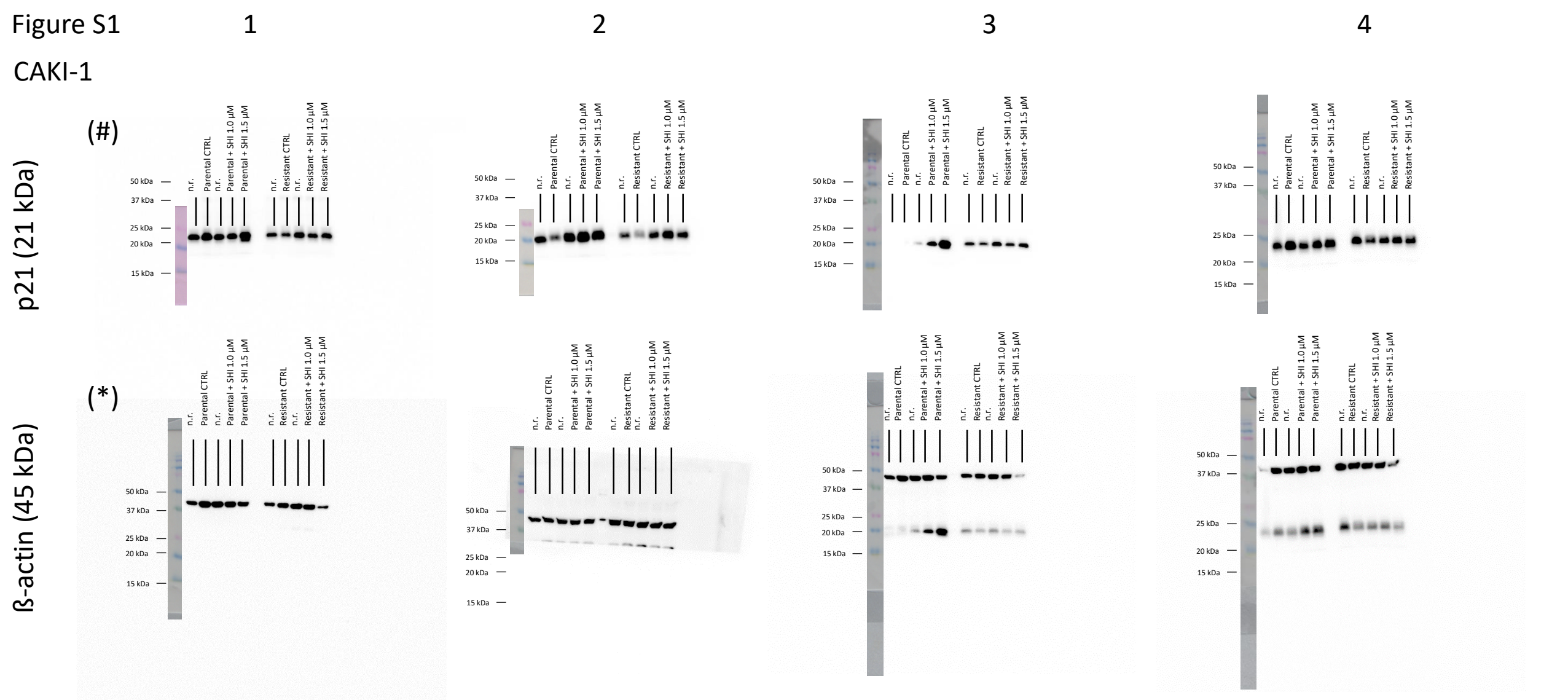


Cell cycle regulating proteins

p21, p27, Cyclin A, Cyclin B, CDK1, pCDK1, CDK2, pCDK2



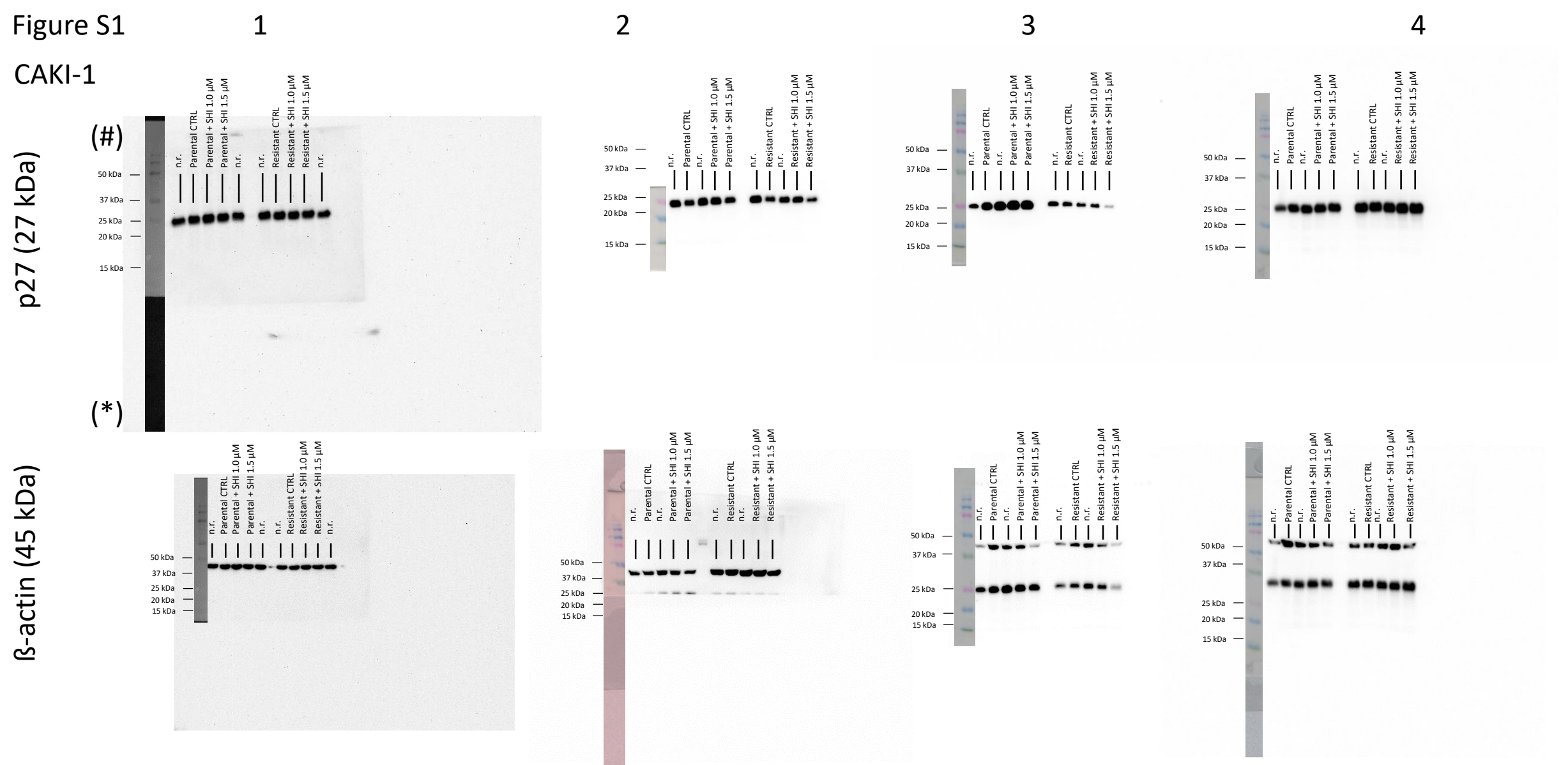


Figure S1b: Detailed information about Figure 4 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant Caki-1 cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of p27 (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

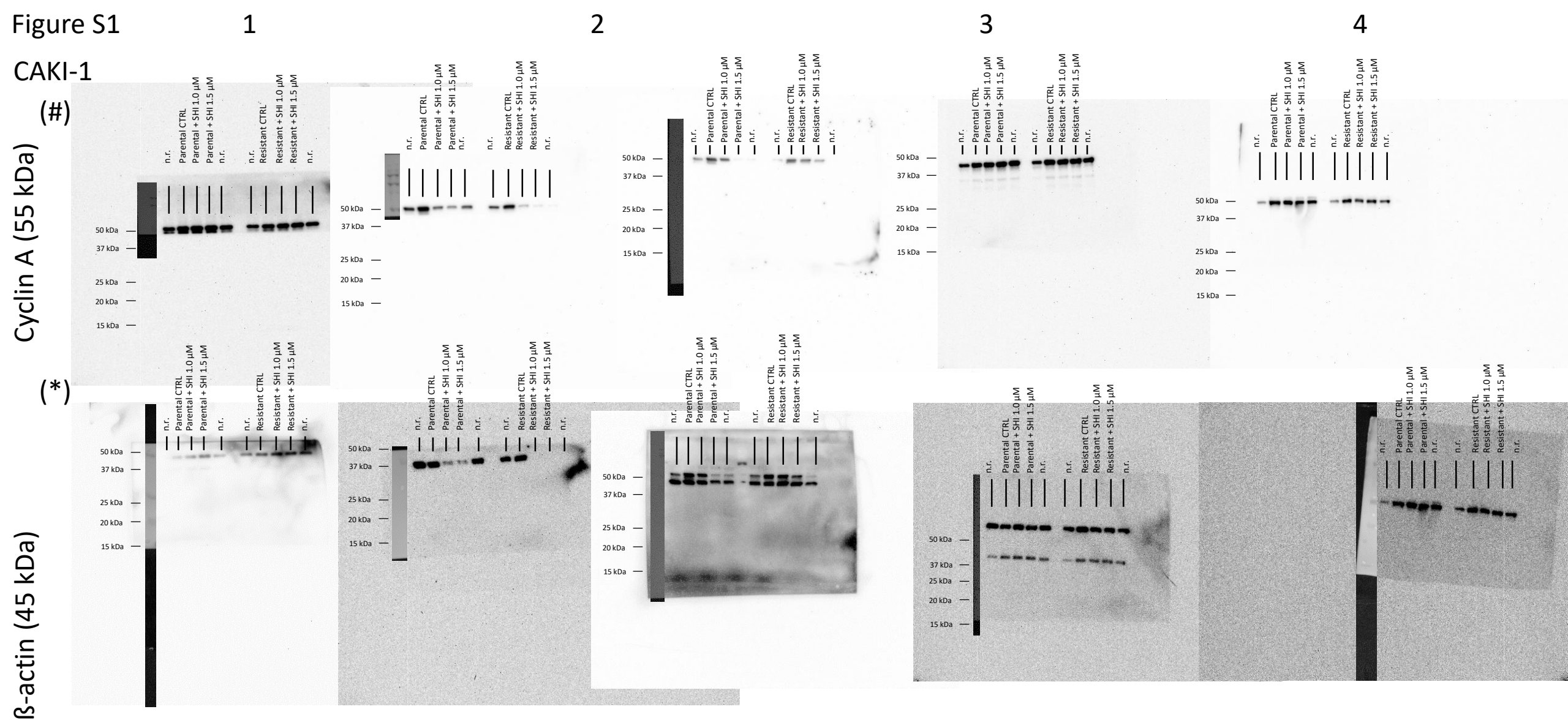


Figure S1c: Detailed information about Figure 4 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant Caki-1 cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of Cyclin A (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

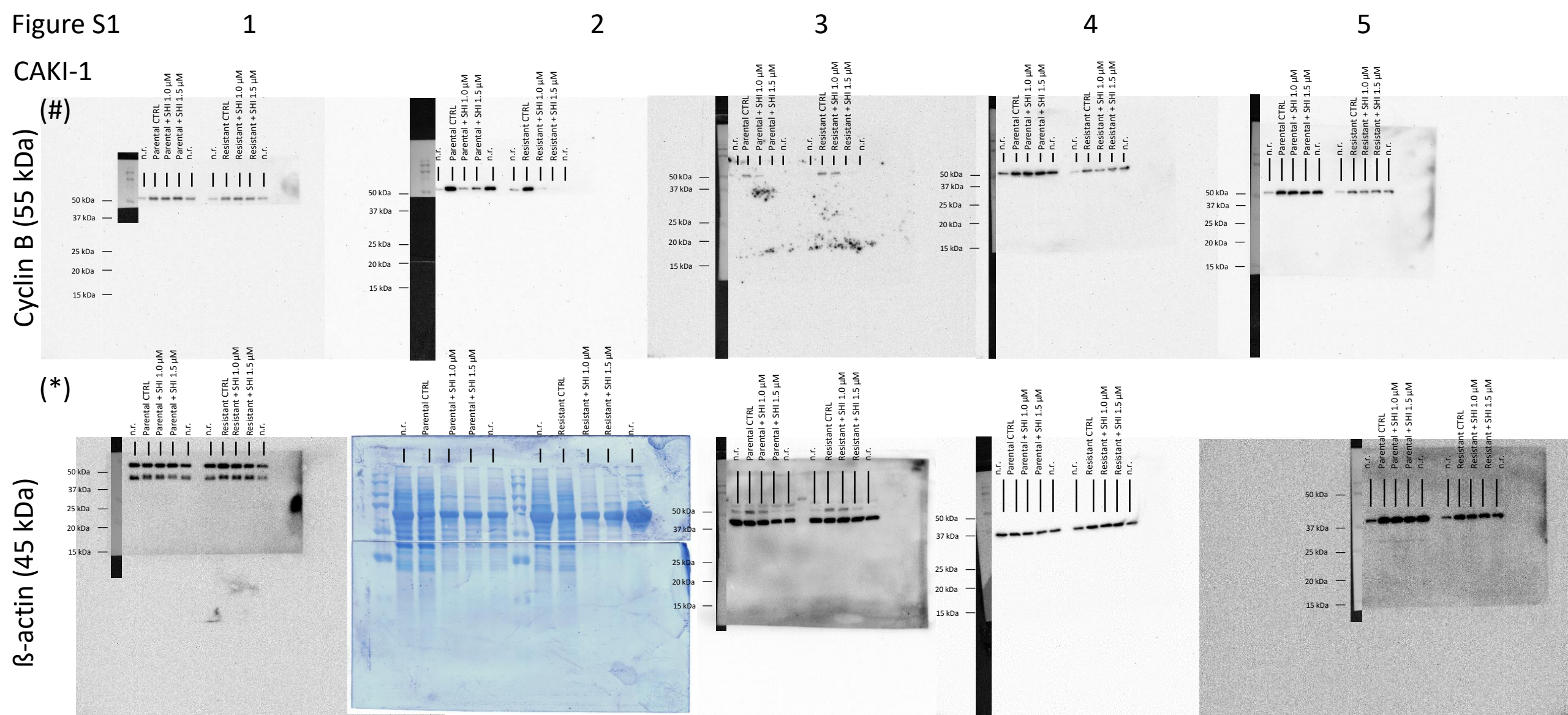


Figure S1d: Detailed information about Figure 4 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant Caki-1 cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of Cyclin B (#), corresponding protein expression of β -actin or Coomassie blue staining of total protein (*). n.r. = not relevant.

