLAG-CAS9
pTrip 2XAARE NLS-FLAG-
13650 bp

FIGURE 3

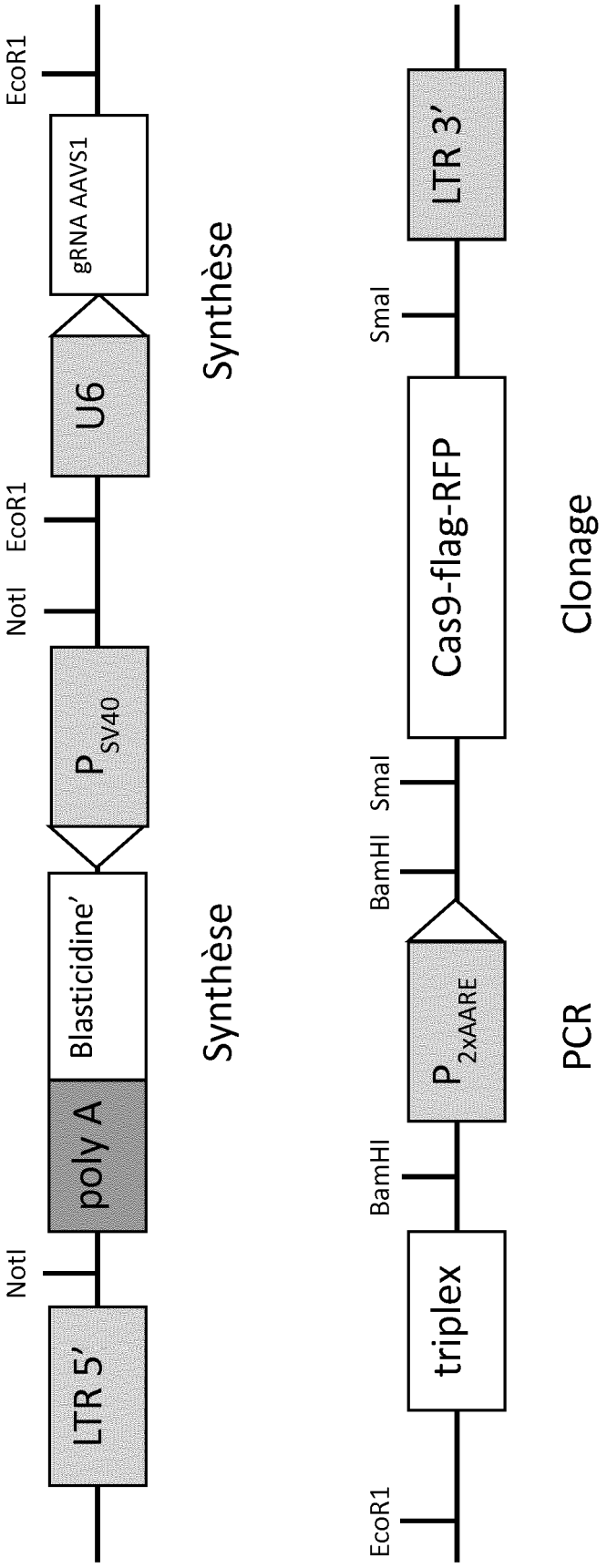

LTR 5'
NotI
poly A
Blasticidine'
Synthèse
NotI
EcoR1
P SV40
U6
EcoR1
gRNA AAVS1
Synthèse
EcoR1
triplex
BamHI
P 2xAARE
BamHI
SmaI
Cas9-flag-RFP
SmaI
LTR 3'
PCR
Clonage

