

## ANESTETICI GENERALI

Gli anestetici generali sono farmaci in grado di indurre una interruzione reversibile di molte funzioni vitali quali: la conoscenza, la sensibilità dolorifica, i riflessi al dolore e alla difesa e la tensione muscolare, inducendo quindi nell'individuo uno stato di analgesia, amnesia, inibizione di riflessi sensoriali e vegetativi ma anche provocando rilassamento della muscolatura scheletrica, tale da consentire interventi chirurgici.

Gli effetti di questi farmaci non dovrebbero interessare altre funzioni vitali quali: il centro del respiro e l'apparato cardio-circolatorio. L'anestetico ideale dovrebbe indurre gradualmente ma rapidamente anestesia consentendo una rapida ripresa dall'effetto anestetico una volta terminata la sua somministrazione, e dovrebbe possedere un ampio margine di sicurezza associato a scarsi effetti collaterali. Poiché nessun singolo anestetico da solo presenta questi requisiti si preferisce l'uso combinato di + anestetici generali per minimizzare gli effetti indesiderati e potenziare le azioni favorevoli. Diversi sono gli stadi dell'anestesia:

*analgesia*, piccole concentrazioni di anestetico a livello del SNC interrompono la trasmissione nocicettiva;

*eccitazione*, iperattività motoria, respiro irregolare dovuto ad inibizione dei centri inibitori o stimolazione della conduzione eccitatoria;

*anestesia chirurgica*, soppressa attività riflessa, rilassamento muscolare;

*paralisi*, *depressione bulbare*, *morte*, tale stadio non deve essere mai raggiunto, per questo è imp. il continuo monitoraggio delle funzioni respiratorie e cardiovascolari per valutare la profondità dell'anestesia.

Il meccanismo d'azione di tali farmaci è ancora oggetto di studio, si è ipotizzato che essi agiscono o inibendo la trasmissione eccitatoria, antagonizzando a livello recettoriale le azioni dei neurotrasmettitori eccitatori (glutammato e glicina), oppure potenziando la trasmissione inibitoria, per interazioni a livello del recettore GABA (essi causano iperpolarizzazione della membrana per apertura dei canali del  $Cl^-$ , prolungando così il periodo refrattario dei neuroni).

Gli anestetici generali possono essere divisi in due classi:

**ANESTETICI INALATORI:** richiedono particolari apparecchiature per la loro somministrazione. Non sono utilizzati da soli, vengono infatti preferite le associazioni di tali farmaci con gli anestetici endovenosi. La velocità di induzione all'anestesia, cioè la velocità con cui si raggiunge una concentrazione efficace nel cervello, dipende dalla loro solubilità. E' stato dimostrato infatti, che più un farmaco è liposolubile, più esso raggiunge con rapidità il SNC ed esplica la sua azione; quindi la liposolubilità è indice di rapidità e potenza. Tuttavia i farmaci liposolubili presentano una breve durata d'azione in quanto sono eliminati più velocemente dall'organismo. I farmaci idrosolubili invece richiedono tempi più lunghi per dare inizio alla loro azione ma hanno una più lunga durata. Alcuni esempi di anestetici gen. inalatori sono:

- **$N_2O$  protossido di azoto**, composto inorganico, gassoso; da solo non è in grado di produrre anestesia chirurgica, viene usato da solo quando non si vuole raggiungere il terzo stadio dell'anestesia, ad esempio in odontoiatria, spesso usato in associazione con altri anestetici per ridurre le concentrazioni di questi;

- **$CH_3CH_2OCH_2CH_3$  etere etilico**, non più utilizzato perché esplosivo ed infiammabile ed inoltre tossico per le vie respiratorie;

- **$CH_3Cl$  cloroformio**, non più usato per il suo stretto margine di sicurezza, induceva tossicità epatica, renale e cardiaca;

-**Alotano**, idrocarburo alogenato, è un etano sostituito. E' un liquido trasparente, dall'odore dolciastro, non è infiammabile. E' oggi il più usato. E' stato ideato in base all'ipotesi che la struttura alogenata potesse conferire alla molecola maggiore stabilità chimica, intermedia solubilità ematica e soprattutto buon potere anestetico. Dal punto di vista chimico è l'unico alogenato ad avere nella sua struttura un atomo di bromo, che probabilmente contribuisce con i tre atomi di fluoro alla sua potenza. L'anestesia è rapida e regredisce altrettanto rapidamente, il che porta all'utilizzo di questo farmaco da solo o in combinazione con  $N_2O$ . Può indurre aritmie per sensibilizzazione alle

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