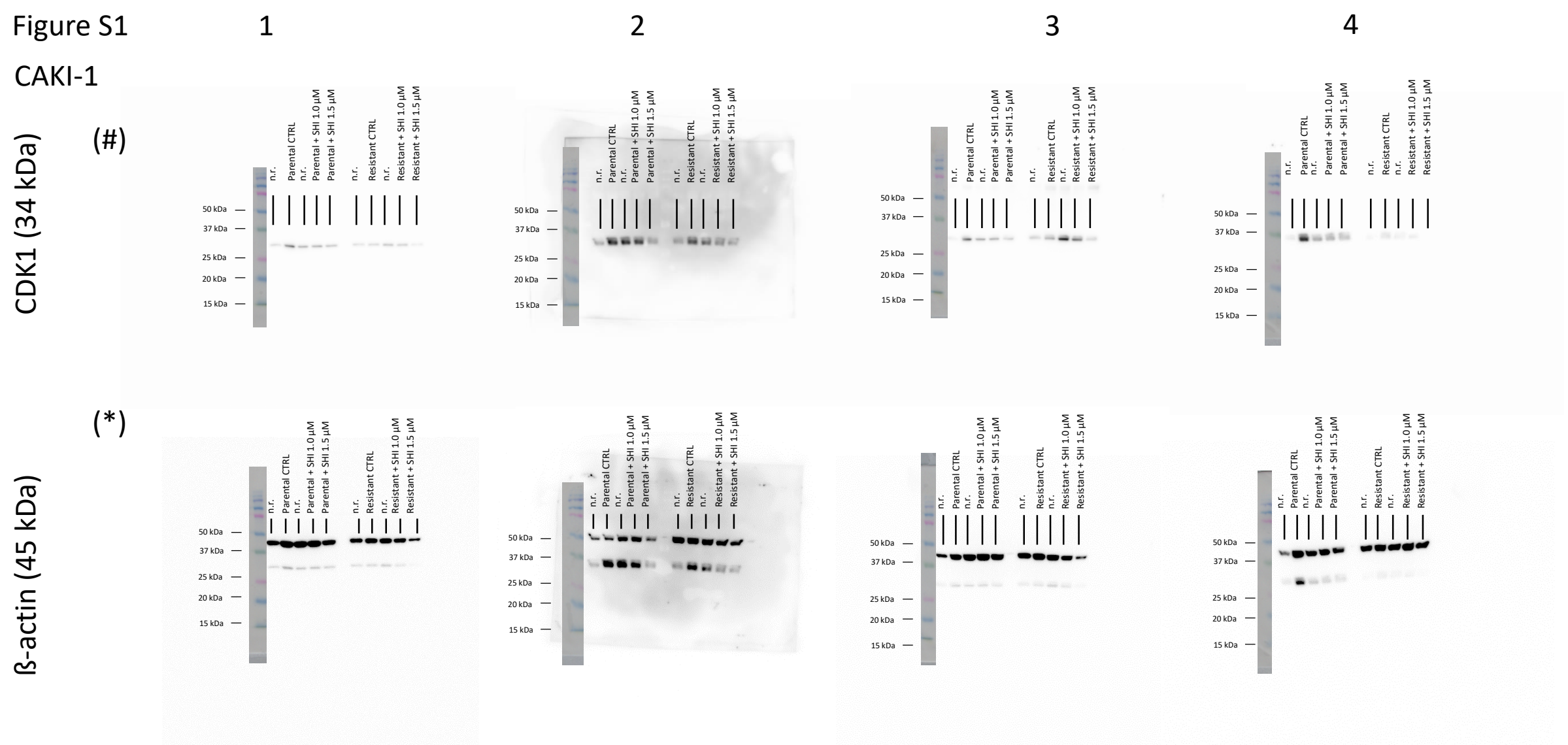
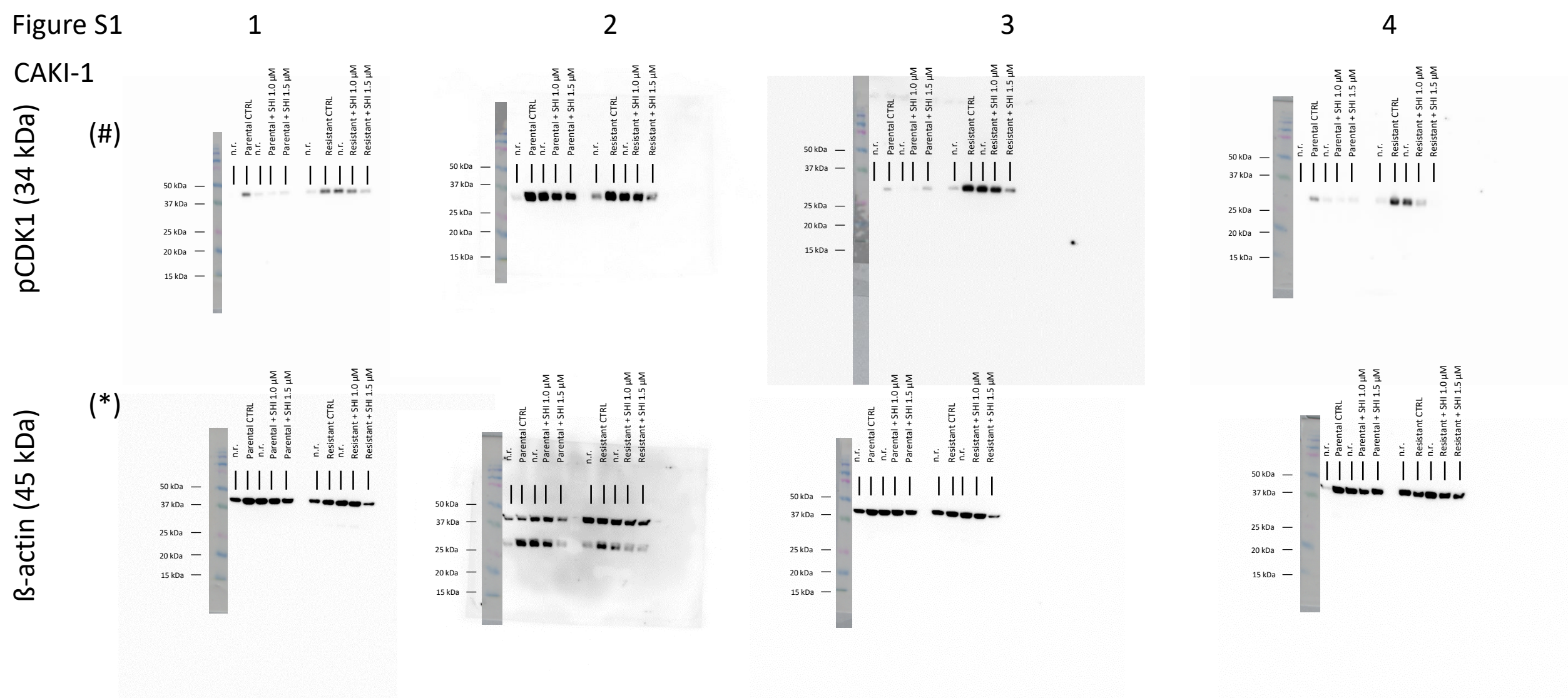


Figure S1e: Detailed information about Figure 4 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant Caki-1 cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of CDK1 (#), corresponding protein expression of β -actin (*). n.r. = not relevant.



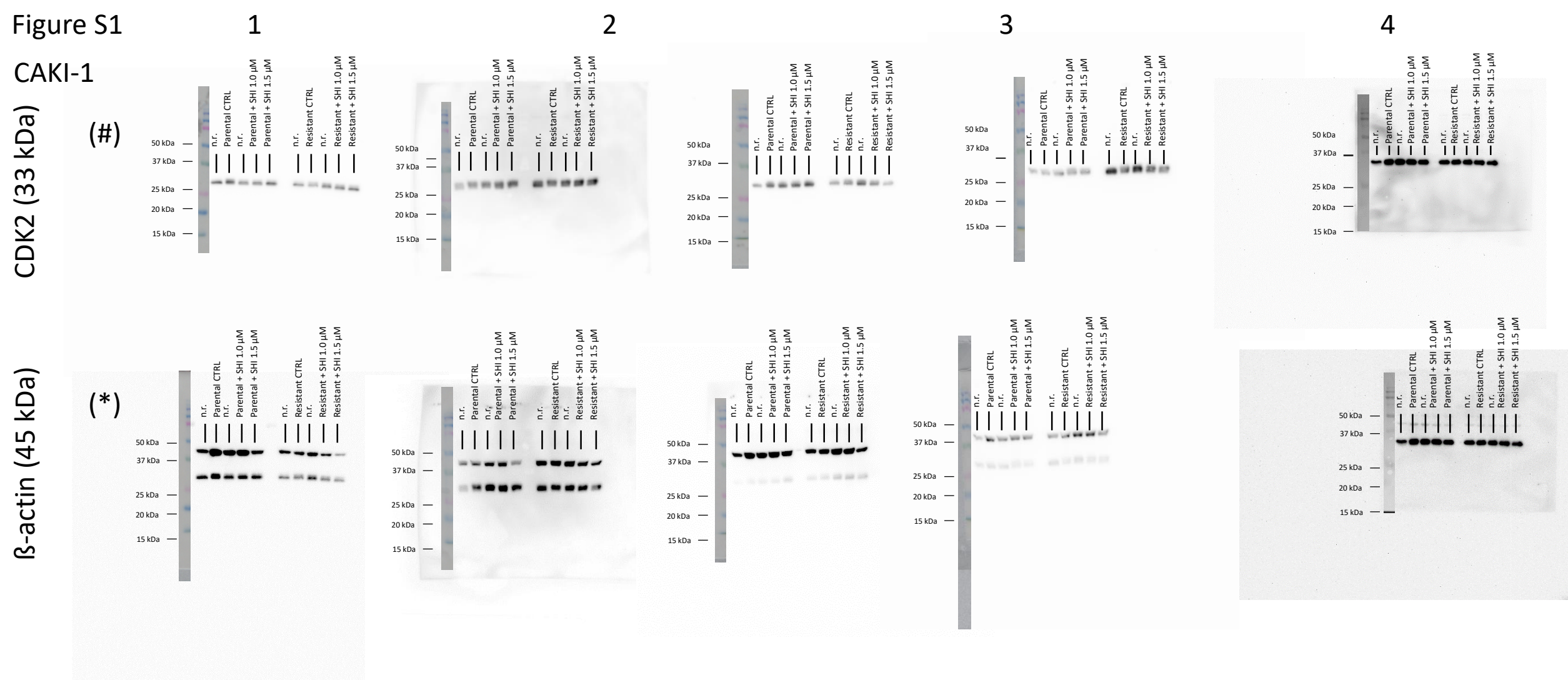


Figure S1g: Detailed information about Figure 4 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant Caki-1 cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of CDK2 (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

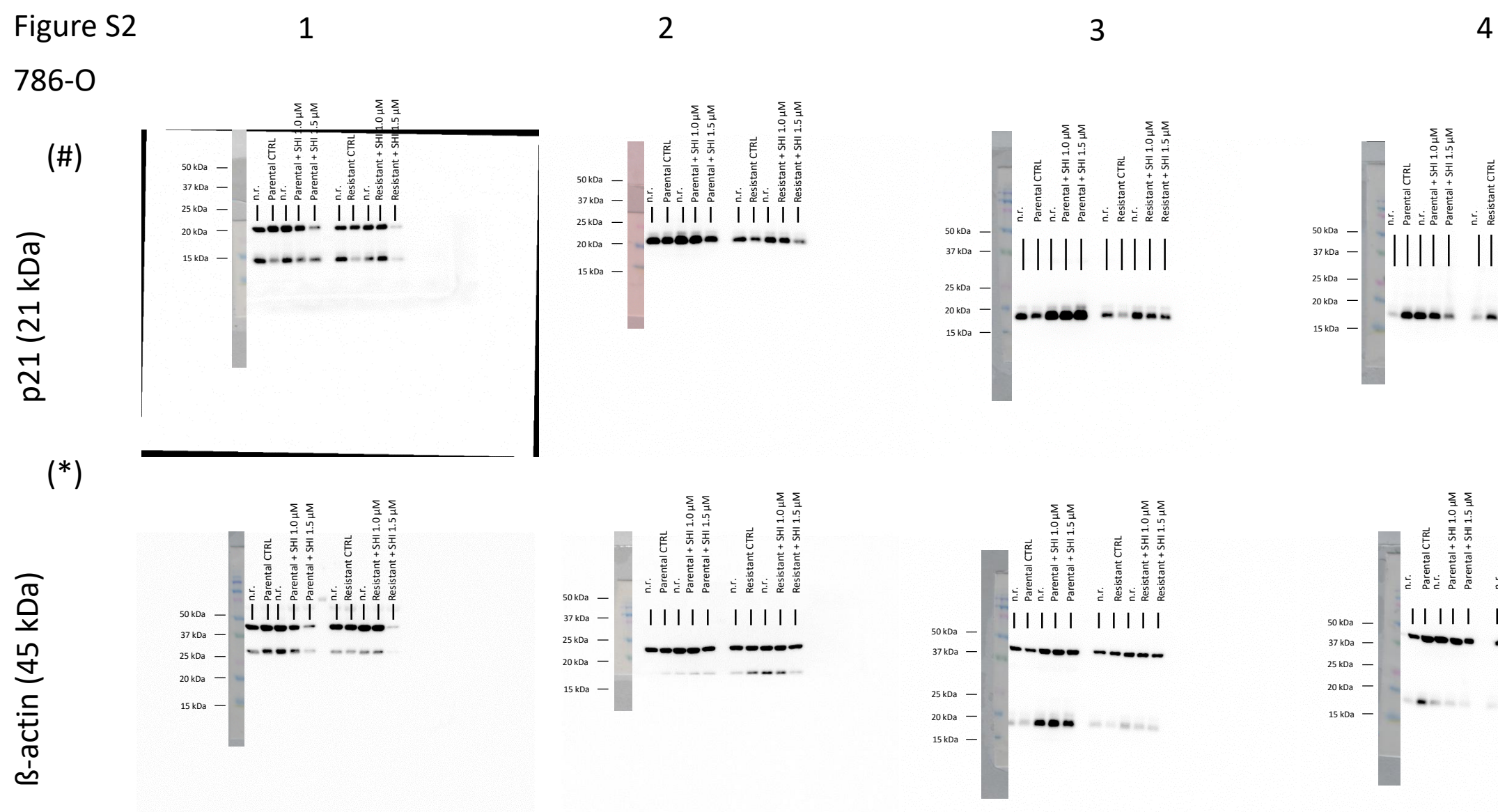


Figure S2a: Detailed information about Figure 5 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant 786-O cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of p21 (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

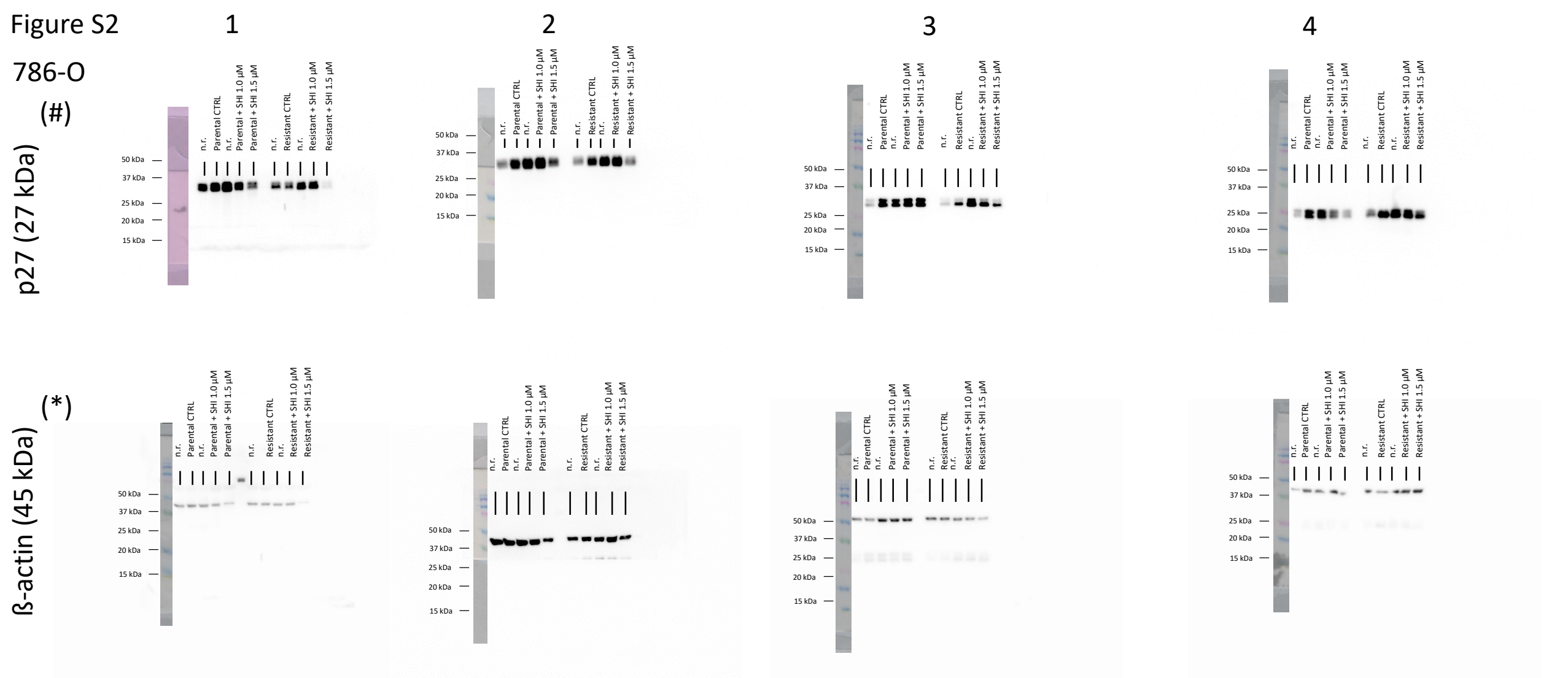


Figure S2b: Detailed information about Figure 5 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant 786-O cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of p27 (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