FIGURE 4

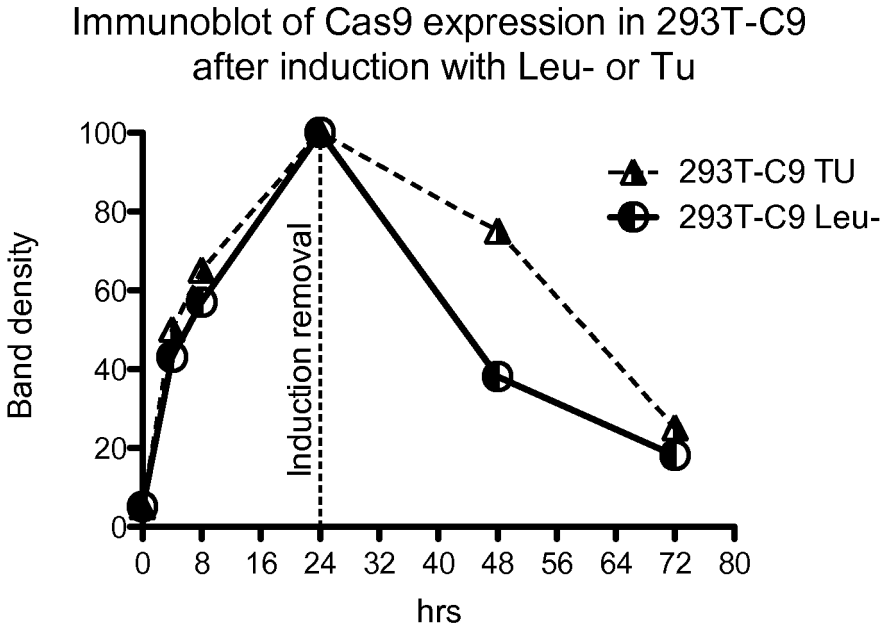


FIGURE 5

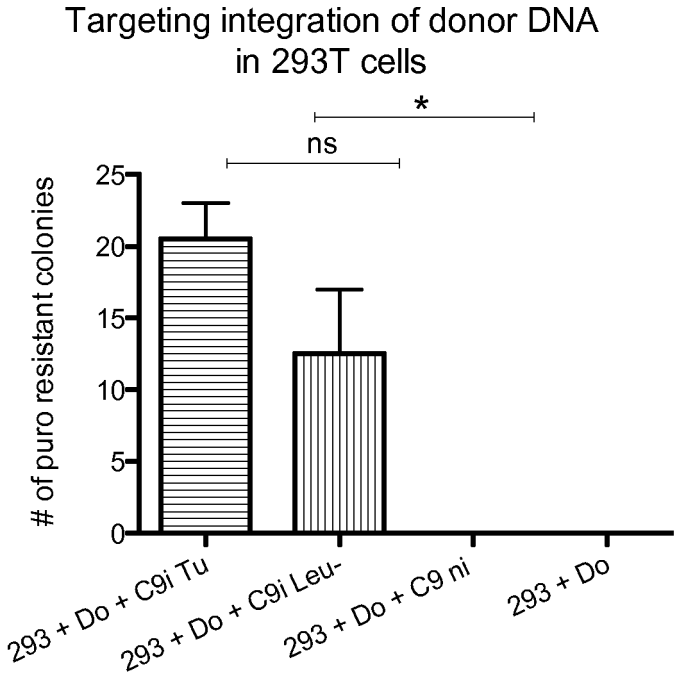


FIGURE 6

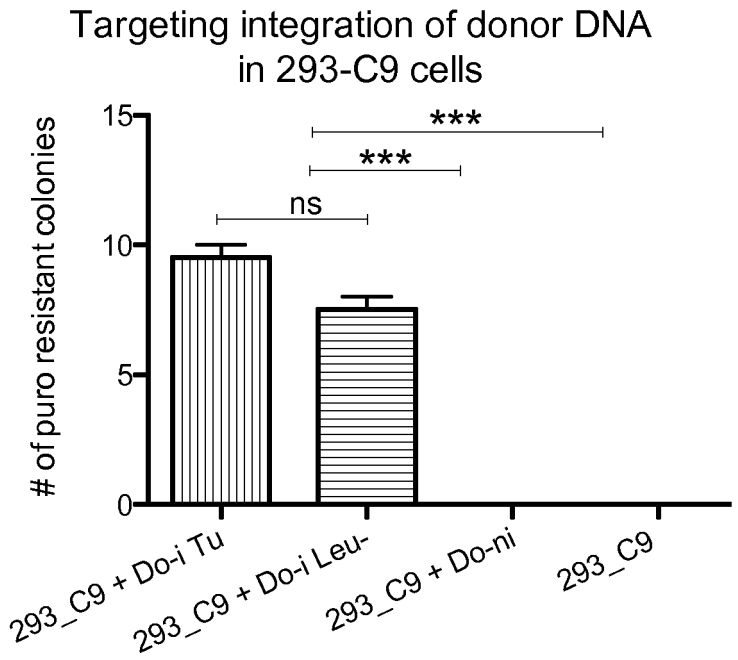

Targeting integration of donor DNA
in 293-C9 cells
# of puro resistant colonies
15
10
5
0
***
***
ns
293_C9 + Do-i Tu
293_C9 + Do-i Leu-
293_C9 + Do-ni
293_C9

FIGURE 7