

Comparison of Hypoxia Inducible Factor 1 α (HIF-1 α) levels in HCT-15 cell line and Human Colon Cancer Stem Cell-like Cells

Vergleich auf Hypoxia Inducible Faktor 1 α (HIF-1 α) spiegel zwischen HCT-15 Zell Linie und Human Dunderm krebs stem cell-like Zellen

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Aim

Hypoxia Inducible Factors (HIFs) are transcription factors that respond to decrease of oxygen level in the cellular environment. Three types of HIFs are known, HIF-1, HIF-2 and HIF-3. All HIFs are heterodimers consisted of α and β subunits. HIF signaling plays a central role in angiogenesis and in the regulation of human metabolism and HIF-1 specifically is responsible for cellular and systemic responses to hypoxia. HIF-1 consists of an α and a β subunit. HIF-1 α is oxygen dependent and HIF-1 β a constitutively-expressed aryl hydrocarbon receptor nuclear translocator (ARNT).

The aim of this study was to quantify and correlate the HIF-1 α levels between Human Colon Cancer Cells and Human Colon Cancer Stem Cell-like Cells (Colon CSCs) growing in normal oxygen concentration and hypoxia.

Materials and methods

Growth curves were generated for three cultures, HCT-15, Colon CSCs growing at physiological O₂ concentration and Colon CSCs growing at absence of O₂. HIF-1 α levels were quantified with both flow cytometry and Real Time PCR.

Results

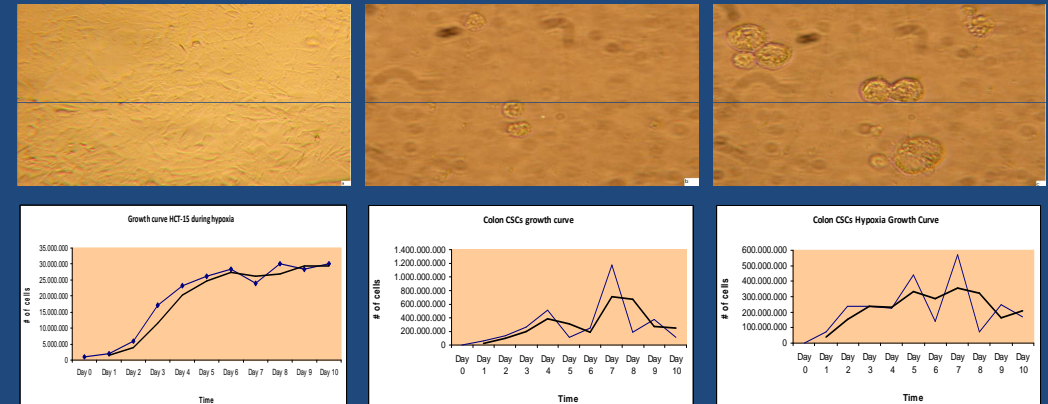
Data were drawn and correlated between the three cultures. HIF-1 α levels were higher in Colon CSCs than in HCT-15, both growing at physiological O₂ concentration. The HIF-1 α levels of Colon CSCs growing at absence of O₂ are indicative of resistance to cell death cause of hypoxia after an adjustment period.

Conclusion

HIF-1 is expressed in most of the oxygen breathing animals and is responsible for cellular and systemic responses to hypoxia. It consists of two subunits (α and β). The α subunit is oxygen dependent and was quantified in HCT-15 cells and in Colon CSCs to investigate the percentage of metabolic changes that hypoxia promotes in Colon CSCs. The results indicate that Colon CSCs, which express higher levels of HIF-1 α , have a higher angiogenic and metabolic profile than Colon Cancer Cells, both during normal O₂ concentrations and hypoxia.

References

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| | HIF-1 α rates from day to day (% of total events) | | |
|-------|--|------------|--------------------|
| | HCT-15 Hypoxia | Colon CSCs | Colon CSCs Hypoxia |
| Day 1 | 3,6 | 1,6 | 1,9 |
| Day 3 | 26,9 | 2,5 | 1,7 |
| Day 5 | 47,2 | 3 | 1,7 |
| Day 7 | 49,6 | 4,6 | 5,4 |

Figures a, b, c represent cell lines HCT-15 (hypoxia), Colon CSCs (Normal O₂), Colon CSCs (Hypoxia) respectively. Underneath every figure there is the growth curve for each cell line.

The table above shows the percentage of HIF-1 α expression measured with flow cytometry for the three cell lines. Colon CSCs during hypoxia express high HIF-1 α levels (according to total number of cells for every cell line). As a result Colon CSCs can adjust to different O₂ concentrations.

Disclosure of Potential Conflicts of Interest

None of the authors of the above study has declared any conflict of interest