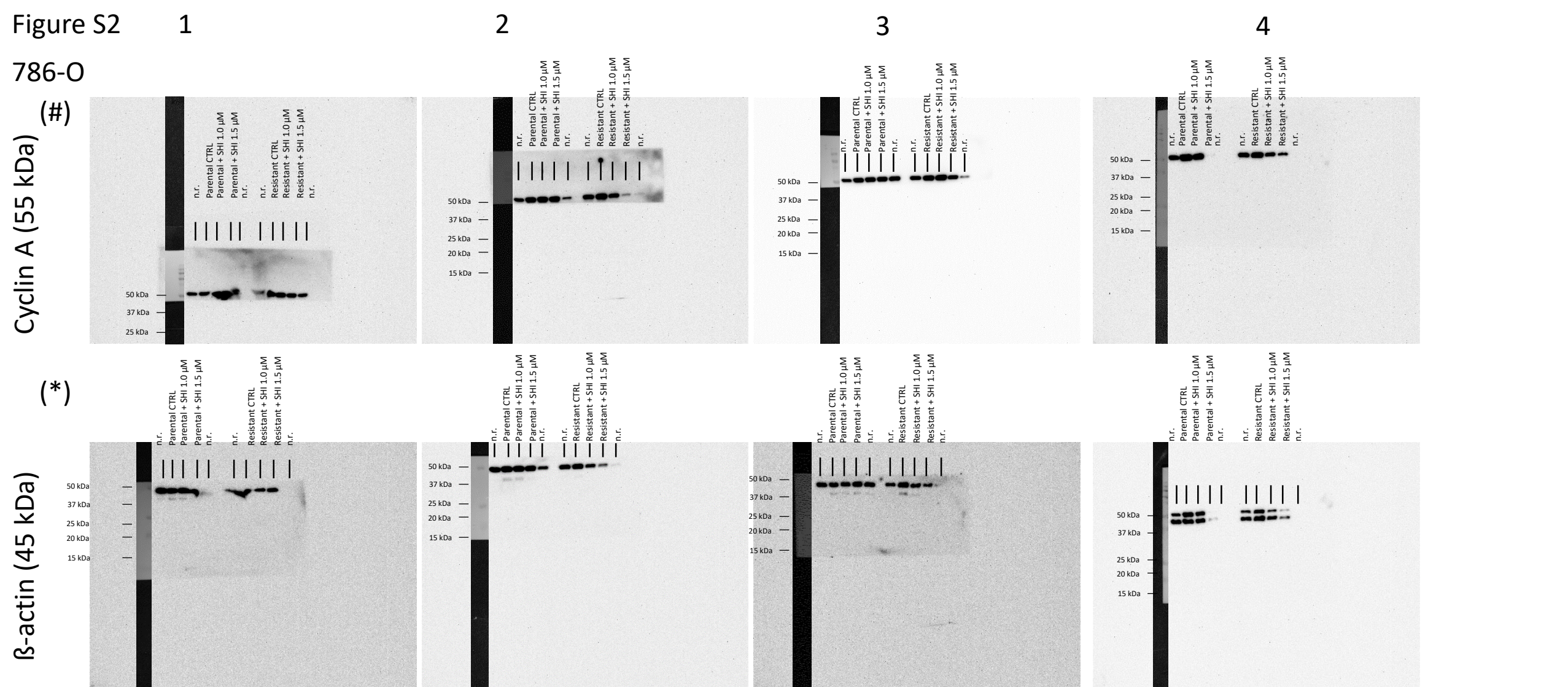


Figure S2c: Detailed information about Figure 5 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant 786-O cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of Cyclin A (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

Figure S2

786-O

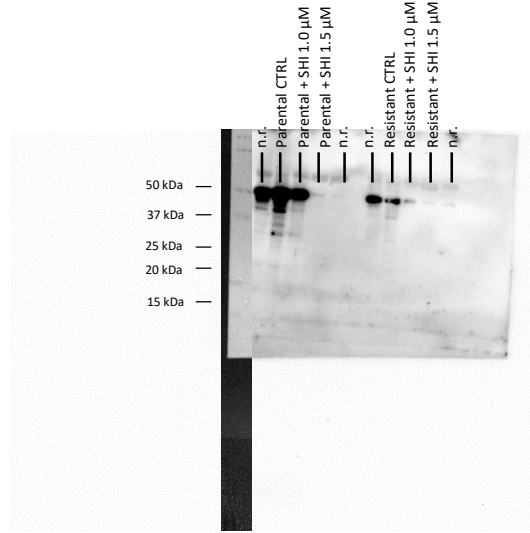
(#)

Cyclin A (55 kDa)

(*)

β -actin (45 kDa)

5



6

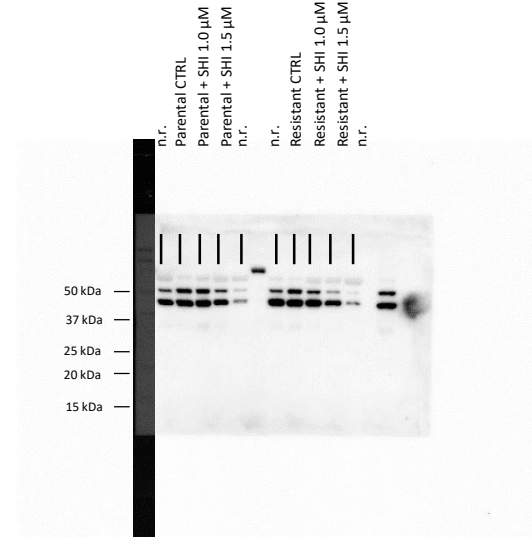
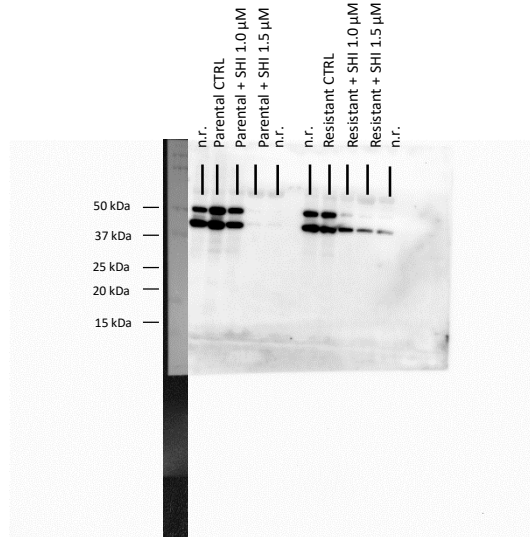
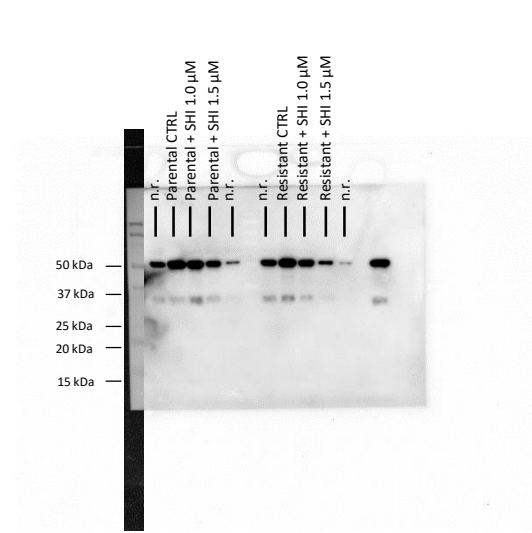


Figure S2c: Detailed information about Figure 5 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant 786-O cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of Cyclin A (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

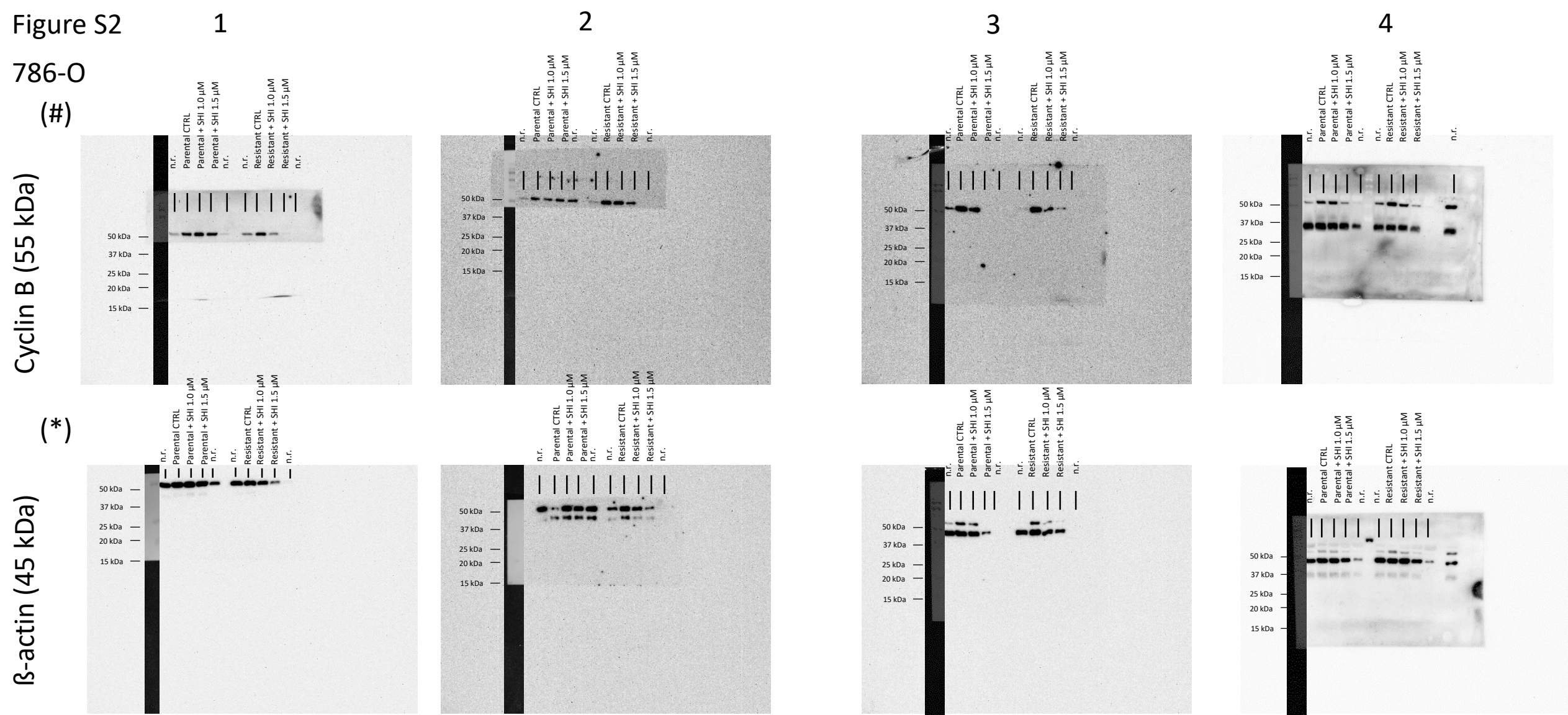


Figure S2d: Detailed information about Figure 5 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant 786-O cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of Cyclin B (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

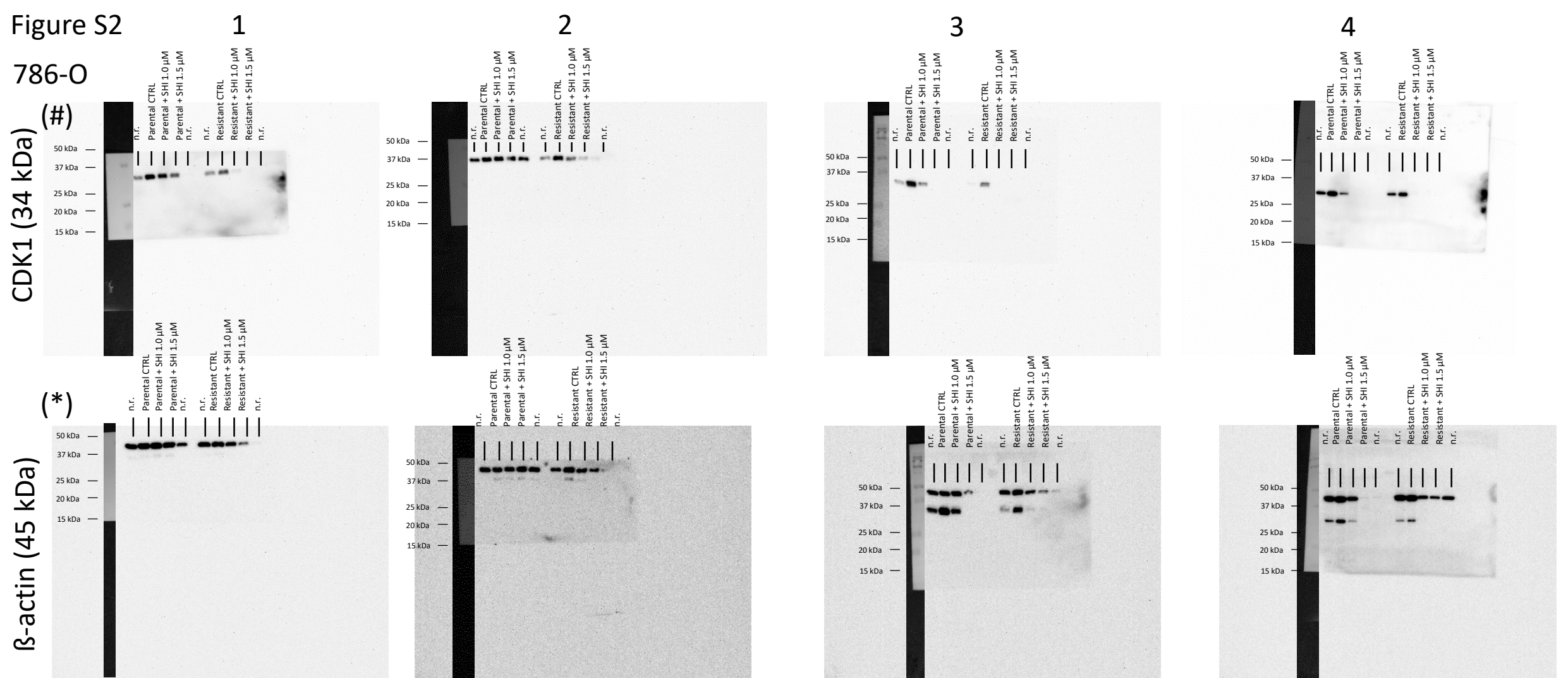


Figure S2e: Detailed information about Figure 5 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant 786-O cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of CDK1 (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

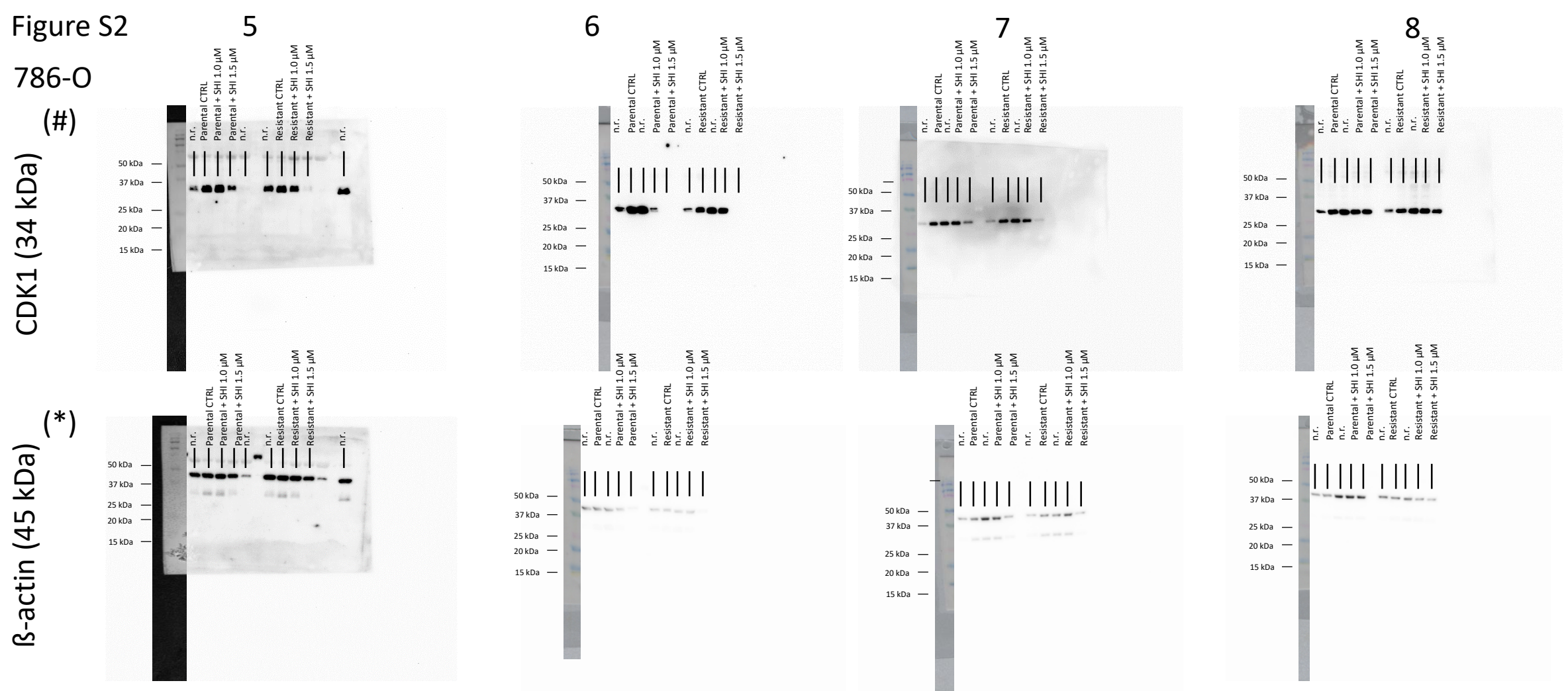


Figure S2e: Detailed information about Figure 5 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant 786-O cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of CDK1 (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

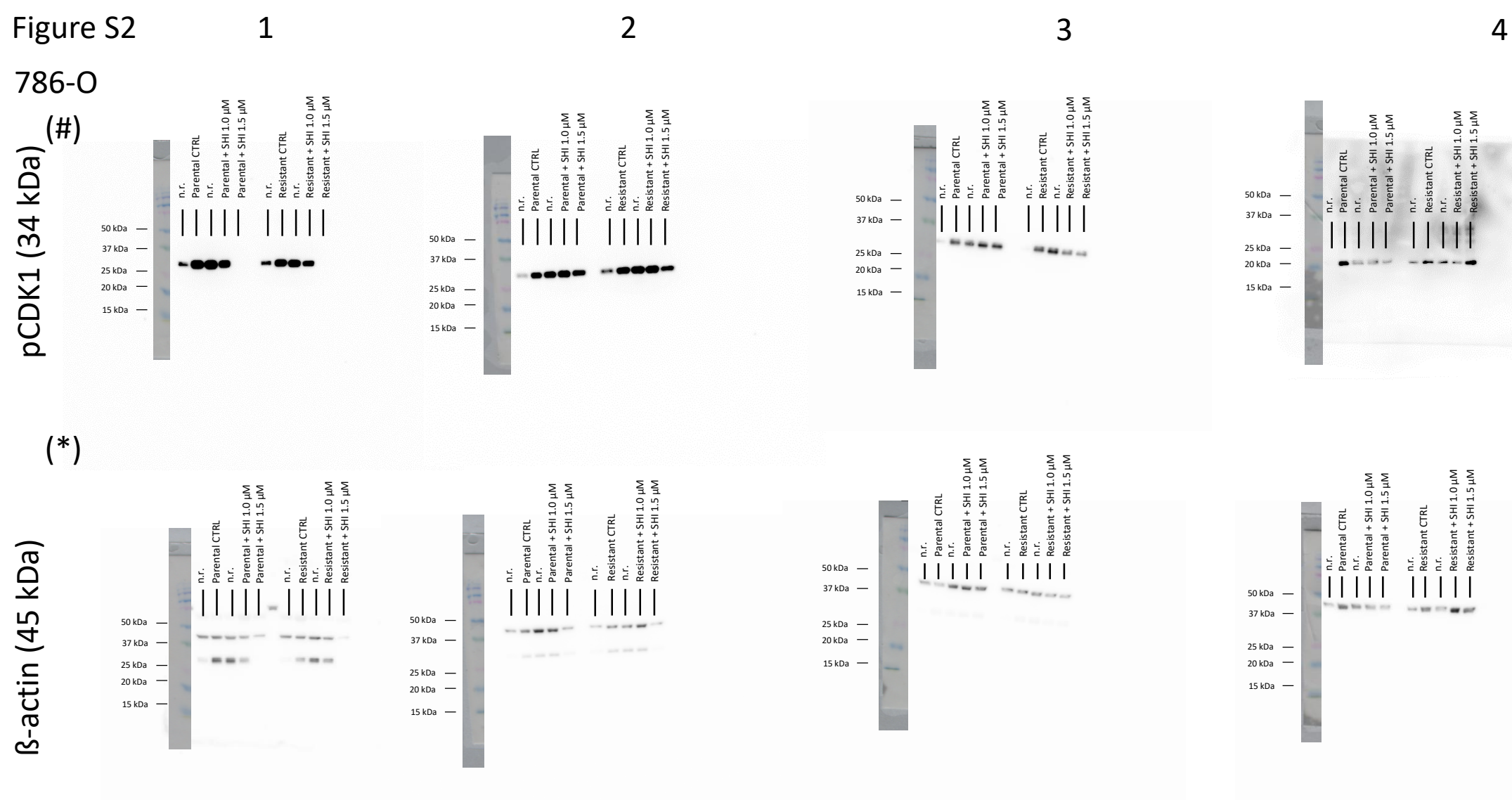


Figure S2f: Detailed information about Figure 5 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant 786-O cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of pCDK1 (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

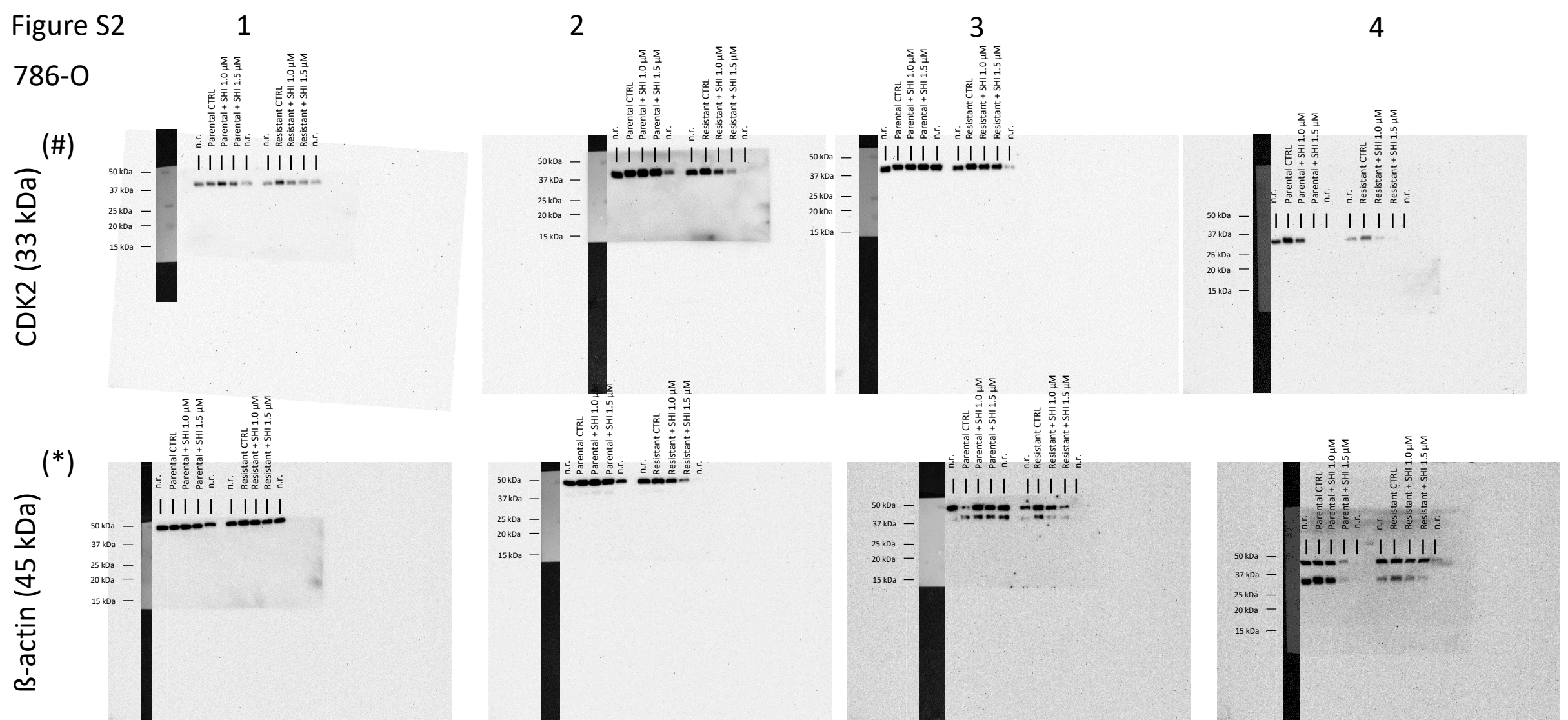


Figure S2g: Detailed information about Figure 5 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant 786-O cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of CDK2 (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

Figure S2

786-O

(#)

CDK2 (33 kDa)

(*)

β -actin (45 kDa)

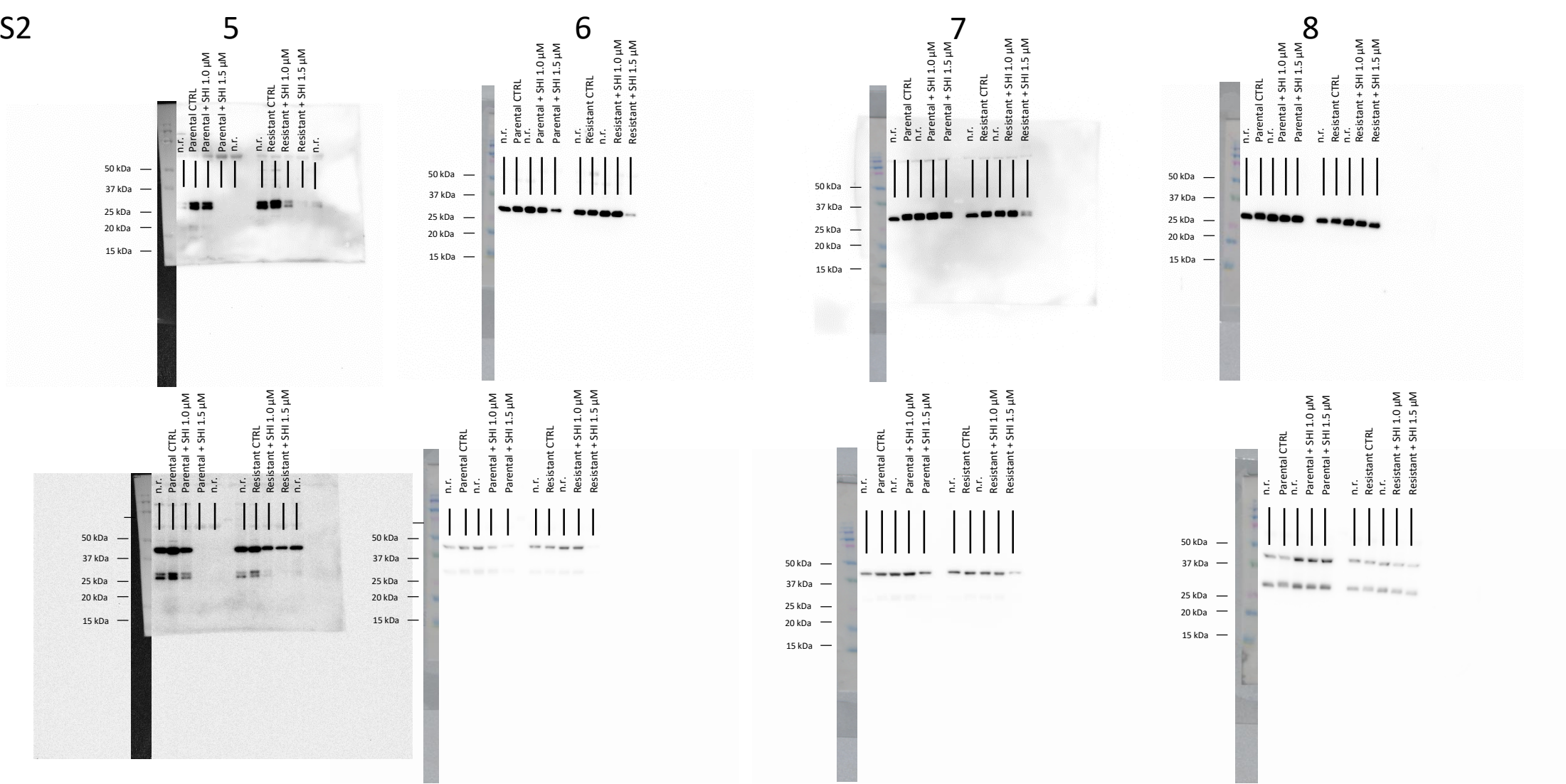


Figure S2g: Detailed information about Figure 5 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant 786-O cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of CDK2 (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

Figure S2

786-O

(#)

(*)

β -actin (45 kDa)

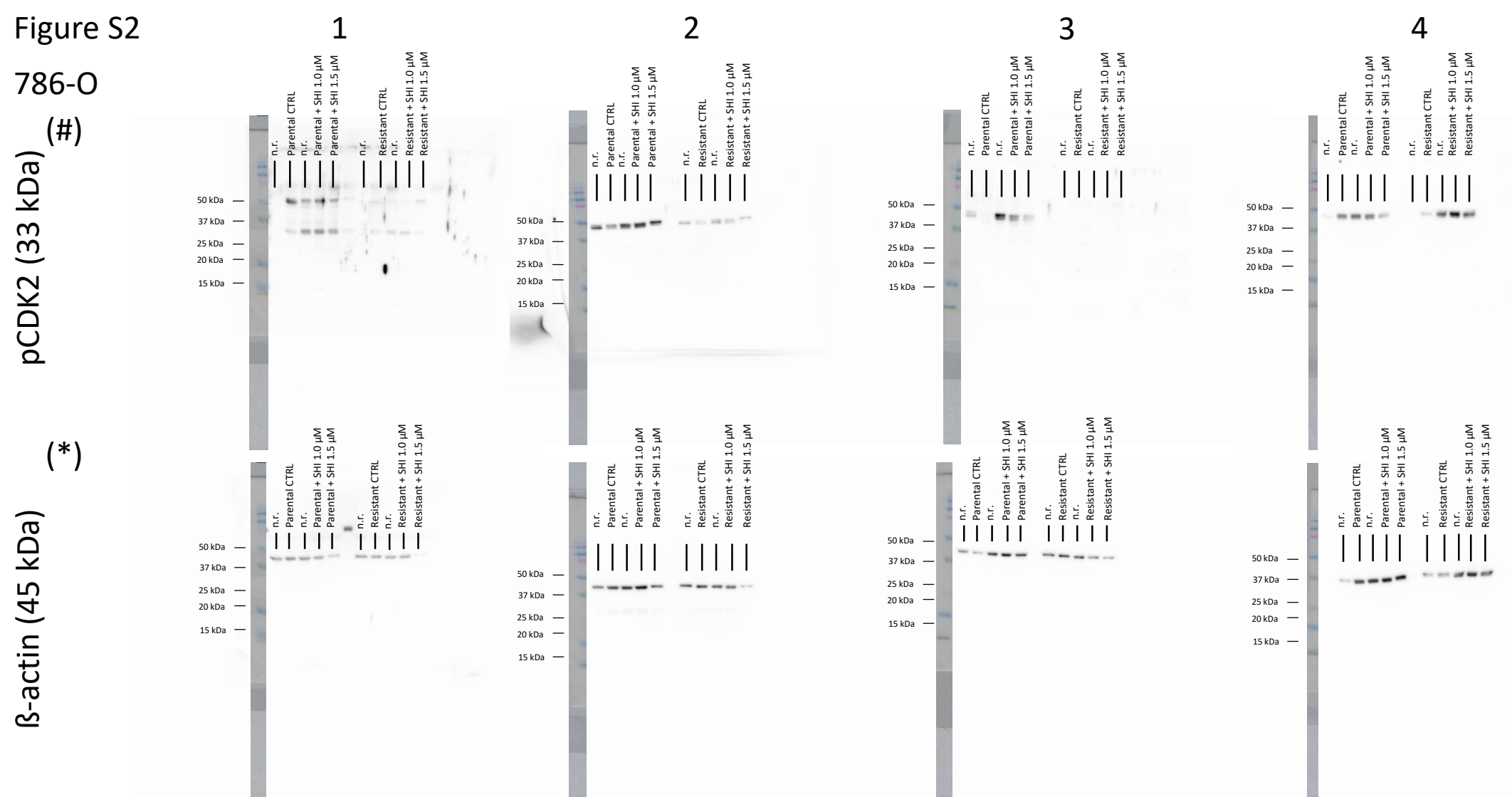


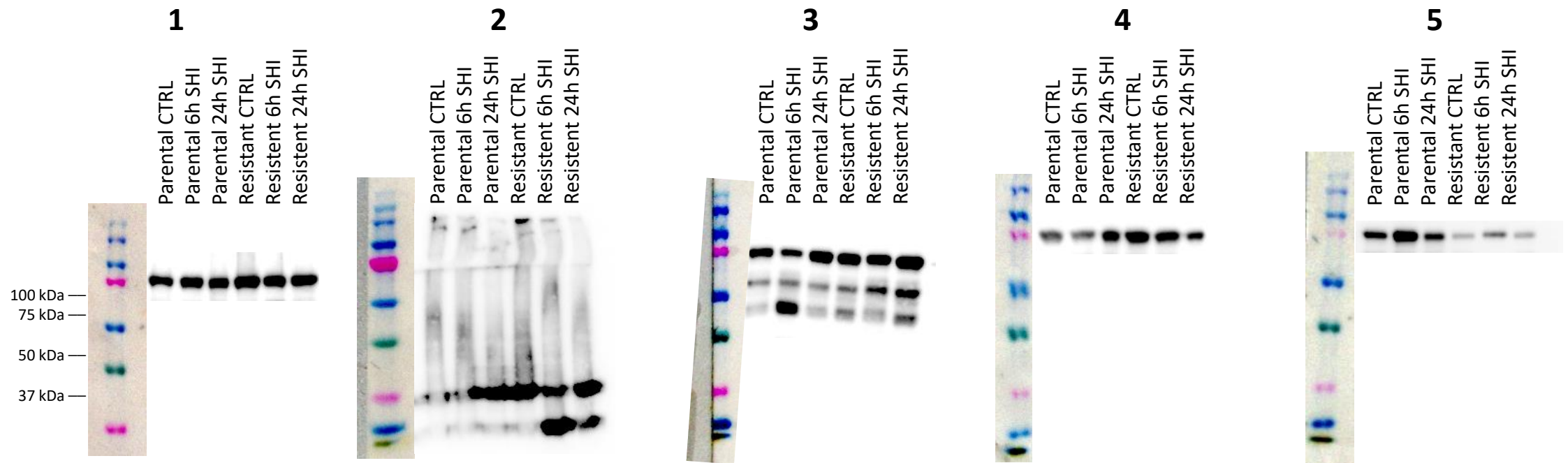
Figure S2f: Detailed information about Figure 5 - Protein expression and activity of cell cycle regulating proteins in parental and sunitinib-resistant 786-O cells after 48 h exposure to 1 or 1.5 μ M SHI. Protein expression of pCDK2 (#), corresponding protein expression of β -actin (*). n.r. = not relevant.

Nekroptose proteins

RIP1, pRIP1, RIP3, pRIP3, MLKL, pMLKL

CAKI-1
(#)

RIP1 (78 kDa)



Total protein
(*)

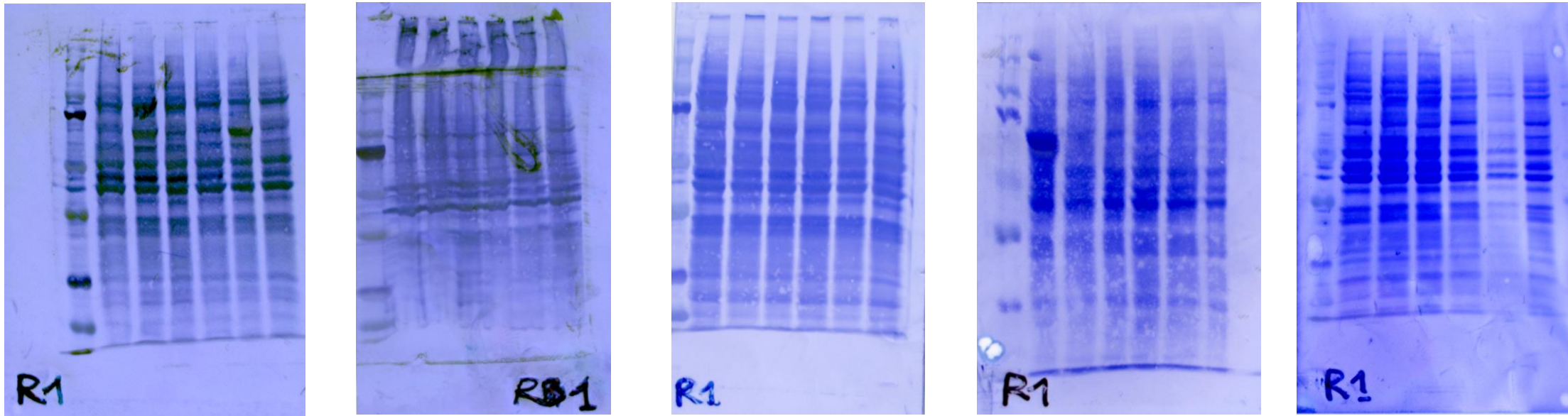
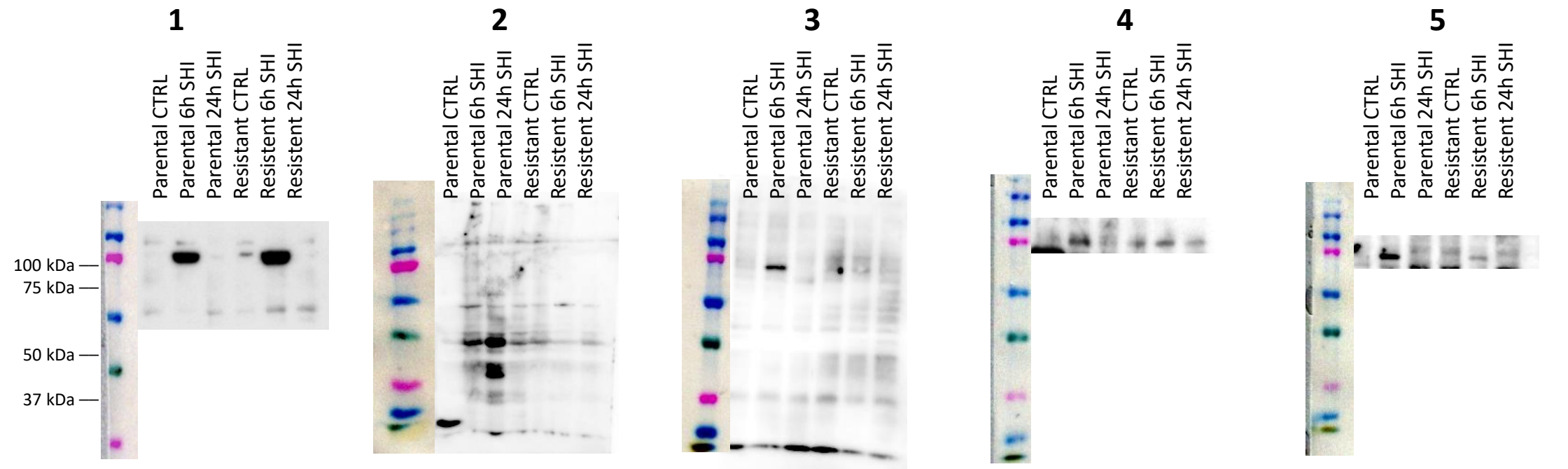


Figure S3a: Detailed information about Figure 10 - Necrosome complex key proteins in parental and sunitinib-resistant Caki-1 cells after 6 and 24 h exposure to 1 μ M SHI. Protein expression of RIP1 (#), corresponding Coomassie blue staining of total protein (*).

CAKI-1

(#)

pRIP1 (78-82 kDa)



(*)

Total protein

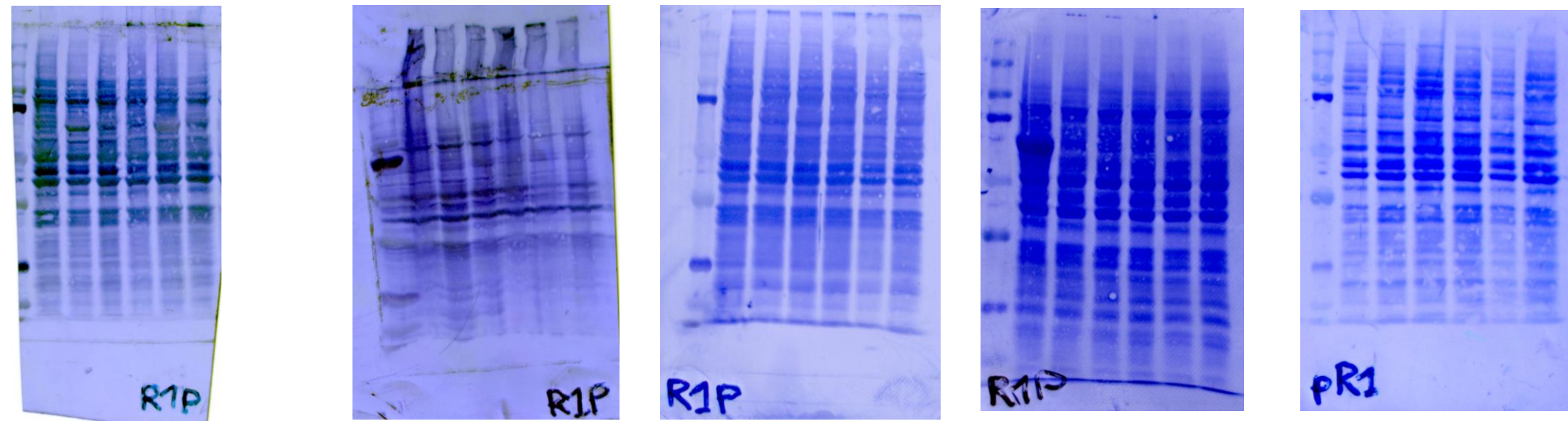


Figure S3b: Detailed information about Figure 10 - Necrosome complex key proteins in parental and sunitinib-resistant Caki-1 cells after 6 and 24 h exposure to 1 μ M SHI. Protein expression of pRIP1 (#), corresponding Coomassie blue staining of total protein (*).

CAKI-1

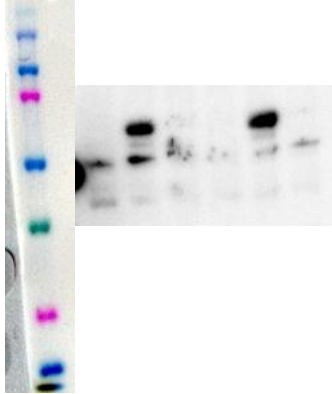
(#)

RIP3 (46-62 kDa)

100 kDa —
75 kDa —
50 kDa —
37 kDa —

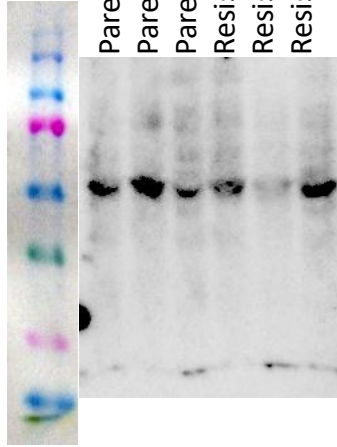
1

Parental CTRL
Parental 6h SHI
Parental 24h SHI
Resistant CTRL
Resistant 6h SHI
Resistant 24h SHI



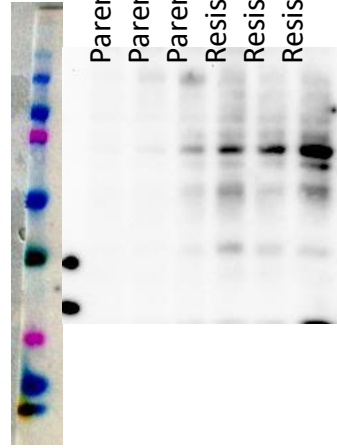
2

Parental CTRL
Parental 6h SHI
Parental 24h SHI
Resistant CTRL
Resistant 6h SHI
Resistant 24h SHI



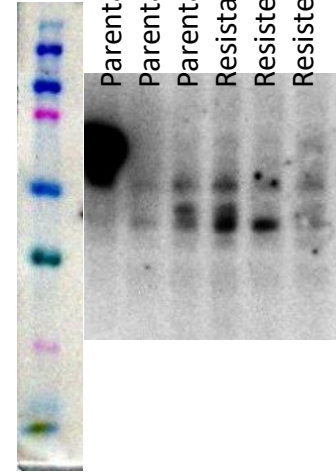
3

Parental CTRL
Parental 6h SHI
Parental 24h SHI
Resistant CTRL
Resistant 6h SHI
Resistant 24h SHI



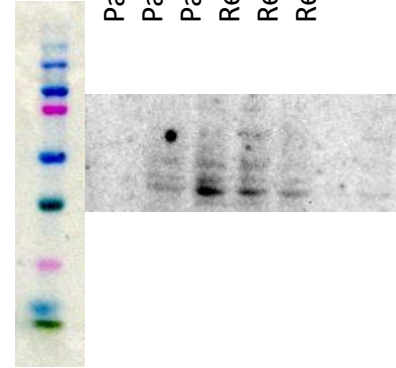
4

Parental CTRL
Parental 6h SHI
Parental 24h SHI
Resistant CTRL
Resistant 6h SHI
Resistant 24h SHI



5

Parental CTRL
Parental 6h SHI
Parental 24h SHI
Resistant CTRL
Resistant 6h SHI
Resistant 24h SHI



(*)

Total protein

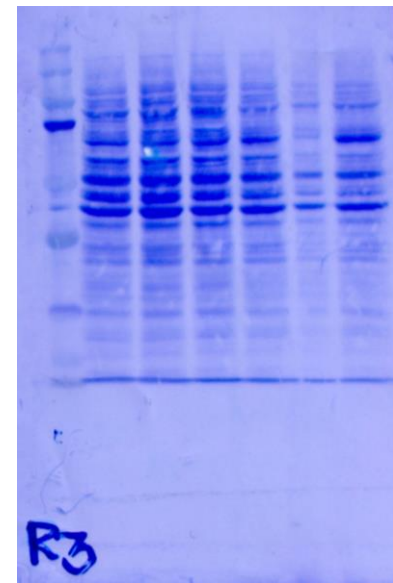
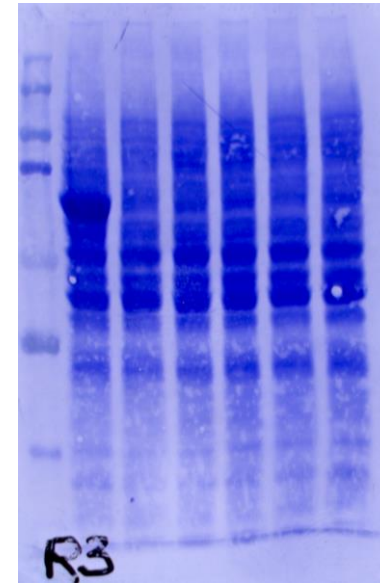
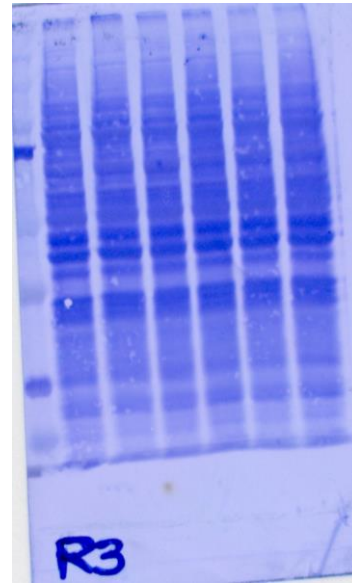
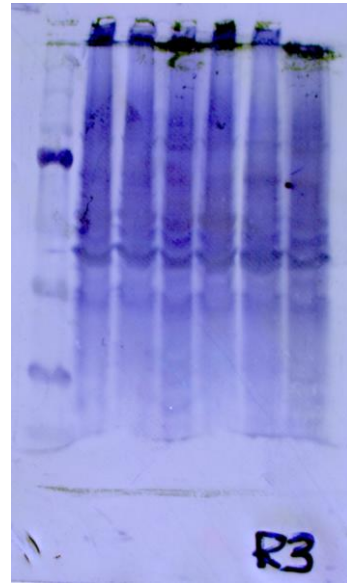
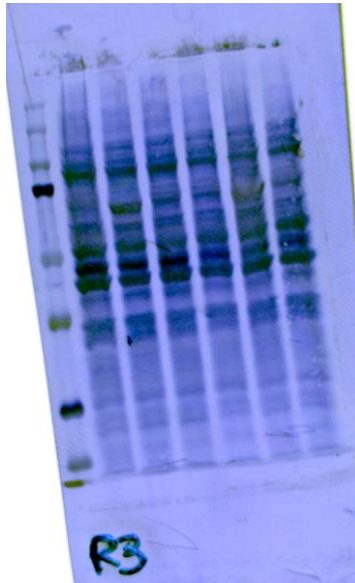
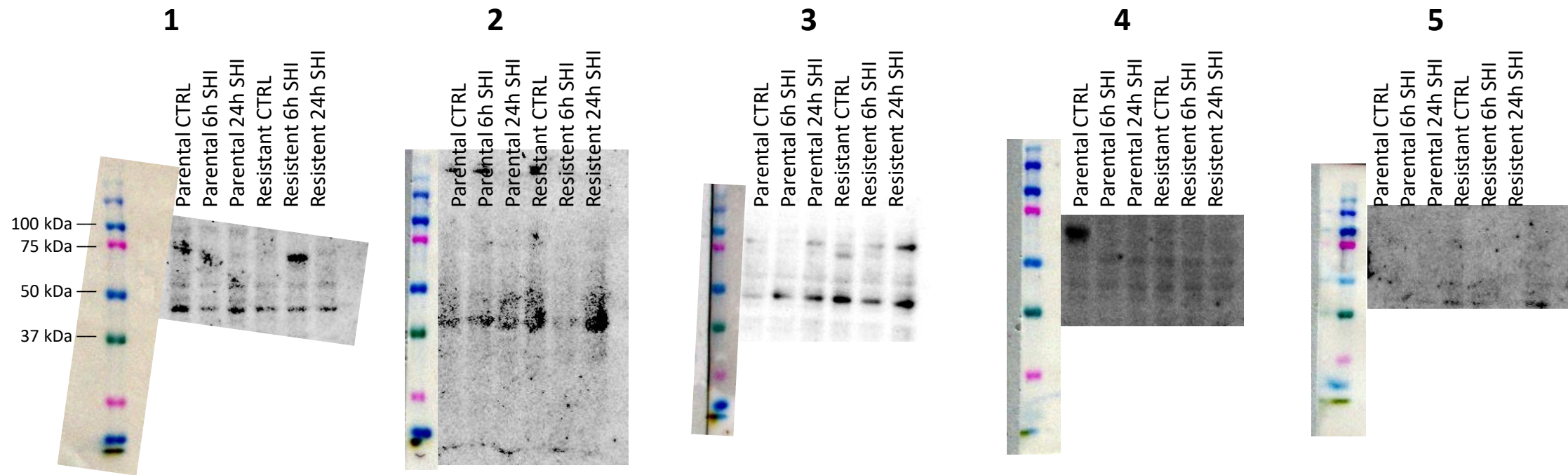


Figure S3c: Detailed information about Figure 10 - Necrosome complex key proteins in parental and sunitinib-resistant Caki-1 cells after 6 and 24 h exposure to 1 μ M SHI. Protein expression of RIP3 (#), corresponding Coomassie blue staining of total protein (*).

CAKI-1

(#)

pRIP3 (46-62 kDa)



(*)

Total protein

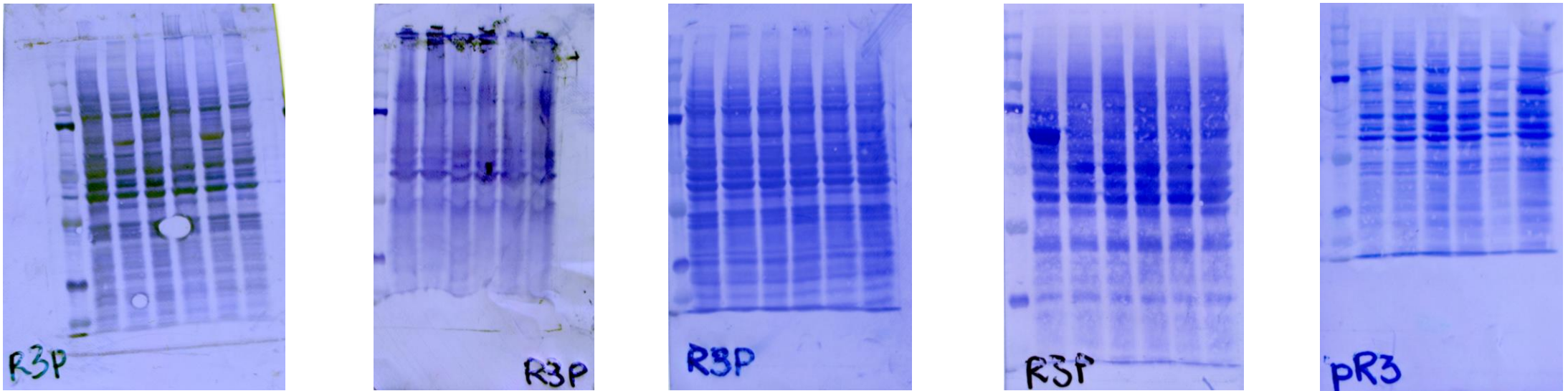
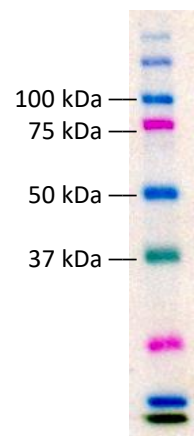


Figure S3d: Detailed information about Figure 10 - Necrosome complex key proteins in parental and sunitinib-resistant Caki-1 cells after 6 and 24 h exposure to 1 μ M SHI. Protein expression of pRIP3 (#), corresponding Coomassie blue staining of total protein (*).

CAKI-1

(#)

MLKL (54 kDa)



1

Parental CTRL
Parental 6h SHI
Parental 24h SHI
Resistant CTRL
Resistant 6h SHI
Resistant 24h SHI

2

Parental CTRL
Parental 6h SHI
Parental 24h SHI
Resistant CTRL
Resistant 6h SHI
Resistant 24h SHI

3

Parental CTRL
Parental 6h SHI
Parental 24h SHI
Resistant CTRL
Resistant 6h SHI
Resistant 24h SHI

4

Parental CTRL
Parental 6h SHI
Parental 24h SHI
Resistant CTRL
Resistant 6h SHI
Resistant 24h SHI

5

Parental CTRL
Parental 6h SHI
Parental 24h SHI
Resistant CTRL
Resistant 6h SHI
Resistant 24h SHI

(*)

Total protein

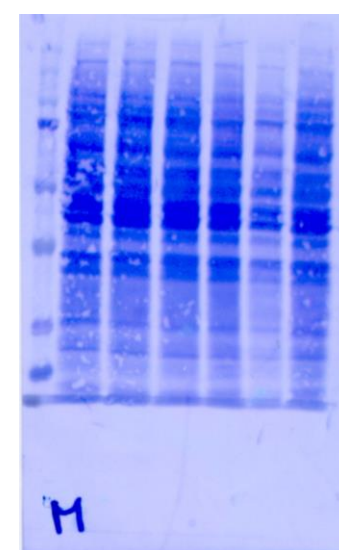
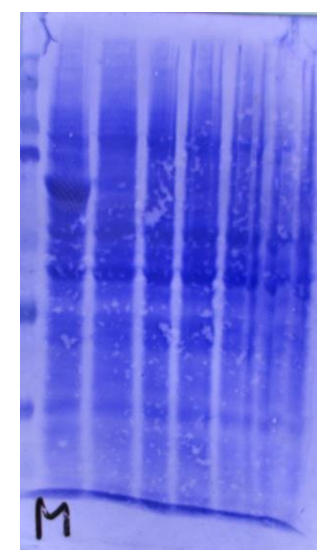
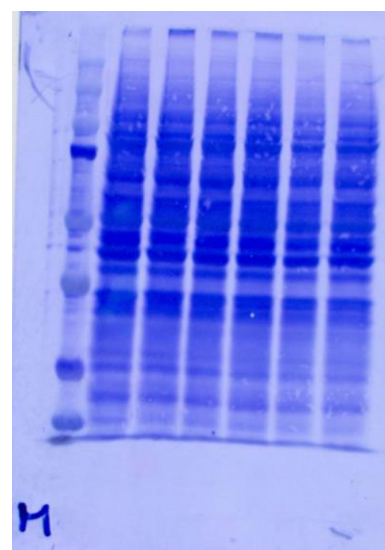
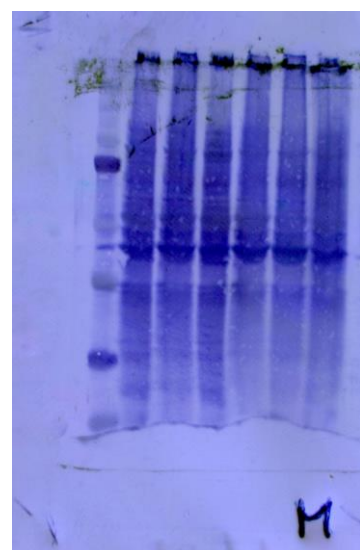
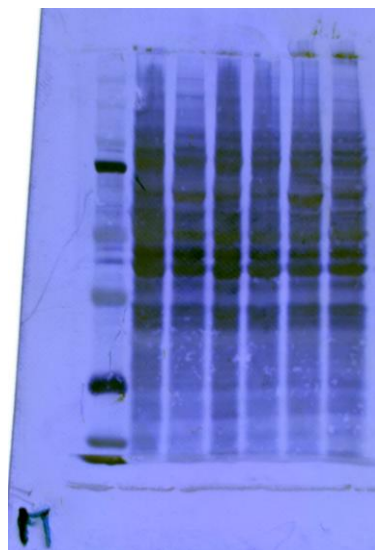
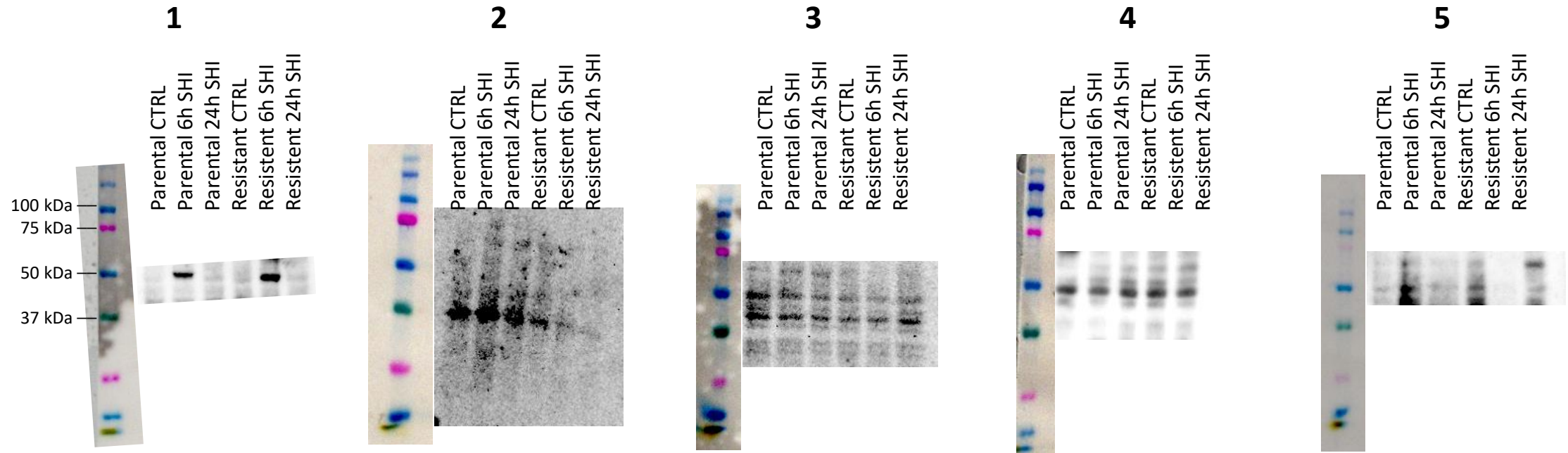


Figure S3e: Detailed information about Figure 10 - Necrosome complex key proteins in parental and sunitinib-resistant Caki-1 cells after 6 and 24 h exposure to 1 μ M SHI. Protein expression of MLKL (#), corresponding Coomassie blue staining of total protein (*).

CAKI-1

(#)

pMLKL (54 kDa)



(*)

Total protein

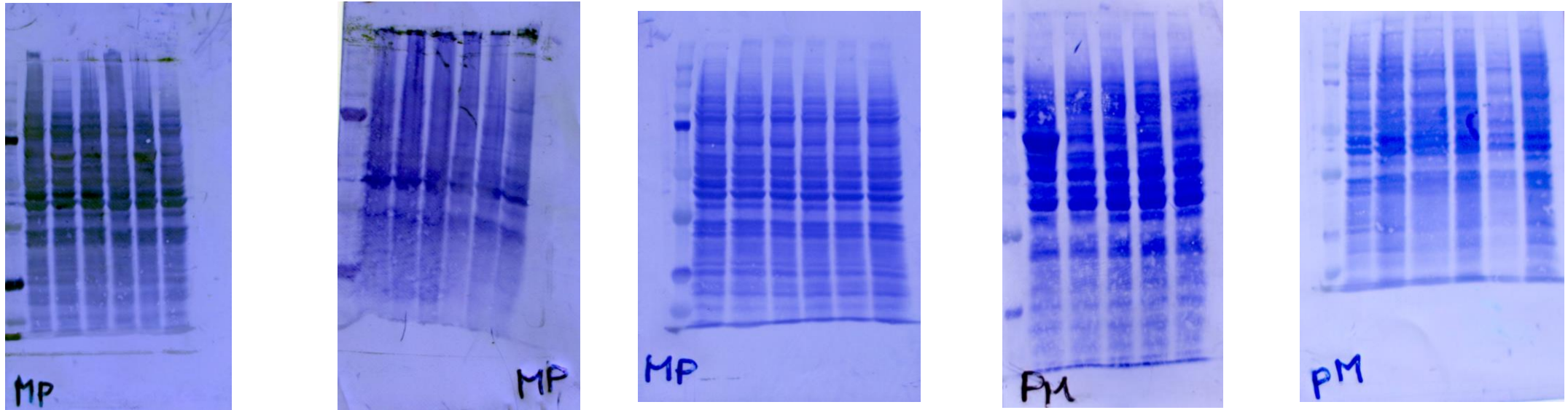


Figure S3f: Detailed information about Figure 10 - Necrosome complex key proteins in parental and sunitinib-resistant Caki-1 cells after 6 and 24 h exposure to 1 μ M SHI. Protein expression of pMLKL (#), corresponding Coomassie blue staining of total protein (*).

Signal proteins

AKT, pAKT, mTOR, pmTOR

CAKI-1

(#)

AKT (60 kDa)

(*)

Total protein

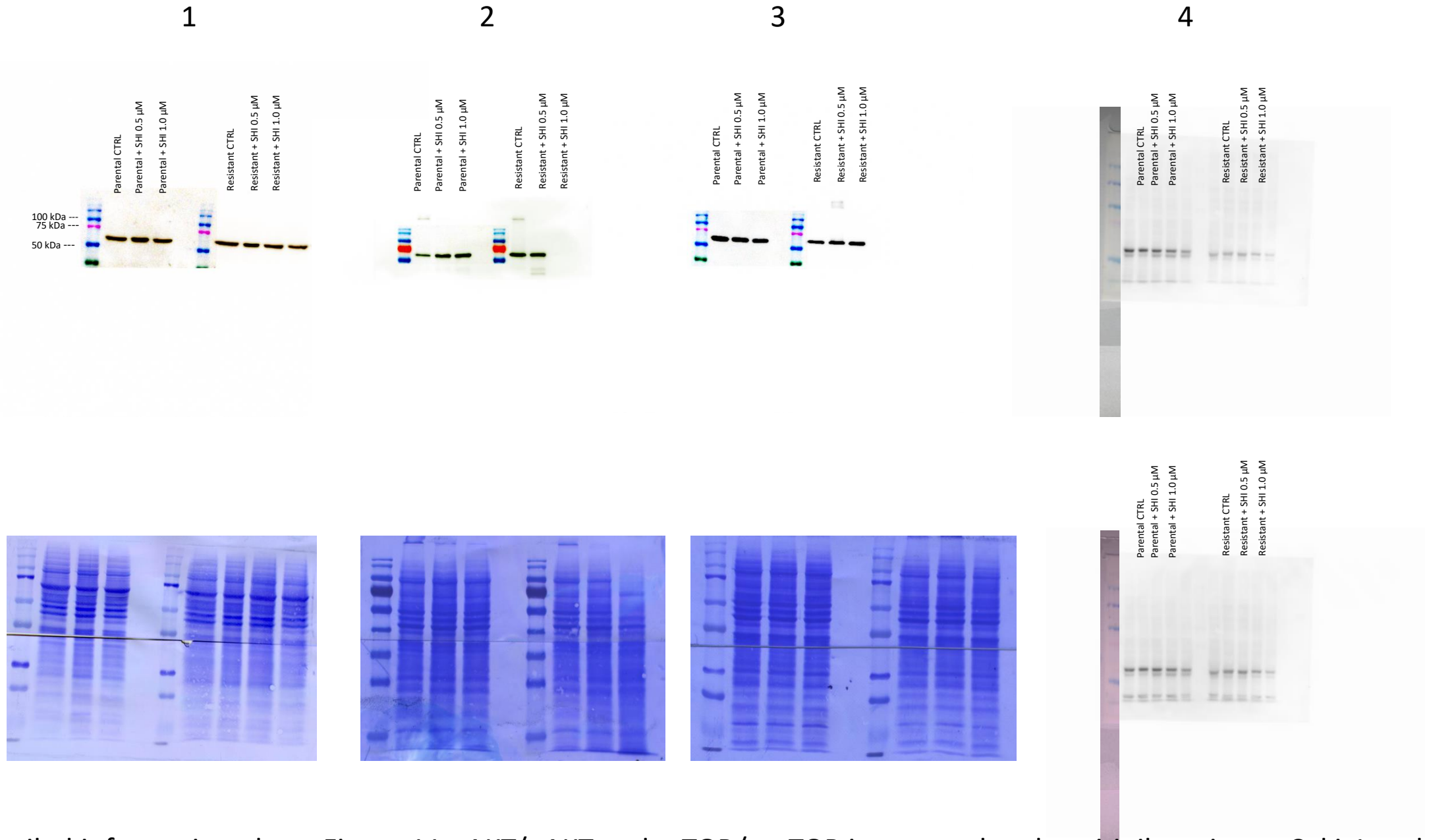


Figure S4a: Detailed information about Figure 11 - AKT/pAKT and mTOR/pmTOR in parental and sunitinib-resistant Caki-1 and 786-O cells after 48 h exposure to SHI [0.5, 1.0 μ M]. Protein expression of AKT (#), corresponding protein expression of β -actin or Coomassie blue staining of total protein (*).

CAKI-1

pAKT (60 kDa)

(#)

(*)

Total protein

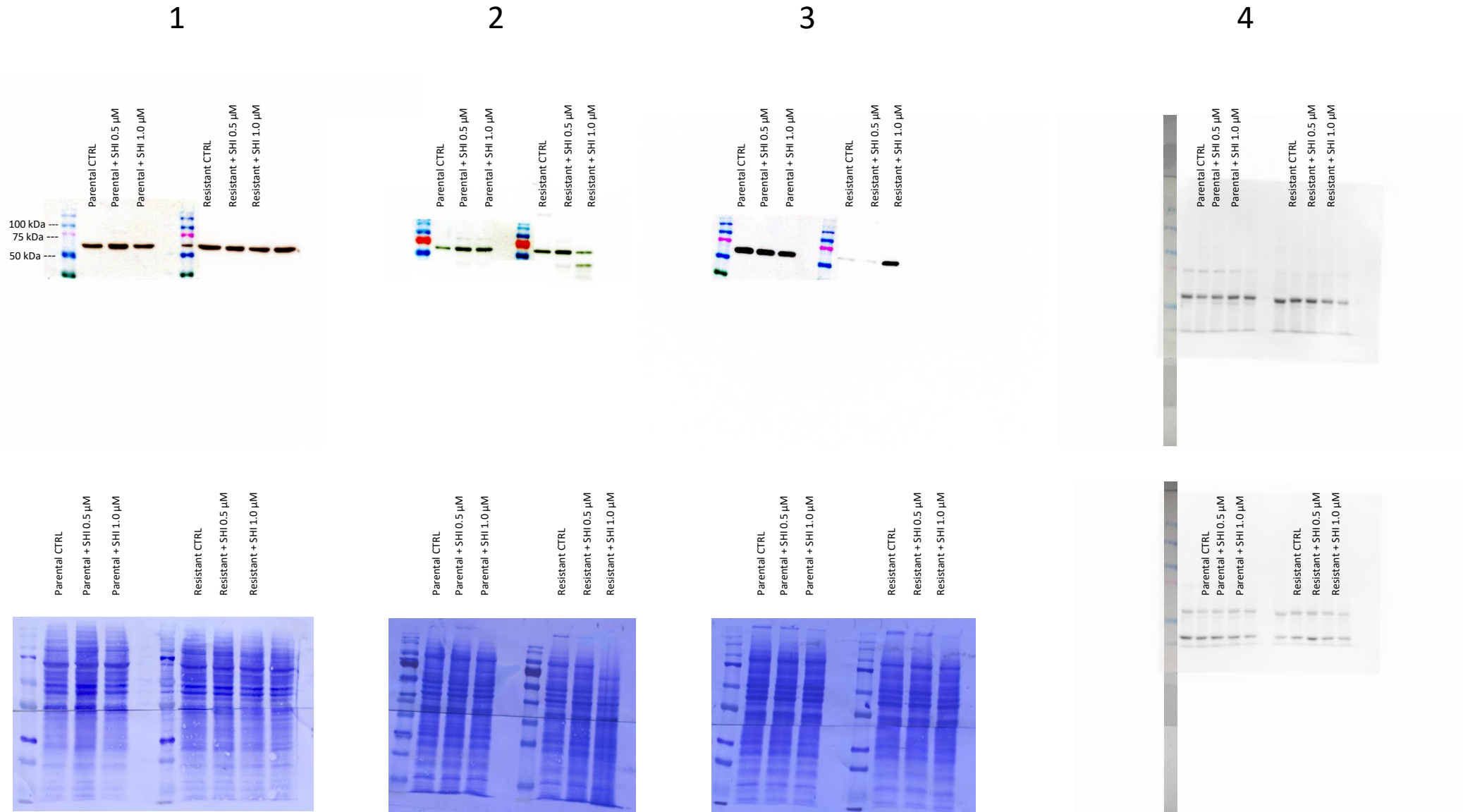


Figure S4b: Detailed information about Figure 11 - AKT/pAKT and mTOR/pmTOR in parental and sunitinib-resistant Caki-1 and 786-O cells after 48 h exposure to SHI [0.5, 1.0 μ M]. Protein expression of pAKT (#), corresponding protein expression of β -actin or Coomassie blue staining of total protein (*).

CAKI-1

(#)

mTOR (289 kDa)

(*)

Total protein

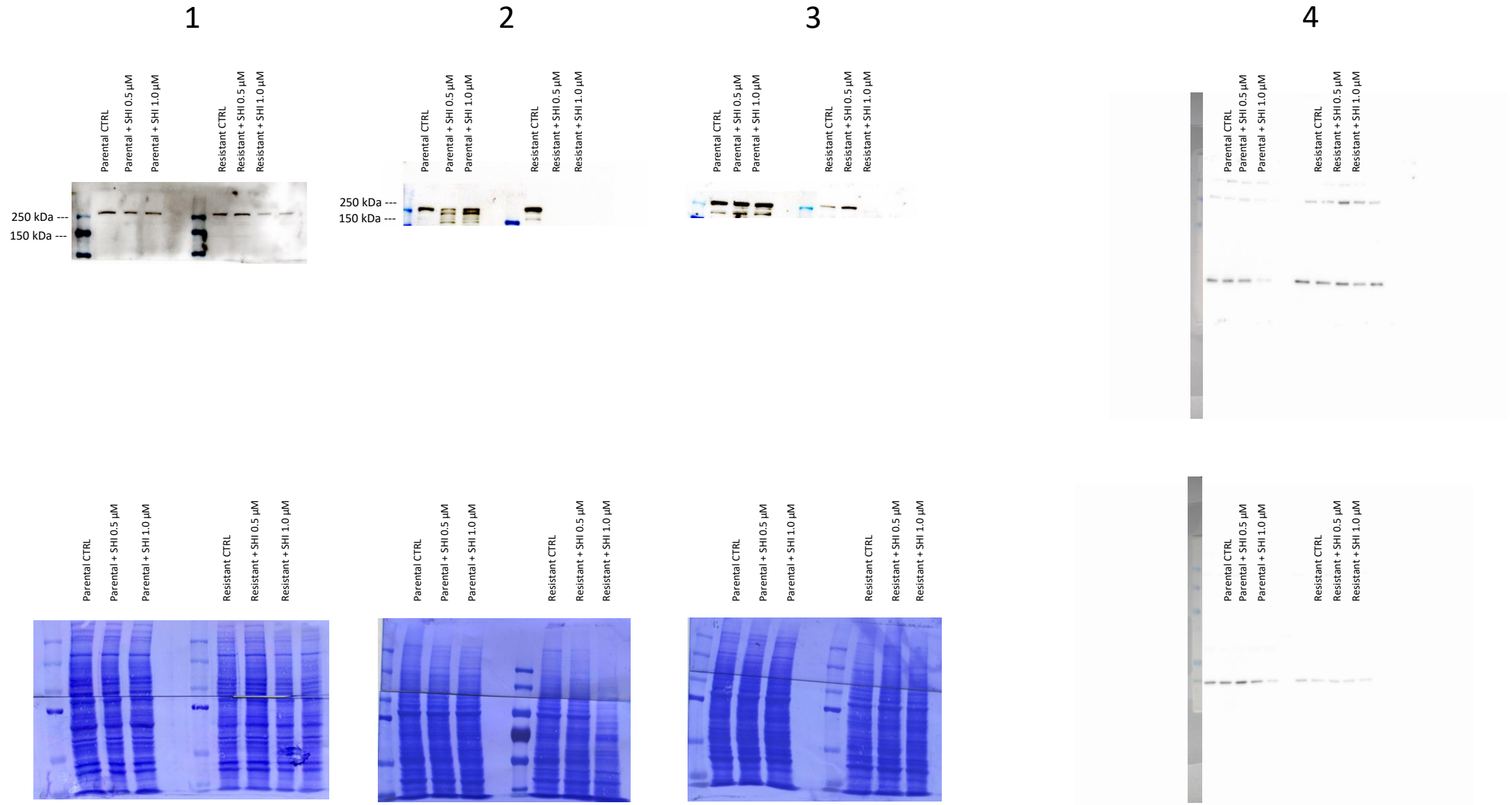


Figure S4c: Detailed information about Figure 11 - AKT/pAKT and mTOR/pmTOR in parental and sunitinib-resistant Caki-1 and 786-O cells after 48 h exposure to SHI [0.5, 1.0 μ M]. Protein expression of mTOR (#), corresponding protein expression of β -actin or Coomassie blue staining of total protein (*).

CAKI-1

(#)

pmTOR (289 kDa)

(*)

Total protein

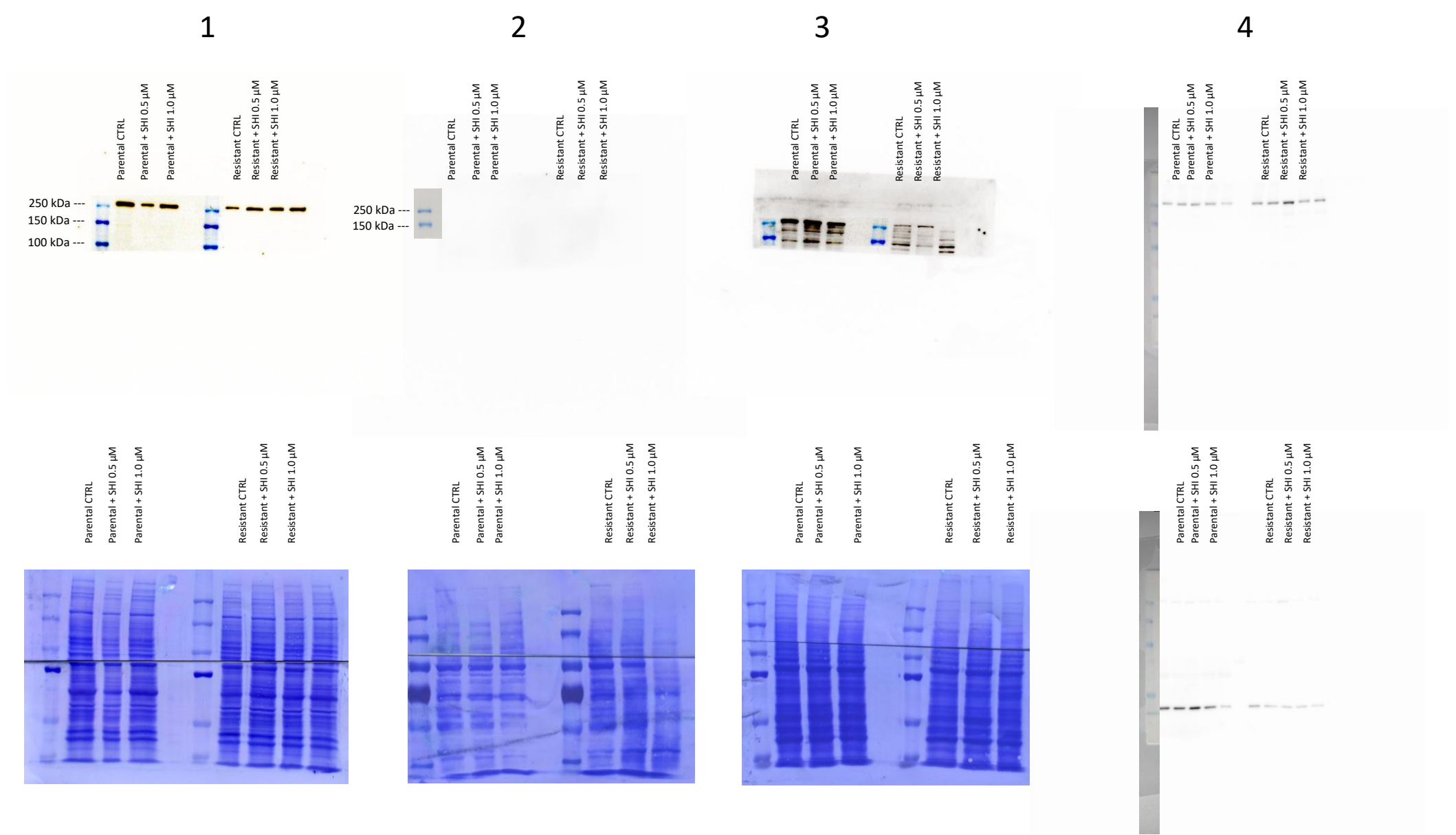


Figure S4d: Detailed information about Figure 11 - AKT/pAKT and mTOR/pmTOR in parental and sunitinib-resistant Caki-1 and 786-O cells after 48 h exposure to SHI [0.5, 1.0 μ M]. Protein expression of pmTOR (#), corresponding protein expression of β -actin or Coomassie blue staining of total protein (*).

786-O

(#)

AKT (60 kDa)

(*)

Total protein

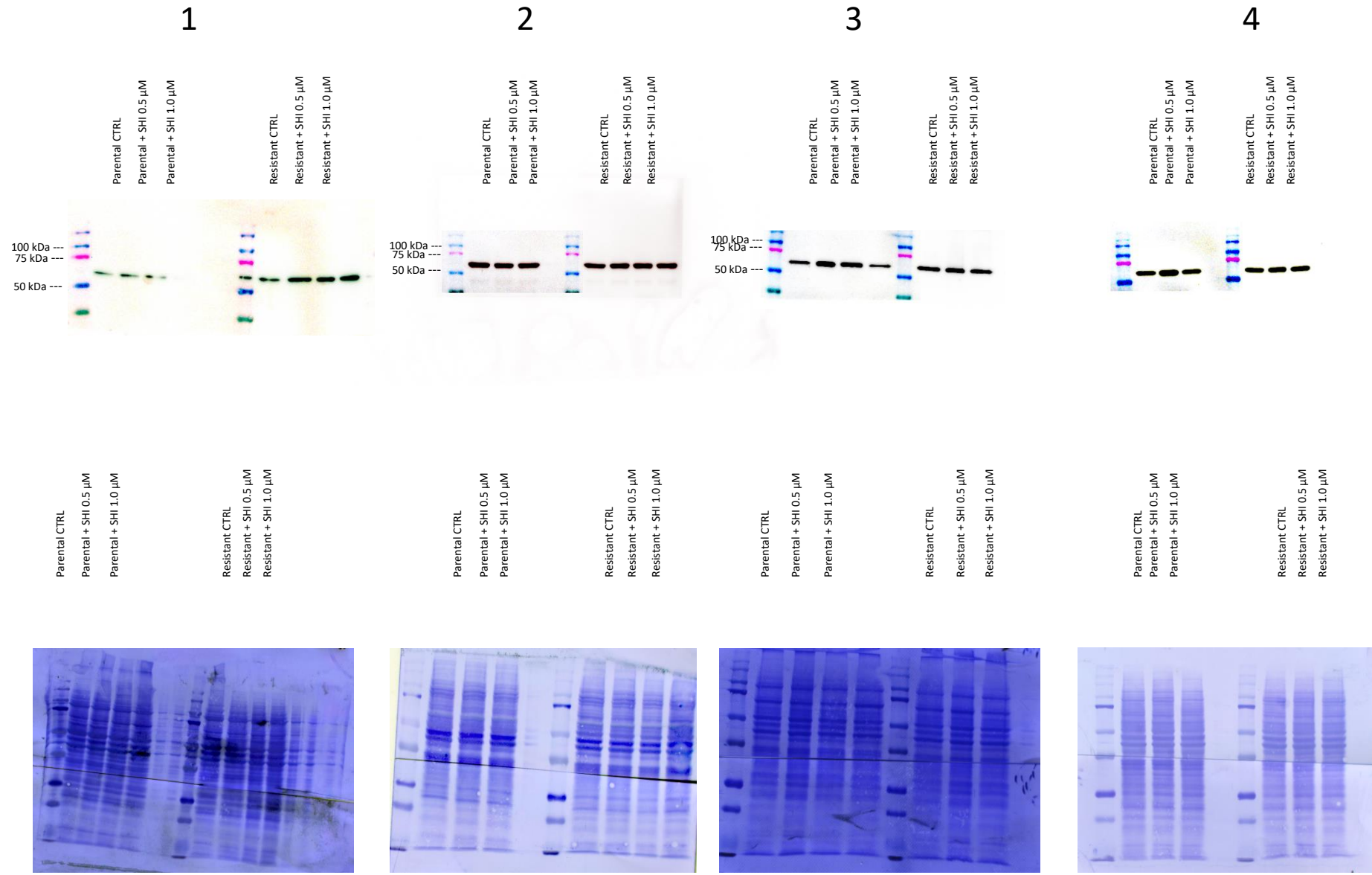


Figure S4e: Detailed information about Figure 11 - AKT/pAKT and mTOR/pmTOR in parental and sunitinib-resistant Caki-1 and 786-O cells after 48 h exposure to SHI [0.5, 1.0 μ M]. Protein expression of AKT (#), corresponding Coomassie blue staining of total protein (*).

786-O

(#)

pAKT (60 kDa)

(*)

Total protein

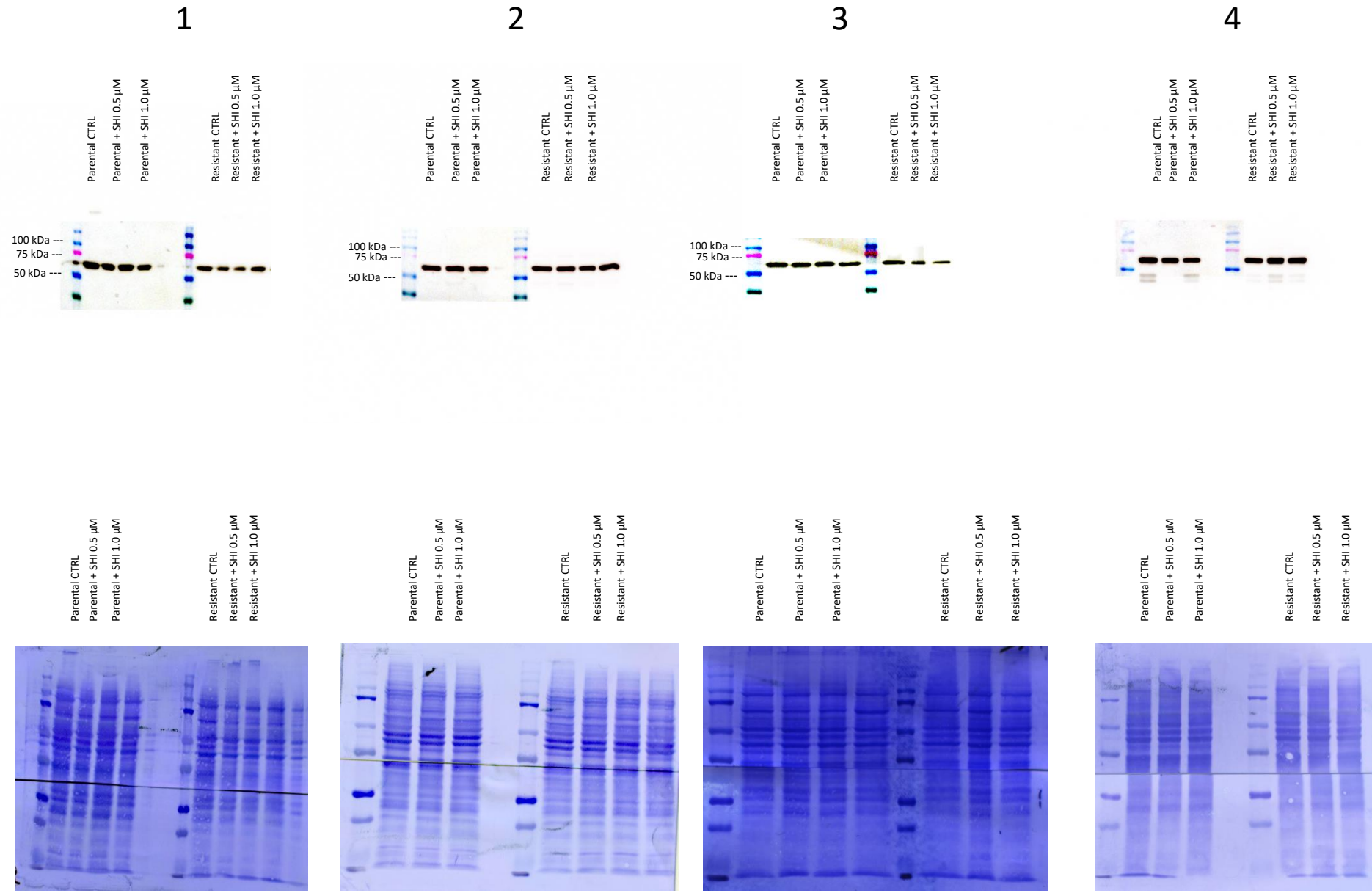


Figure S4f: Detailed information about Figure 11 - AKT/pAKT and mTOR/pmTOR in parental and sunitinib-resistant Caki-1 and 786-O cells after 48 h exposure to SHI [0.5, 1.0 μ M]. Protein expression of pAKT (#), corresponding Coomassie blue staining of total protein (*).

786-O

(#)

mTOR (289 kDa)

(*)

Total protein

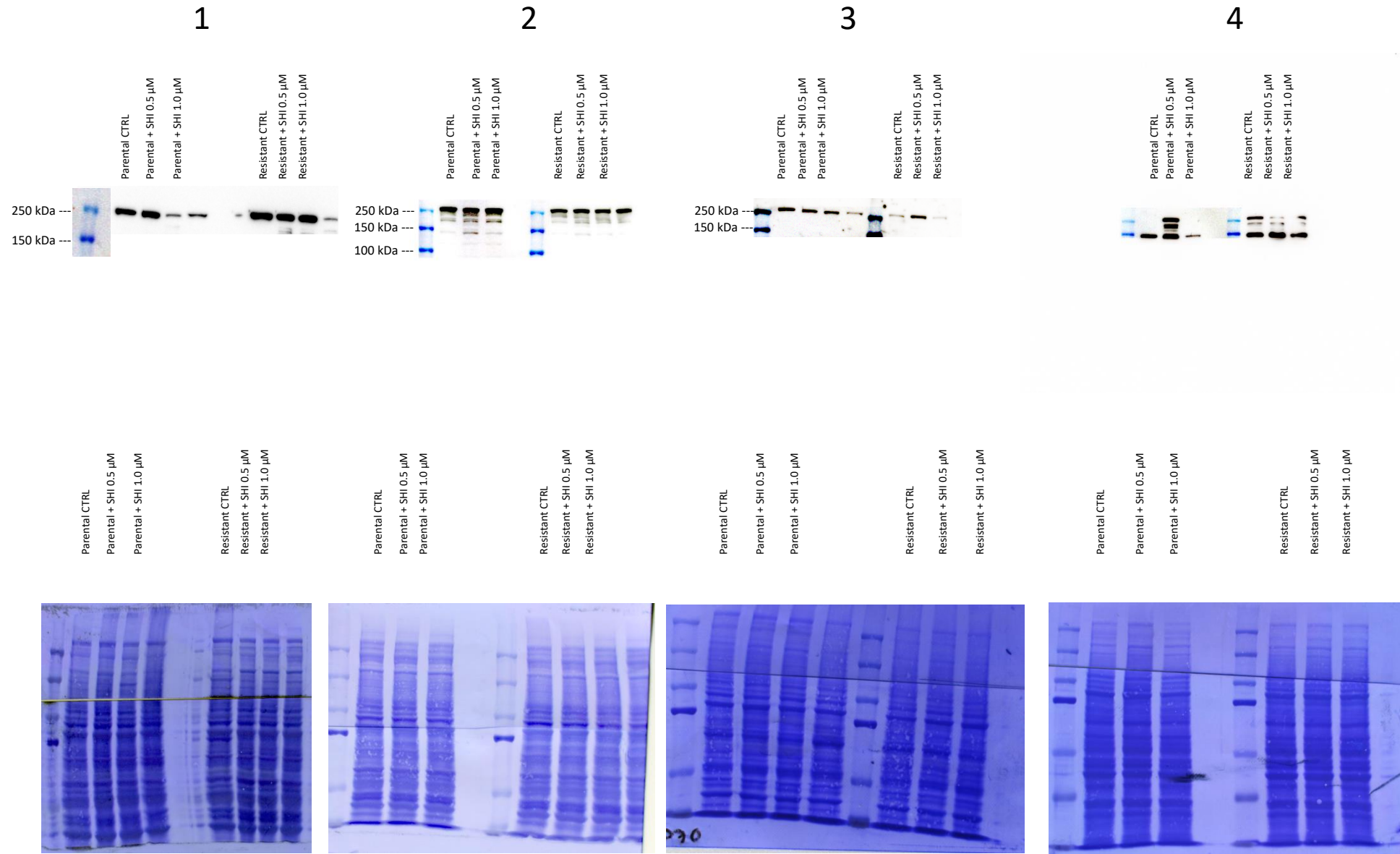


Figure S4g: Detailed information about Figure 11 - AKT/pAKT and mTOR/pmTOR in parental and sunitinib-resistant Caki-1 and 786-O cells after 48 h exposure to SHI [0.5, 1.0 μ M]. Protein expression of mTOR (#), corresponding Coomassie blue staining of total protein (*).

786-O

(#)

pmTOR (289 kDa)

(*)

Total protein

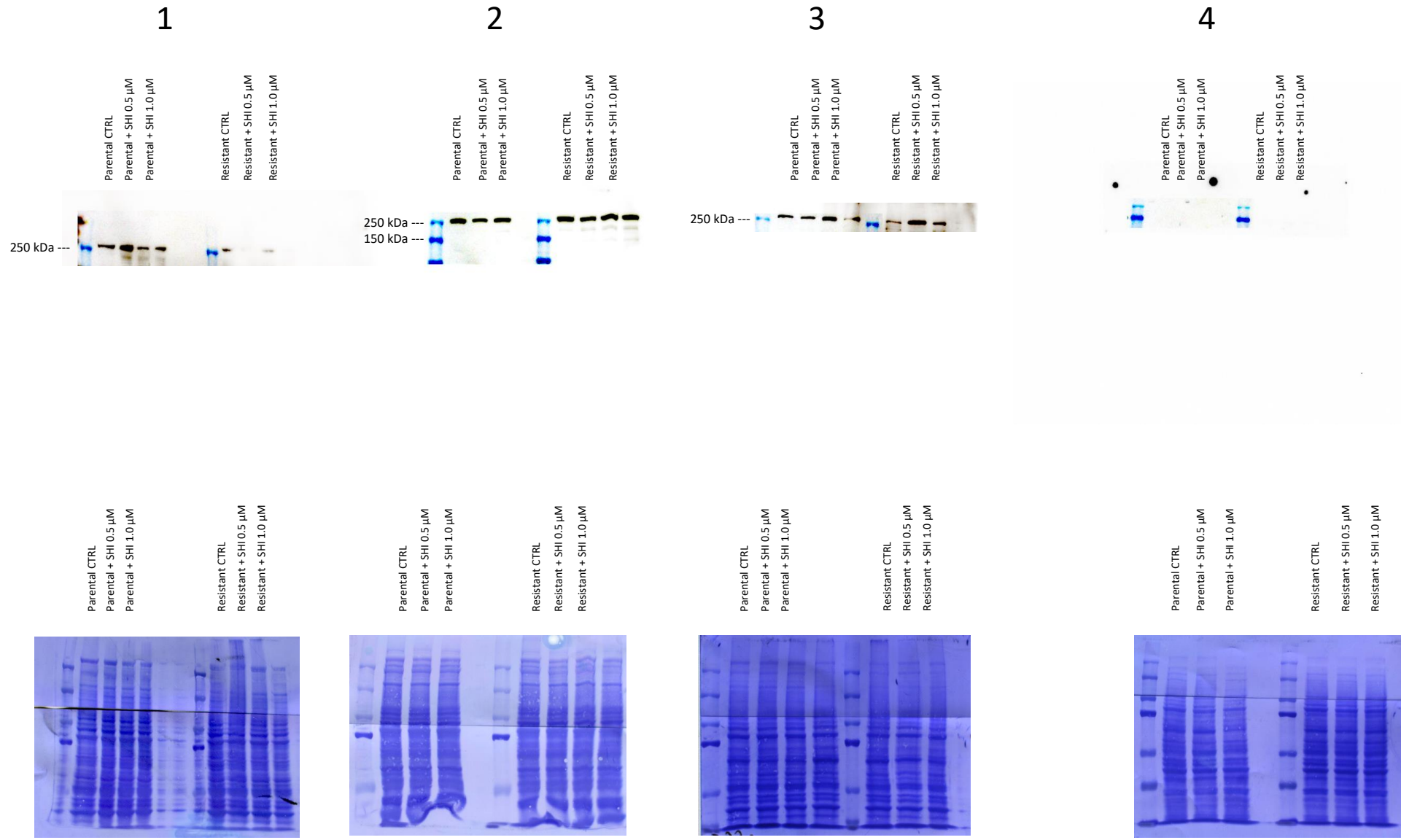


Figure S4h: Detailed information about Figure 11 - AKT/pAKT and mTOR/pmTOR in parental and sunitinib-resistant Caki-1 and 786-O cells after 48 h exposure to SHI [0.5, 1.0 μ M]. Protein expression of pmTOR (#), corresponding Coomassie blue staining of total protein (*).